Original Article

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# **Use of Artificial Intelligence in Pulmonary Embolism Prediction**

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#### **Abstract**

Aim: The purpose of this study was to use artificial intelligence to predict the risk of pulmonary embolism (PE) in patients with suspected PE admitted to the emergency room based on physical examination, laboratory, and clinical probability prediction scores without computed tomography angiography.

Materials and Methods: A comprehensive analysis was conducted on a total of 156 individuals who were admitted to the emergency room with PE. Seventy-eight patients were diagnosed with PE through anamnesis, physical examination, clinical likelihood prediction scores, investigations, and imaging. These patients were then included in the PE group. The data set includes gender, age, shock index, vital signs, complaints at arrival to the emergency department, comorbidities, medications used, medical history, radiological examinations, presence of deep vein thrombosis, electrocardiography, echocardiography findings, Wells score, Geneva score, PERC score, and laboratory tests performed.

Results: The average age of the patients in the study was 69.46±15 years. Dyspnea was the most prevalent presentation, affecting 88 patients (56.4%). The most prevalent comorbidities were hypertension in 52 patients (33.1%), cancer in 51 patients (32.7%), and coronary artery disease in 35 patients (22.4%). The Wells score, D-dimer, low partial carbon dioxide pressure, and tachycardia were discovered to be important factors in the diagnosis of PE. Statistically significant parameters were investigated using a multilayer perceptron artificial intelligence model. The diagnosis of PE was correct with 96% accuracy and 89% specificity.

**Conclusion:** According to the findings of our study, a thorough review of the patient's anamnesis, physical examination, laboratory and imaging data, and the application of scores are all crucial in the diagnosis of PE. Furthermore, it was determined that artificial intelligence can be used to diagnose PE before using imaging modalities.

Keywords: Artificial intelligence, diagnostic algorithm, pulmonary embolism

# Introduction

Venous thromboembolism (VTE) consists of deep vein thrombosis (DVT) and pulmonary embolism (PE). PE is a clinical condition that occurs when a thrombus passes from the venous circulation to the pulmonary arteries and clots. The clinical presentation varies from asymptomatic to fatal. For this reason, it is difficult to determine the true incidence of PE. Nevertheless, the incidence has increased over the years (1). PE is a critical condition that, along with myocardial infarction and stroke, is among the leading causes of cardiovascular-related death (2). PE-related

mortality may vary depending on the patient's age, comorbid diseases, disease burden, and duration of effective treatment. Thirty-day all-cause mortality in patients with PE is 6.6%. PErelated seven-day mortality was 1.1%, while thirty-day mortality was 1.8% (3). The annual cost of PE to the European Union countries was found to be 8.5 billion Euros, including indirect costs such as pre-hospital prevention, in-hospital treatment, and post-hospital care. Both the aging of the population, the increase in incidence, and the decrease in mortality will increase the financial burden of VTE events on governments in Europe and other countries of the world (4).

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Although PE is a common disease, there are no pathognomonic findings or diagnostic tests. For this reason, the clinician should be the primary authority for making the diagnosis. It is difficult for the clinician to diagnose the disease as it has a wide range of clinical presentations, from asymptomatic to fatal outcomes. It is emphasized that PE can be fatal if there are delays in diagnosis and treatment (5). The relatively high prevalence of PE makes it a common and potentially life-threatening disease (6). Early diagnosis of PE is crucial, as even patients with minor symptoms are at risk of recurrent PE (7). Currently, the gold standard diagnostic method is computed tomography-pulmonary angiography (CTPA) (8). Because of the risks associated with CTPA, including contrast agent allergy, contrast nephropathy, radiation exposure, and economic reasons, diagnostic algorithms have been proposed and clinical probability prediction scores have been developed to diagnose PE before imaging (9-15). Two of these scores are the Wells Clinical score and the Geneva score. The Wells clinical score is a widely recognized and validated tool for assessing the clinical probability of PE. It includes physical findings and risk factors such as DVT, lack of alternative diagnoses, tachycardia, immobilization or recent surgery, history of DVT or PE, hemoptysis, and malignancy (16). To help diagnose PE, the Geneva score, like the Wells score, is a standardized tool to help determine the clinical probability of PE based on several criteria, including heart rate, clinical signs of DVT, hemoptysis, and previous PE or DVT (17).

