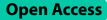
RESEARCH

BMC Infectious Diseases



Machine learning models for prediction of invasion Klebsiella pneumoniae liver abscess syndrome in diabetes mellitus: a singled centered retrospective study



Chengyi Feng¹, Jia Di¹, Shufang Jiang¹, Xuemei Li¹ and Fei Hua^{1*}

Abstract

Objective This study aimed to develop and validate a machine learning algorithm-based model for predicting invasive Klebsiella pneumoniae liver abscess syndrome(IKPLAS) in diabetes mellitus and compare the performance of different models.

Methods The clinical signs and data on the admission of 213 diabetic patients with Klebsiella pneumoniae liver abscesses were collected as variables. The optimal feature variables were screened out, and then Artificial Neural Network, Support Vector Machine, Logistic Regression, Random Forest, K-Nearest Neighbor, Decision Tree, and XGBoost models were established. Finally, the model's prediction performance was evaluated by the ROC curve, sensitivity (recall), specificity, accuracy, precision, F1-score, Average Precision, calibration curve, and DCA curve.

Results Four features of hemoglobin, platelet, D-dimer, and SOFA score were screened by the recursive elimination method, and seven prediction models were established based on these variables. The AUC (0.969), F1-Score(0.737), Sensitivity(0.875) and AP(0.890) of the SVM model were the highest among the seven models. The KNN model showed the highest specificity (1.000). Except that the XGB and DT models over-estimates the occurrence of IKPLAS risk, the other models' calibration curves are a good fit with the actual observed results. Decision Curve Analysis showed that when the risk threshold was between 0.4 and 0.8, the net rate of intervention of the SVM model was significantly higher than that of other models. In the feature importance ranking, the SOFA score impacted the model significantly.

Conclusion An effective prediction model of invasion Klebsiella pneumoniae liver abscess syndrome in diabetes mellitus could be established by a machine learning algorithm, which had potential application value.

Keywords Machine learning, Diabetes mellitus, Pyogenic liver abscess, Klebsiella pneumoniae, Prediction model

Fei Hua huafei1970@suda.edu.cn

*Correspondence:

¹ Department of Infection Control, The Third Affiliated Hospital of Soochow University, Changzhou 213003, China

Introduction

Pyogenic liver abscess is a purulent lesion caused by the invasion of pathogenic bacteria into the liver. The incidence of PLA varies slightly in various regions of the world and is increasing yearly [1]. The incidence rate in European and American countries was about $(1.0 \sim 4.1)/100,000$, the annual incidence rate in some Asian countries was $(12 \sim 18)/100,000$, and the

© The Author(s) 2023. Open Access This article is licensed under a Creative Commons Attribution 4.0 International License, which permits use, sharing, adaptation, distribution and reproduction in any medium or format, as long as you give appropriate credit to the original author(s) and the source, provide a link to the Creative Commons licence, and indicate if changes were made. The images or other third party material in this article are included in the article's Creative Commons licence, unless indicated otherwise in a credit line to the material. If material is not included in the article's Creative Commons licence and your intended use is not permitted by statutory regulation or exceeds the permitted use, you will need to obtain permission directly from the copyright holder. To view a copy of this licence, visit http://creativecommons.org/licenses/by/4.0/. The Creative Commons Public Domain Dedication waiver (http://creativeco mmons.org/publicdomain/zero/1.0/) applies to the data made available in this article, unless otherwise stated in a credit line to the data.

annual incidence rate in mainland China was about $(1.1 \sim 5.4)/100,000$ [1-4]. Incidence is higher in males than females (3.3/100,000 vs. 1.3/100,000) [5]. Although the diagnosis and cure rates of PLA had improved significantly with the development of medical technology, the mortality rate was still around 10% [6]. In China and throughout the Asia-Pacific region, the primary pathogen of PLA is Klebsiella pneumoniae [7], which is prevalent in diabetic patients [4]. Diabetic patients with Klebsiella pneumoniae liver abscess are likelier to develop the invasive syndrome, IKPLAS [8]. IKPLAS refers to Klebsiella pneumoniae liver abscess with metastatic infection such as lung abscess, endophthalmitis, meningitis, necrotizing fasciitis, etc.IKPLAS has the characteristics of acute onset, rapid disease progression, and lack of specific clinical manifestations. If patients are not diagnosed and treated in time, the prognosis is generally poor [9]. Although there have been some studies on IKPLAS in the past, most of them are case reports [10-13], and there is no relevant literature report on its clinical prediction model.

Compared with traditional statistical methods, machine learning, as a branch of artificial intelligence, can analyze and obtain rules from existing data and continuously improve and build models based on algorithms and data [14]. Furthermore, it shows apparent advantages in clinical diagnosis and prognosis prediction [15, 16]. This study compared the performance of seven different machine learning methods in predicting the progression of the invasive Klebsiella pneumoniae liver abscess syndrome. Then, a model that can effectively identify high-risk patients is selected, which can help clinical decision-making and provide new perspectives for research in this field.

Materials and methods

Patients pre-processing

This study included patients with diabetes and Klebsiella pneumoniae liver abscesses admitted to Changzhou First People's Hospital from January 1, 20,15 to December 31, 2021. The inclusion criteria were (1) Imaging showed liver abscess, and the puncture fluid or microbial blood culture was Klebsiella pneumoniae. (2) Diabetes diagnosis was based on the "Chinese Guidelines for the Prevention and Treatment of Type 2 Diabetes, 2020 Edition". The exclusion criteria were (1) Patients who died on admission. (2) Patients are automatically discharged or referred midway through. (3) Liver abscess secondary to primary or metastatic liver tumors. (4) Patients with abnormal coagulation function, platelet count, or dysfunction in the past. (5) The age is less than 18 years old. The primary observation was a diagnosis of IKPLAS during hospitalization. The diagnostic criteria of IKPLAS were liver abscess caused by Klebsiella pneumoniae and metastatic infection such as lung abscess, endophthalmitis, meningitis, necrotizing fasciitis, etc. The diagnosis of IKPLAS was judged by two physicians with senior professional titles in the clinic. Both physicians needed to be diagnosed with IKPLAS before establishing the diagnosis. Secondary observation indicators include general information (such as age, gender, comorbidities, etc.), the first laboratory (blood routine, liver and kidney function, etc.) and imaging (abdominal B-ultrasound) related indicators, treatment plans, etc. after admission. Among them, the medical history collection and routine blood test were collected on the day of admission, and the results of the first examination after admission by abdominal B-ultrasound, the treatment plan, and the prognosis were collected retrospectively after the patients were discharged from the hospital.

Data pre-processing

Statistical analysis was performed using EmpowerStats software and Python 3.9, and the procalcitonin with too many missing values (number of missing values \geq 30%) was deleted. Multiple imputations were performed for C-reactive protein, triglyceride, and cholesterol with a few missing values (number of missing values \leq 30%) using the miceforest package in Python. Since different indicators are not comparable due to their different dimensions, we use the Z-score method to standardize continuous variables. The formula is:

$$z = \frac{\chi - u}{\sigma}$$

Where μ is the average of the continuous variable across all samples, and α is the standard deviation. The influence of dimensions on the data can be eliminated after data standardization. K-S-L test and Q-Q plot were used to test the normality of measurement data. The binary variables were described as counts, and percentages were evaluated using the Chi-square test or Fisher's exact test. If the continuous variables conformed to a normal distribution, they were compared using a t-test and expressed as means \pm SEM. For a non-normal distribution, the Mann–Whitney U test was used. *P*<0.05 was considered statistically significant.

Model training and evaluation

This research uses the python3.9 version, anaconda3 integrated development environment. Based on the train_test_split module, the parameter is set to test_ size = 0.3, and the complete data is divided into a training set of 149 cases and a test set of 64 cases by stratified random sampling in a ratio of 7:3. This study used recursive feature elimination (RFE) for feature screening [17].

RFE can effectively eliminate the redundancy between features and select the optimal feature combination. It takes the prediction accuracy as the evaluation standard and eliminates the minimum relevant variables through each iteration. Then cross-validation is used to find the optimal number of features. In this study, random forest (RF) was used as the primary classifier for RFE, and feature selection was performed on the training set. The Scikit-learn python software package was used to build seven machine learning prediction models. The logistic regression model(LR) [18] was selected for the linear model. The Multilayer Perceptron (MLP) [19] model, also called artificial neural network (ANN), was chosen as an essential nonlinear prediction model. For the kernelbased model, Support Vector Machine (SVM) [20] with Gaussian kernel (RBF) was selected. For the decision tree approach, the random forest(RF) [21] model, the Decision Tree (DT) [22]model and the XGBoost(XGB) [23] model have also been used in clinical research. Finally, we chose a basic prediction model, the K-Nearest Neighbor algorithm (KNN) [24]. After the model was established, Bayesian optimization algorithm was used to find the maximum model Area Under Curve(AUC) value according to the Settings for parameter optimization.

The specific optimized parameters were the C value of LR model, max_depth, min_samples_split, min_samples_ leaf, min_weight_fraction_leaf of DT model, and max_ depth, min_samples_leaf of LR model, n_estimators, max_features, max_depth, min_weight_fraction_leaf of RF model, n_estimators, max_leaves, max_depth, max_ bin of XGB model, C-value and gamma of SVM model, hidden1, hidden2, learning_rate_int of ANN model and n_neighbors of KNN model.A fivefold cross-validation method was used to evaluate the model's generality in the training set. The model performance was evaluated using the test set, and the evaluation indicators were accuracy, precision, specificity, sensitivity (recall), F1-score, confusion matrix and AUC. A schematic overview of the study design and model development is depicted in Fig. 1.

Results

Patients and variables

After screening by inclusion and exclusion criteria, 213 patients were included in this study, all in line with the diagnosis of type 2 diabetes mellitus and Klebsiella pneumoniae liver abscess. Patients were grouped by the occurrence of IKPLAS, with 25 cases progressing to IKPLAS as the IKP-LAS group and 188 cases as the NIKPLAS group. There