With the increasing awareness of PE among physicians and the increasing availability of diagnostic tests and imaging, the need to avoid unnecessary tests has become evident. The aim was to avoid complications of the tests and to reduce the excessive cost and length of hospitalization. For this purpose, they defined "pulmonary embolism exclusion criteria (PEEC)." It is a rule based on clinical criteria to exclude this condition in patients suspected of having it. The PEEC rule aims to prevent unnecessary additional testing in low-risk patients by assessing whether patients have certain clinical characteristics. Besides all these algorithms, only D-dimer has been validated as a biomarker to aid in the decision to exclude PE. Although not specific for PE, elevated white blood cell count, serum lactate dehydrogenase (LDH), C-reactive protein (CRP), aspartate aminotransferase (AST), and increased sedimentation rate may be detected. The diagnosis of PE plays a critical role in the management of this life-threatening condition, alongside the use of many methods and algorithms. The use of advanced imaging techniques such as CTPA and the application of algorithms, together with a high index of suspicion and rapid intervention, is essential in providing a timely and accurate diagnosis that can significantly affect patient outcomes. In this study, our aim is to investigate the feasibility of using artificial intelligence (AI) approaches in the diagnosis of PE; to identify possible risk factors; and to ensure that CTPA, the gold standard in the diagnosis of PE, is used in appropriate patients based on Al findings.

### **Materials and Methods**

The appropriateness of this study was approved by the İnönü University Scientific Research and Publication Ethics Committee with (decision number: 2022/45, date: 20.04.2022). In addition, the study was supported by İnönü University Scientific Research Projects Unit with project number 3002.

#### **Dataset**

In this study, 156 patients admitted to the Department of Emergency Medicine of İnönü University Faculty of Medicine Turgut Özal Medical Center from 13.10.2022-14.10.2024, with PE symptoms, were prospectively analyzed. Adult patients presenting to the emergency department with PE symptoms were included in the study. Pediatric patients under 18 years of age, as well as pregnant and recently delivered patients, were excluded. Anamnesis, physical examination, computerized order tracking system (COTS), and laboratory tests were evaluated. All patients underwent bolus-tracking pulmonary angiography, the gold standard imaging method in PE. Seventy-eight patients were diagnosed with PE and then enrolled in the PE group. The 78 patients with alternative diagnoses in whom PE was ruled out were enrolled as the control group. In the medical records of patient admissions, the admission number, name-surname, gender, age, shock index, vital signs (temperature, pulse, systolic and diastolic blood pressures, saturation values), complaints at presentation to the emergency department, comorbidities, medications used, medical history, radiological examinations, presence of DVT, electrocardiography (ECG), echocardiography findings [ejection fraction (EF)], pulmonary artery pressure (PAP), right ventricular volume (RVV), Wells score, Geneva score, PERC score, laboratory tests hemoglobin, hematocrit, mean cellular volume, monocyte count, platelet, activated partial thromboplastin time, international normalized ratio values, CRP, prothrombin time, platelet distribution width, erythrocyte distribution width, liver enzymes [alanine aminotransferas, AST, creatine kinase (CK), CK myocardial band, renal function tests [blood urea nitrogen (BUN) and creatinine], total protein, albumin, LDH, triglycerides, cholesterol, low-density lipoprotein (LDL), blood gas parameters (pH, PCO<sub>2</sub>, PO<sub>2</sub> lactate, HCO<sub>3</sub>), D-dimer, fibrinogen, pro-brain natriuretic peptide, procalcitonin (PCT), high sensitivity troponin (HS troponin), PCT triglycerides, total cholesterol, high-density lipoprotein (HDL-cholesterol), LDL-cholesterol, plasmin, vitamin K, fibrinopeptide A, factor V Leiden, and protein S were examined.