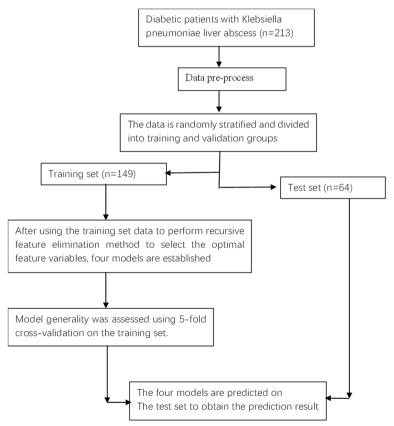


Fig. 1 Overview of study design and model development

Table 1 Baseline statistics for 213 patients Line 143

Characteristics	ALL(N=213)	NIKLAS (<i>N</i> = 188)	IKLAS (N=25)	<i>P</i> -value
Clinical findings				
AGE	61.5 ± 12.2	61.43 ± 12.05	61.92 ± 13.62	0.851
Gender				0.150
Female	60 (28.2%)	56 (29.79%)	4 (16.00%)	
Male	153 (71.8%)	132 (70.21%)	21 (84.00%)	
Smoke	30 (14.1%)	24 (12.77%)	6 (24.00%)	0.129
Alcohol	23 (10.8%)	18 (9.57%)	5 (20.00%)	0.115
DM(Year)	3.0 (2.0–10.0)	3.0 (2.0–9.2)	5.0 (2.0–10.0)	0.250
Underlying disease				
Biliary diseases	46 (21.6%)	39 (20.74%)	7 (28.00%)	0.408
CHD	19 (8.9%)	18 (9.57%)	1 (4.00%)	0.358
Liver diseases	16 (7.5%)	15 (7.98%)	1 (4.00%)	0.478
Symptom at presentation		() () () () ()	. (0.170
Body temperature	38.6 ± 1.1	38.62 ± 1.09	38.61 ± 1.01	0.962
Weakness	82 (38.5%)	62 (32.98%)	20 (80.00%)	< 0.001
Diarrhea	8 (3.8%)	5 (2.66%)	3 (12.00%)	0.021
Vomiting	22 (10.3%)	20 (10.64%)	2 (8.00%)	0.684
Abdominal pain	80 (37.6%)	72 (38.30%)	8 (32.00%)	0.541
Chills	91 (42.7%)	72 (38.30%) 78 (41.49%)	13 (52.00%)	0.541
Admission data	91 (42.770)	78 (41:49%)	13 (32.00%)	0.580
SBP	125.5±18.0	125.18±16.93	127.72 ± 25.23	0.510
DBP	75.4±10.6	75.25 ± 9.87	76.92 ± 15.32	0.310
GLU	10.3±3.6	10.03±3.38	12.67±4.17	< 0.001
WBC	11.6 (8.9–15.4)	11.6 (8.8–15.1)	12.5 (9.3–19.4)	0.055
NE	9.9 (7.2–13.4)	9.8 (7.1–13.1)	10.6 (7.8–17.9)	0.159
HB	116.0 (105.0–128.0)	117.0 (106.8–130.0)	106.0 (97.0–114.0)	< 0.001
PLT	191.0 (119.0–273.0)	205.0 (124.0–290.0)	125.0 (51.0–182.0)	0.003
ALT	57.0 (35.0–93.0)	57.0 (35.0–93.0)	58.0 (27.3–90.9)	0.509
AST	40.0 (25.7–78.2)	39.0 (25.6–73.2)	52.0 (26.0–113.0)	0.207
ALP	155.0 (103.0–240.0)	146.5 (103.0–240.0)	184.0 (133.0–239.0)	0.116
LDH	221.0 (176.0–299.0)	219.5 (176.8–289.2)	265.0 (170.0–343.0)	0.052
ALB	29.5 (26.7–33.2)	29.9 (27.2–33.5)	27.3 (24.1–29.9)	0.021
TBIL	12.9 (8.6–21.4)	12.6 (8.5–19.8)	15.8 (9.8–31.1)	0.013
DBIL	6.5 (4.2–10.9)	6.2 (4.0–10.4)	9.5 (5.6–13.5)	0.009
IBIL	5.7 (3.8–9.2)	5.7 (3.8–8.4)	5.7 (3.8–11.7)	0.209
BUN	5.0 (3.6–7.5)	4.9 (3.6–7.3)	5.8 (4.1–9.3)	0.082
TC	3.2 (2.6–3.9)	3.3 (2.7–3.9)	2.7 (2.2–3.1)	0.014
TG	1.4 (1.0–1.9)	1.4 (1.0–2.0)	1.3 (1.2–1.8)	0.755
CRP	114.0 (69.7–184.6)	112.5 (69.7–173.4)	163.0 (72.3–222.4)	0.019
PT	13.2 (12.4–14.1)	13.1 (12.3–13.9)	13.6 (13.0–15.3)	0.017
D.DIMER	2.5 (1.2–4.7)	2.2 (1.2–4.2)	4.4 (3.7–8.2)	< 0.001
MDRO	13 (6.1%)	10 (5.32%)	3 (12.00%)	0.190
SOFA	1.3 ± 2.3	0.87 ± 1.50	4.88 ± 3.66	< 0.001
Radiologic findings				
Abscess location				0.672
Right lobe	160 (75.1%)	140 (74.47%)	20 (80.00%)	
Left lobe	39 (18.3%)	36 (19.15%)	3 (12.00%)	
Both lobes	14 (6.6%)	12 (6.38%)	2 (8.00%)	
Abscess size (cm)	7.3 ± 2.5	7.41 ± 2.59	6.81 ± 2.04	0.271