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#### **Artificial Intelligence**

Al is increasingly being integrated into various aspects of healthcare, revolutionizing the field and providing new opportunities for better patient care and outcomes. Al applications in healthcare cover a wide range of areas, from diagnosis and treatment to administrative tasks and patient engagement. Machine learning (ML) techniques such as support vector machines, neural networks, and deep learning have been instrumental in leveraging structured and unstructured data to improve decision-making in healthcare (18). In the field of medical imaging, AI has played an important role in improving diagnostic accuracy and treatment strategies. ML algorithms have been used to predict outcomes and help analyze medical images such as magnetic resonance imaging and computed tomography scans, leading to improved diagnostic capabilities (19). The integration of AI into healthcare has been positively received by both patients and healthcare professionals, highlighting the potential benefits of AI in improving healthcare delivery and patient outcomes (20). However, it is crucial to ensure the interpretability and ethical use of AI in healthcare to protect patient safety and data privacy (21). Overall, the development of Al in healthcare holds great promise for transforming the sector, increasing diagnostic accuracy, improving treatment outcomes, and optimizing healthcare delivery processes.

# Statistical Analysis

Data analysis was performed using IBM® SPSS® Statistics (version 25 for Windows, IBM Corporation, Armonk, New York, USA). Shapiro-Wilk test, histogram distribution, and skewness-kurtosis parameters were used for normality analysis. Descriptive statistics are presented as mean ± standard deviation for variables with normal distribution, median (minimum-maximum) for variables with non-normal distribution, and count of cases and (%) for nominal variables. The chi-square test and the Fisher's exact test were used to analyze the relationship between categorical variables. In the evaluation of the relationship between continuous variables, the Mann-Whitney U test was used if the variables were nonparametric, and the Student t test was used if the variables were parametric. Results were considered statistically significant for p<0.05.

#### Al Modeling

The multilayer perceptron (MLP) artificial neural network model was used with the variables that were statistically different between the PE and control groups. Gradient descent was used as the optimization function for the model. 70% of the data was used in the training of the model, while 30% was used in the testing phase.

#### Results

#### **Biostatistical Analysis**

The mean age of the patients included in the study was  $69.46\pm15$  years. Of the patients, 79 were male (50.6%) and 77 were female (49.4%). When the presenting complaints of the patients were analyzed, 88 patients (56.4%) presented with dyspnea, 11 patients (7.1%) with palpitations, 8 patients (5.1%) with chest pain, 4 patients (2.6%) with syncope, 3 patients (1.9%) with hemoptysis, and 40 patients (25.6%) with other reasons. The general data of the patients included in the study are shown in Table 1.

Table 1. The descriptive statistics of all patients					
	Mean ± SD	Minimum- Maximum			
Age (years)	69.46±15	27-98			
Vital signs					
Systolic blood pressure	133.31±25.7	77-264			
Diastolic blood pressure	80.58±15.21	36-130			
SaO <sub>2</sub>	89.10±8.38	40-100			
Heart rate	93.68±20.64	54-161			
Fever	36.27±0.24	36-37.2			
	Count (n)	Percent (%)			
Gender	'	1			
Female	77	49.4			
Male	79	50.6			
Hospital application complaint					
Palpitations	11	7.1			
Shortness of breath	88	56.4			
Chest pain	8	5.1			
Hemoptysis	3	1.9			
Syncope	4	2.6			
Other	40	25.6			
Comorbidity					
DM	12	7.7			
HT	52	33.3			
CAD	35	22.4			
COPD	21	13.5			
CHF	23	14.7			
CRF	7	4.5			
SVE	4	2.6			
Malignite	51	32.7			
DVT	26	16.7			
PE	9	5.8			
Other	63	40.4			

DM: Diabetes mellitus, HT: Hypertension, CAD: Coronary artery disease, COPD: Chronic obstructive pulmonary disease, CHF: Congestive heart failure, CRF: Chronic renal failure, SVE: Serobrovascular events, DVT: Deep vein thrombosis, PE: Pulmonary embolism, SD: Standard deviation, EA-EY: Minimum-highest, TA: Tension arterial, SaO,: Saturation

The mean age of the patients with PE was  $68.48\pm13.4$  years, while the mean age of our control group was  $70.44\pm13.4$  years. In the PE group, 45 patients (57.7%) were female and 33 patients (42.3%) were male. In the control group, 32 patients (41%) were female and 46 patients (59%) were male. The comparison of risk factors for PE and control groups is given in Table 2.