Characteristics	ALL(N=213)	NIKLAS (<i>N</i> = 188)	IKLAS (<i>N</i> =25)	P-value	
No. of abscesses				0.151	
Multiple	52 (24.4%)	43 (22.87%)	9 (36.00%)		
Solitary	161 (75.6%)	145 (77.13%)	16 (64.00%)		
Treatment					
Drainage mode				0.003	
None	32 (15.0%)	23 (12.23%)	9 (36.00%)		
Catheterization	135 (63.4%)	120 (63.83%)	15 (60.00%)		
Puncture	40 (18.8%)	40 (21.28%)	0 (0.00%)		
Surgery	6 (2.8%)	5 (2.66%)	1 (4.00%)		

Table 1 (continued)

were 60 females and 153 males, as shown in Table 1. Through stratified random sampling, the data set was divided into the training set and test set. As shown in Table S1, there was no statistically significant difference between training set and test set($P \ge 0.05$). Clinical findings, Symptom at presentation, Admission data, and Radiologic findings in Table S1 will all be screened as variables. As shown in Fig. 2, When the number of feature variables is four, the recursive feature elimination method with random forest as classifier has the highest cross validation score. These four variables are hemoglobin, platelets, D-dimer, and SOFA score.Spearman correlation analysis was performed on these four features, as shown in Fig. 3, indicating no highly correlated redundant features.

Tuning of parameters

The four variables selected from the training set were put into the machine learning classifier to construct the prediction model. Through Bayesian algorithm optimization, the parameters were adjusted with the average optimal AUC value, and the specific parameter Settings are shown in Table S2. The five-fold cross-validation ROC curve of the training set can be seen in Fig. 4, where it can be seen that the SVM model and LR model have better performance.

Evaluation of prediction models

The ROC curve of the test set can be seen in Fig. 5. The AUC values of most models are higher than 0,850, among which SVM (0.969) and LR (0.967) rank the top two, but the AUC values of XGB (0.799) and DT (0.800) are lower. Studies have shown that precision recall curve (PRC) has advantages over ROC in evaluating imbalanced datasets [25]. The dataset included in this study is also imbalanced, so PRC is also a valuable indicator. Figure 6 shows the PRC of the test set, and the Average Precision(AP) value was used as a criterion to evaluate the PR curve

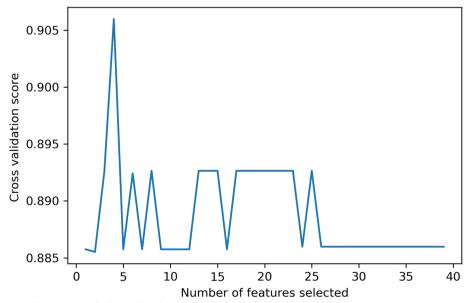


Fig. 2 Recursive feature elimination method variable selection diagram

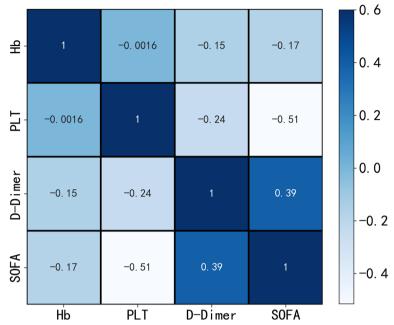


Fig. 3 Spearman Correlation Analysis Heatmap

[26]. The APs of the LR,SVM models were all above 0.800. The confusion matrix was also calculated for all seven models (Table 2), and the DT model generated a large number of FPs (n=19) during the prediction process, while the other models were relatively few. DT, LR, and SVM models produced the least FNs (n=1), and the KNN model produced the least FPs (n=0). Table 3 shows each model evaluation result's sensitivity (recall), specificity, accuracy, precision, f1, AP and AUC.

As shown in Table 3, there were significant performance differences between the models. The AUC (0.969), F1-Score(0.737) and AP(0.890) of the SVM model were the highest among the seven models, and the all-around performance was the best. At the same time, its sensitivity (0.875) is the highest and can effectively identify the occurrence of IKPLAS in the early stage. The KNN model had the best specificity (1.000) and could be used to reduce the occurrence of overdiagnosis and treatment.

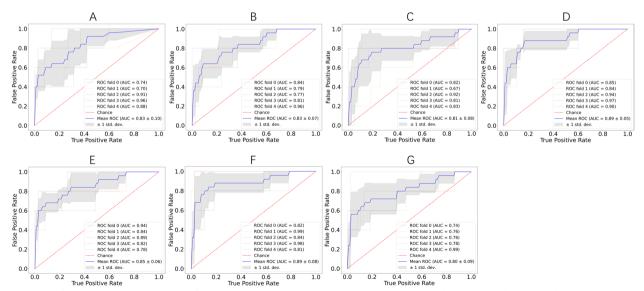


Fig. 4 Five-fold cross-validation ROC curve for the training set. A ANN model. B DT model. C KNN model. D LR model. E RF model. F SVM model. G XGB model

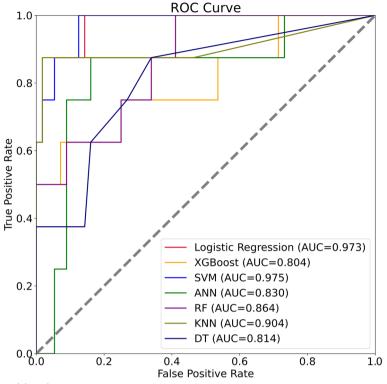


Fig. 5 ROC curves of seven models in the test set

Figure 7 shows the calibration curves of the seven models. Except that the XGB and DT models over-estimates the occurrence of IKPLAS risk, the other models' calibration curves are a good fit with the actual observed results.