According to Table 2, when PE patients were compared with the control group, PE patients were more likely to be female (p=0.037), and complaints such as palpitations (p=0.005) and shortness of breath (p=0.001) were more common. In terms of vital signs, PE patients had lower systolic and diastolic blood pressure

Table 2.	The	comparison	of	risk	factors	for	PE	by	group
categorie	s (PF.	control)							

	Group		
	PE (n=78)	Control (n=78)	p value
	Mean ± SD	Mean ± SD	
Age (years)	68.48±16.4	70.44±13.4	0.605
Vital signs			
Systolic blood pressure	124.83±26.58	141.69±22.08	<0.001
Diastolic blood pressure	75.94±16.06	85.16±12.85	<0.001
SaO <sub>2</sub>	89.33±6.89	88.88±9.66	0.594
Heart rate	99.89±21.59	87.55±17.77	<0.001
Fever	36.28±0.26	36.27±0.22	0.990
Gender			
Female	45 (57.7)	32 (41.0)	0.027
Male	33 (42.3)	46 (59.0)	0.037
Hospital application complaint	Count (%)	Count (%)	
Palpitations	10 (12.8)	1 (1.3)	0.005
Shortness of breath	34 (43.6)	54 (69.2)	0.001
Chest pain	6 (7.7)	2 (2.6)	0.147
Hemoptysis	2 (2.6)	1 (1.3)	0.506
Syncope	3 (3.8)	1 (1.3)	0.311
Other	21 (29.5)	19 (24.4)	0.714
Comorbidity			
DM	12 (15.4)	0 (0)	<0.001
HT	19 (24.4)	33 (42.3)	0.017
CAD	10 (12.8)	25 (32.1)	0.004
CRF	1 (1.3)	6 (7.7)	0.117
SVE	1 (1.3)	3 (3.8)	0.620
Malignite	29 (37.2)	22 (28.2)	0.232
DVT	24 (30.8)	2 (2.6)	<0.001
PE	6 (7.7)	3 (3.8)	0.495
Other	33 (42.3)	30 (38.5)	0.624

DM: Diabetes mellitus, DVT: Deep vein thrombosis, HT: Hypertension, CAD: Coronary artery disease, CRF: Chronic renal failure, PE: Pulmonary embolism, SaO<sub>2</sub>: Saturation, SD: Standard deviation, SVE: Serobrovascular events

(p<0.001) and higher heart rate (p<0.001). Comorbidities such as diabetes (p<0.001), coronary artery disease (p=0.004), and DVT (p<0.001) were more common in PE patients. These data suggest that PE patients differ from the control group in terms of certain demographic and clinical characteristics. The examination of cardiac markers in our study between PE patients and the control group is given in Table 3.

According to Table 3, in ECG findings, normal sinus rhythm was found to be 65.4% in PE patients while 74.4% in the control group, and this difference was not statistically significant (p=0.222). Syncope or tachycardia was 17.9% in the PE group and 6.4% in the control group, and this difference was significant (p=0.028). There were no significant differences between the groups in terms of atrial fibrillation, block, and other ECG findings.

When EF were analyzed, mean EF, was not statistically significant between PE and control groups (p=0.069). The mean of PAP was also similar between the groups (p=0.545). RVV was 29.5% in the PE group and 18.4% in the control group, and this difference was not statistically significant (p=0.108). These data show that ECG and EF of PE patients had some differences compared to the control group, but most of these differences were not statistically significant. The results of the statistical analysis of the utility scores used in PE estimation are given in Table 4.