Figure 8 shows the Decision Curve Analysis of the seven models, which was first proposed in 2006 and

has been used for prognostic decision analysis in cancer [27] and other fields [23]. The DCA curve shows a model compared to the Net Benefit situation under different High-Risk Thresholds between the two strategies of intervention in all patients (ALL) and no intervention in all patients (NONE). As shown in Fig. 6, there is no

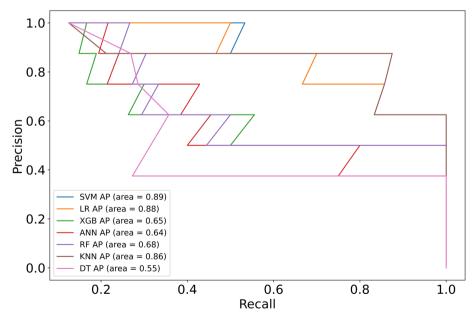


Fig. 6 Precision Recall Curves for the seven models in the test set

Table 2 Confusion matrices of 7 models

Confusion matrix	Actual	Prediction			
		Negative	Positive		
SVM	Negative	52	4		
	Positive	1	7		
LR	Negative	49	7		
	Positive	1	7		
RF	Negative	53	3		
	Positive	4	4		
XGB	Negative	55	1		
	Positive	4	4		
ANN	Negative	51	5		
	Positive	5	3		
DT	Negative	37	19		
	Positive	1	7		
KNN	Negative	56	0		
	Positive	6	2		

significant difference in the benefits of treatment intervention based on SVM and LR model between the risk threshold of 0.0 and 0.4. However, when the risk threshold was between 0.4 and 0.8, the SVM model's net intervention rate was significantly higher than that of other models and the overall benefit rate was high. Model explanation.

To explain the output of our models, we used the SHapley Additive exPlanations(SHAP) algorithm to help us understand how a single feature affects the output of the models [28]. Its most significant advantage is that it can reflect the influence of the features in each sample, and it also shows the positive and negative effects of the influence. Each row represents a feature, sorted by feature importance from top to bottom. The abscissa is the SHAP value. A point represents a sample, and the color represents the eigenvalue (red for high, blue for low). The SVM prediction model with the best all-around performance was selected to interpret the feature importance.

As shown in Fig. 9, the SOFA score ranks first in the feature importance of SVM model, and the higher the value, the higher the probability of the patient progressing to IKPLAS. Platelet and hemoglobin were the second and third most important predictors of the SVM model, and both were negatively correlated with the outcome. D-dimer ranked last and was positively associated with the risk of IKPLAS.

Discussion

The high incidence of IKPLAS is mainly in the Asian population, which may be related to the fact that the Asian population is more likely to colonize the intestine with K1/K2 serotype Klebsiella pneumoniae [29, 30]. Diabetes is considered a significant risk factor for IKPLAS, and up to 63% of patients with a bacterial liver abscess in Taiwan have diabetes. This may be related to the impaired phagocytosis of K1/K2 Klebsiella pneumoniae in diabetic patients [31] and the more excellent vascular permeability in diabetic patients, which is conducive to bacterial invasion [11]. The above two serotypes of Klebsiella pneumoniae are also highly virulent Klebsiella pneumoniae, which show high viscosity in the String test [9]. Although the highly virulent Klebsiella pneumoniae is sensitive to most antibiotics, patients often have a poor prognosis if they are not recognized and treated early [32].

This study screened four characteristic variables: hemoglobin, platelets, D-dimer, and SOFA score. We interpreted the importance of the model characteristic variables by using the SHAP package, in which the SOFA score ranked first among all four models.

The SOFA score is a scoring system that measures the degree of impairment of significant organ function in patients with sepsis or suspected sepsis to determine prognosis [33]. Several studies have confirmed its predictive value in the prognosis of infected patients [34, 35]. This study also suggests that the SOFA score is a significant predictor of diabetes complicated by IKPLAS. As can be seen from the SHAP plot, the higher the SOFA score, the greater the risk of progression to IKPLAS.

Model	Sensitivity (recall)	Specificity	Accuracy	Precision	F1-Score	AP	AUC
SVM	0.875	0.929	0.922	0.636	0.737	0.890	0.969
LR	0.875	0.875	0.875	0.500	0.636	0.880	0.967
RF	0.500	0.946	0.891	0.571	0.533	0.680	0.879
XGB	0.500	0.982	0.922	0.800	0.615	0.650	0.799
ANN	0.375	0.911	0.844	0.375	0.375	0.640	0.897
DT	0.875	0.661	0.688	0.269	0.412	0.620	0.800
KNN	0.250	1.000	0.906	1.000	0.400	0.860	0.900

Table 3 Performance summary in terms of sensitivity (recall), specificity, accuracy, precision,F1-score,AUC

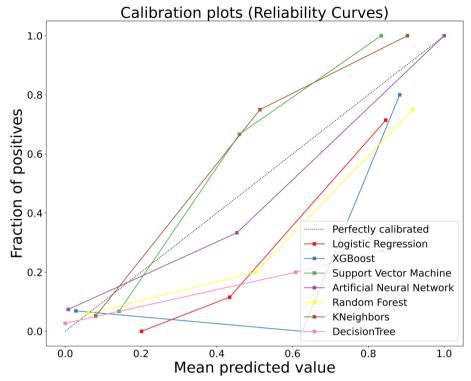


Fig. 7 Seven machine learning model calibration curves

Although the pathogenesis of IKPLAS is currently unclear, the study by Chen-Guang Zhang et al. shows that most diabetic patients with IKPLAS are prone to sepsis [11]. Blood-borne transmission may be one of the more important ways.

In the feature importance ranking, platelets' influence on SVM model ranked second.Jai Hoon Yoon et al. showed that thrombocytopenia is an independent risk factor for invasive syndrome in diabetic patients with Klebsiella pneumoniae liver abscess [10]. This is also consistent with the conclusions about platelets in the SVM model established in this study. The mechanism of platelet reduction in diabetes combined with IKPLAS may be that when the body is infected, platelets are stimulated and activated to participate in the body's inflammatory response by inducing the expression of membrane proteins and the production of mediators and play the role of anti-infection and pathogen removal. Activated platelets produce and release pro-inflammatory, anti-inflammatory, chemokines, antimicrobial, and other mediators to regulate the body's innate immune or adaptive immune response [36]. The interaction between platelets and pathogens or their products, endothelial cells, and immune cells promotes endothelial cell damage and leukocyte activation. As a result, the adhesion of platelets to it is enhanced, platelets are continuously activated in the circulation, and the body continuously produces anti-platelet antibodies and macrophage-colony stimulating factors, which accelerates the destruction and consumption of platelets [37].

The SHAP plot shows that hemoglobin is the third most important characteristic variable after the SOFA score, and the lower its value, the higher the risk of progression to IKPLAS. It has been shown that hemoglobin can be an indicator to assess the severity of the disease in infected patients, probably due to a systemic inflammatory response leading to decreased erythropoiesis, increased destruction of erythrocytes due to hemolysis, and hemorrhage, which leads to a reduced ability of blood to transport oxygen and carbon dioxide and insufficient oxygen supply to the body, resulting in multiorgan damage [38].

D-dimer is a specific molecular marker for secondary hyperfibrinolysis in vivo and is an effective indicator to reflect the coagulation state of the body. The coagulation and fibrinolytic systems are usually closely linked to the development of inflammation. Infection can lead to damage of vascular endothelial cells and alveolar epithelial cells, which stimulates the coagulation system, resulting in impairment of coagulation function and abnormal coagulation indexes in patients, further aggravated by elevated D-dimer along with infection [39, 40]. The above two promote each other, forming a vicious circle. The autoimmune function of diabetic

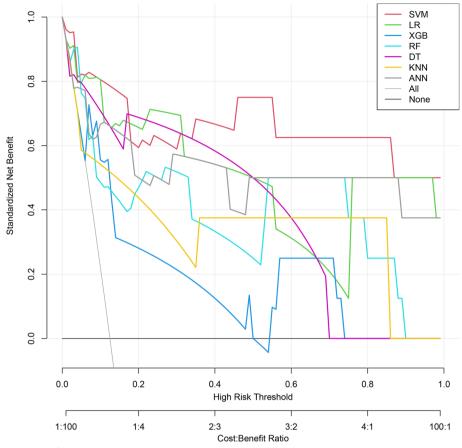


Fig. 8 Decision curve analysis of seven machine learning models

patients is weakened, and the inflammatory response is enhanced after infection. Patients with diabetes complicated with IKPLAS can have noticeable D-dimer changes in the early stage. In the SVM model, D-dimer was positively associated with the risk of developing diabetes with IKPLAS, which is consistent with the above findings. In the field of IKPLAS, more studies are focused on the risk factors of IKPLAS. The study by Shixiao Li et al. [41] showed that patients with IKPLAS were more likely to develop chronic renal insufficiency, thrombocytopenia, and increased total bilirubin than patients with non-IKPLAS. Hairui Wang et al. [42]. A logistic regression prediction model was used to predict

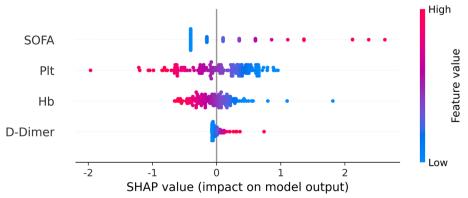


Fig. 9 SHAP feature analysis of the SVM model

the incidence of IKPLAS by incorporating clinical and CT features, with an AUC value of 0.842 in the validation set, and did not compare other prediction models.Unlike many studies, we first used seven machine learning models for prediction. Through parameter adjustment and verification, the SVM model with the best performance was selected, with an AUC value of 0.969 and an AP value of 0.890, indicating that it was a reliable IKPLAS prediction model. At the same time, the variables included in this model are clinical indicators, which are easy to collect and can be used by clinicians to conveniently judge the possibility of IKPLAS in patients with diabetes mellitus complicated with Klebsiella pneumoniae liver abscess.

Machine learning algorithms can build complex models that perform satisfactorily enough when the amount of data is sufficient. However, in specific applications, the amount of data is often insufficient, so it is essential to analyze these machine learning algorithms and obtain good results with relatively small sample sizes. In this study, the Power analysis was satisfied by calculating a power value of > 0.80, although we only used a small data set of 213 patients. The main reason for the excellent performance of the SVM model in this study is that it is a nonlinear learner that is more suitable for small samples, can ideally separate samples, and has better generalization.

There are still some limitations in this study. First, this is a single-center regression study, and some potential biases cannot be avoided. Secondly, for machine learning, the sample size of this study is insufficient. In order to further improve the accuracy of the model, we will collect more clinical data and further optimize the parameters.

Conclusion

In this study, we established and compared seven models to compare the performance of predicting the progression of diabetes with Klebsiella pneumoniae to IKPLAS and found that the SVM model had the highest overall predictive power. We also found that SOFA score, platelets, hemoglobin, and D-dimer significantly affected the model's predictions. In the future, we will expand the dataset to improve further the model's accuracy and better plan diagnosis and treatment for clinicians.

Supplementary Information

The online version contains supplementary material available at https://doi.org/10.1186/s12879-023-08235-7

Additional file 1: Table S1. Characteristics of patients in the training and test sets.

Additional file 2: Table S2. Tuning parameters of the predictive models.

Acknowledgements

Not applicable.

Authors' contributions

CY F and JD had roles in the study design, data analysis, literature search, and writing of the manuscript. SF J and FH had roles in research guiding, and clinical management. CY F and XM L had roles in datacollection, and data interpretation. All authors have read and agreed with the final manuscript.

Funding

No fund was received for this study.

Availability of data and materials

The datasets used and analyzed during the current study are available from the corresponding author on reasonable request.

Declarations

Ethics approval and consent to participate

This study was carried out in accordance with the Declaration of Helsinki and approved by the Ethics Committee of the Third Affiliated Hospital of Soochow University, China (registration number 2021/166). The Ethics Committee of the Third Affiliated Hospital of Soochow considered that informed consent of patients was waived for participation in this observational study.

Consent for publication

Not applicable.

Competing interests

The authors declare no competing interests.

Received: 10 October 2022 Accepted: 9 April 2023 Published online: 04 May 2023

References

- 1. Meddings L, Myers RP, Hubbard J, et al. A population-based study of pyogenic liver abscesses in the United States: incidence, mortality, and temporal trends. Am J Gastroenterol. 2010;105(1):117–24.
- Li W, Chen H, Wu S, et al. A comparison of pyogenic liver abscess in patients with or without diabetes: a retrospective study of 246 cases. BMC Gastroenterol. 2018;18(1):144.
- Zhang J, Gao Y, Du Z, et al. Clinical Features and Prognosis of Gas-Forming and Non-Gas-Forming Pyogenic Liver Abscess: A Comparative Study. Surg Infect (Larchmt). 2021;22(4):427–33.
- Song H, Wang X, Lian Y, et al. Analysis of the clinical characteristics of 202 patients with liver abscess associated with diabetes mellitus and biliary tract disease. J Int Med Res. 2020;48(8):300060520949404.
- Tsai FC, Huang YT, Chang LY, et al. Pyogenic liver abscess as endemic disease, Taiwan. Emerg Infect Dis. 2008;14(10):1592–600.
- 6. Akhondi H, Sabih D E. Liver Abscess [M]. StatPearls. Treasure Island (FL). 2022.
- Luo M, Yang XX, Tan B, et al. Distribution of common pathogens in patients with pyogenic liver abscess in China: a meta-analysis. Eur J Clin Microbiol Infect Dis. 2016;35(10):1557–65.
- Lin YT, Wang FD, Wu PF, et al. Klebsiella pneumoniae liver abscess in diabetic patients: association of glycemic control with the clinical characteristics. BMC Infect Dis. 2013;13:56.
- 9. Jun JB. Klebsiella pneumoniae Liver Abscess. Infect Chemother. 2018;50(3):210–8.
- Yoon JH, Kim YJ, Jun YH, et al. Liver abscess due to Klebsiella pneumoniae: risk factors for metastatic infection. Scand J Infect Dis. 2014;46(1):21–6.
- 11. Zhang CG, Wang Y, Duan M, et al. Klebsiella pneumoniae invasion syndrome: a case of liver abscess combined with lung abscess, endophthalmitis, and brain abscess. J Int Med Res. 2022;50(3):3000605221084881.
- Kassam NM, Aziz OM, Somji SS, et al. Invasive liver abscess syndrome caused by Klebsiella pneumoniae: first Tanzanian experience. Pan Afr Med J. 2020;36:191.

- Vila A, Cassata A, Pagella H, et al. Appearance of Klebsiella pneumoniae liver abscess syndrome in Argentina: case report and review of molecular mechanisms of pathogenesis. Open Microbiol J. 2011;5:107–13.
- 14. Van Calster B, Wynants L. Machine Learning in Medicine. N Engl J Med. 2019;380(26):2588.
- Eaton JE, Vesterhus M, Mccauley BM, et al. Primary sclerosing cholangitis risk estimate tool (PREsTo) predicts outcomes of the disease: a derivation and validation study using machine learning. Hepatology. 2020;71(1):214–24.
- Bae S, Choi YS, Ahn SS, et al. Radiomic MRI phenotyping of glioblastoma: improving survival prediction. Radiology. 2018;289(3):797–806.
- Darst BF, Malecki KC, Engelman CD. Using recursive feature elimination in random forest to account for correlated variables in high dimensional data. BMC Genet. 2018;19(Suppl 1):65.
- Meurer WJ, Tolles J. Logistic Regression Diagnostics: Understanding How Well a Model Predicts Outcomes. JAMA. 2017;317(10):1068–9.
- 19. Yan J, Wang X, Cai J, et al. Medical image segmentation model based on triple gate MultiLayer perceptron. Sci Rep. 2022;12(1):6103.
- Wang H, Shao Y, Zhou S, et al. Support vector machine classifier via L(0/1) Soft-Margin Loss. IEEE Trans Pattern Anal Mach Intell. 2022;44(10):7253–65.
- 21. Rigatti SJ. Random forest. J Insur Med. 2017;47(1):31-9.
- 22. Flayer CH, Perner C, Sokol CL. A decision tree model for neuroimmune guidance of allergic immunity. Immunol Cell Biol. 2021;99(9):936–48.
- Hou N, Li M, He L, et al. Predicting 30-days mortality for MIMIC-III patients with sepsis-3: a machine learning approach using XGboost. J Transl Med. 2020;18(1):462.
- 24. Gao S, Cai T, Fang K. Gravity-matching algorithm based on k-nearest neighbor. Sensors (Basel). 2022;22(12):4454.
- 25. Fu GH, Yi LZ, Pan J. Tuning model parameters in class-imbalanced learning with precision-recall curve. Biom J. 2019;61(3):652–64.
- 26. Saito T, Rehmsmeier M. The precision-recall plot is more informative than the ROC plot when evaluating binary classifiers on imbalanced datasets. PLoS ONE. 2015;10(3):e0118432.
- Hu C, Yang J, Huang Z, et al. Diagnostic and prognostic nomograms for bone metastasis in hepatocellular carcinoma. BMC Cancer. 2020;20(1):494.
- Lundberg SM, Erion G, Chen H, et al. From local explanations to global understanding with explainable AI for trees. Nat Mach Intell. 2020;2(1):56–67.
- Kim JH, Jeong Y, Lee CK, et al. Characteristics of Klebsiella pneumoniae isolates from stool samples of patients with liver abscess caused by hypervirulent K. pneumoniae. J Korean Med Sci. 2020;35(2):e18.
- Zhang S, Zhang X, Wu Q, et al. Clinical, microbiological, and molecular epidemiological characteristics of Klebsiella pneumoniae-induced pyogenic liver abscess in southeastern China. Antimicrob Resist Infect Control. 2019;8:166.
- Zheng S, Florescu S, Mendoza M. Klebsiella pneumoniae invasive syndrome in a diabetic patient with gallbladder abscess. Clin Case Rep. 2020;8(10):1940–2.
- Kim JK, Chung DR, Wie SH, et al. Risk factor analysis of invasive liver abscess caused by the K1 serotype Klebsiella pneumoniae. Eur J Clin Microbiol Infect Dis. 2009;28(1):109–11.
- Esposito S, De Simone G, Boccia G, et al. Sepsis and septic shock: New definitions, new diagnostic and therapeutic approaches. J Glob Antimicrob Resist. 2017;10:204–12.
- Gupta T, Puskarich MA, Devos E, et al. Sequential organ failure assessment component score prediction of in-hospital mortality from sepsis. J Intensive Care Med. 2020;35(8):810–7.
- Pawar RD, Shih JA, Balaji L, et al. Variation in SOFA (Sequential Organ Failure Assessment) score performance in different infectious States. J Intensive Care Med. 2021;36(10):1217–22.
- 36. Li C, Li J, Ni H. Crosstalk between platelets and microbial pathogens. Front Immunol. 1962;2020:11.
- Larkin CM, Santos-Martinez MJ, Ryan T, et al. Sepsis-associated thrombocytopenia. Thromb Res. 2016;141:11–6.
- Muady GF, Bitterman H, Laor A, et al. Hemoglobin levels and blood transfusion in patients with sepsis in internal medicine departments. BMC Infect Dis. 2016;16(1):569.

- Duarte JC, Tavares E Castro A, Silva R, et al. Prognostic value of plasma level in adults with community-acquired pneumonia a prospective study. Rev Port Pneumol (2006). 2015;21(4):218–9.
- NastasijevicBorovac D, Radjenovic Petkovic T, Pejcic T, et al. Role of D-dimer in predicting mortality in patients with community-acquired pneumonia. Med Glas (Zenica). 2014;11(1):37–43.
- Li S, Yu S, Peng M, et al. Clinical features and development of Sepsis in Klebsiella pneumoniae infected liver abscess patients: a retrospective analysis of 135 cases. BMC Infect Dis. 2021;21(1):597.
- Wang H, Guo Y, Yan B, et al. Development and validation of a prediction model based on clinical and CT features for invasiveness of K. pneumoniae liver abscess. Eur Radiol. 2022;32(9):6397–406.

Publisher's Note

Springer Nature remains neutral with regard to jurisdictional claims in published maps and institutional affiliations.

Ready to submit your research? Choose BMC and benefit from:

- fast, convenient online submission
- thorough peer review by experienced researchers in your field
- rapid publication on acceptance
- support for research data, including large and complex data types
- gold Open Access which fosters wider collaboration and increased citations
- maximum visibility for your research: over 100M website views per year

At BMC, research is always in progress.

Learn more biomedcentral.com/submissions