The results of hemogram, coagulation, and blood gas parameters of the patients in PE and control groups are given in Table 5.

According to Table 5, the coagulation parameter D-dimer and the blood gas parameter PaCO<sub>2</sub> were statistically different

Table 3. The examination of cardiac markers in our study in terms of PE patients and control group

	Group		
	PE (n=78)	Control (n=78)	p value
ECG	Count (%)	Count (%)	
NSR	51 (65.4)	58 (74.4)	0.222
ST	14 (17.9)	5 (6.4)	0.028
AF	7 (9.0)	12 (15.4)	0.221
BLOK	6 (7.7)	2 (2.6)	0.147
S1Q3T3	0 (0)	1 (1.3)	0.316
Other	2 (2.6)	4 (5.1)	0.405
Echocardiography	Mean ± SD	Mean ± SD	
EF	57.37±7.01	55.85±8.13	0.069
PAP	32.98±10.93	34.32±12.55	0.545
	Count (%)	Count (%)	
RVV	23 (29.5)	14 (18.4)	0.108

ECG: Electrocardiography, AF: Atrial fibrillation, EF: Ejection fraction, NSR: Normal sinus rhythm, PAP: Pulmonary artery pressure, RVV: Right ventricular volume, SD: Standard deviation, ST: Sinus tachycardia, PE: Pulmonary embolism

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between the groups. D-dimer was found to be high in the PE group, while PaCO<sub>2</sub> was found to be low. The descriptive statistics of the biochemical parameters between the groups included in the study are shown in Table 6.

According to Table 6, statistically significant differences were observed in some parameters in the biochemistry analysis between PE patients and the control group. BUN levels were significantly lower in PE patients (26.72±18.63) compared to

Table 4. Examination of useful scores in PE prediction				
	Group			
	PE	PE Control		
	Mean ± SD	Mean ± SD		
Shock index	0.82±0.25	0.62±0.12	<0.001	
Wells score	5.73±4.20	0.66±0.95	<0.001	
Geneva score	eneva score 4.20±2.96 2.11±2.20 <b>&lt;0.001</b>			
PE: Pulmonary embolism, SD: Standard deviation				

Table 5. Comparison of hematologic parameters in PE and control groups

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	Group			
	PE	PE Control		
	Mean ± SD	Mean ± SD	p value	
Hemogram				
HGB	12.41±2.30	12.40±2.64	0.945	
MCV	86.80±9.64	85.89±7.61	0.296	
НСТ	38.48±6.75	38.04±7.88	0.717	
PLT	223.94±124.97	245.46±115.99	0.326	
PDW	11.98±4.44	11.63±2.12	0.420	
Monocyte	0.90±1.17	0.94±1.31	0.838	
RDW	17.0±6.93	17.41±10.18	0.714	
Coagulation				
APTT	31.18±20.68	35.97±24.62	0.089	
PT	19.35 ± 16.69	17.06±12.76	0.481	
INR	1.62±2.30	1.38±0.79	0.490	
Fibrinogen	361.02±164.41	396.51±179.27	0.468	
D-dimer	8.78±7.99	2.49±2.54	<0.01	
Blood gas				
Ph	7.39±0.13	7.38±0.08	0.344	
PaCO <sub>2</sub>	34.79±11.60	39.96±18.28	0.027	
PaO <sub>2</sub>	65.38±36.31	55.71±24.63	0.162	
HCO <sub>3</sub>	22.99±9.68	23.08±8.94	0.916	
Lactate	2.70±2.72	2.82±3.65	0.589	

APTT: Activated partial thromboplastin time test, HCO<sub>3</sub>: Bicarbonate, HCT: Hematocrit, INR: International normalized prothrombin time, MCV: Mean corpuscular erythrocyte volume, PE: Pulmonary embolism, PLT: Platelet, PDW: Platelet distribution width, RDW: Erythrocyte distribution width, PaCO<sub>2</sub>: Partial carbon dioxide pressure, PaO<sub>2</sub>: Partial oxygen pressure, Ph: Potential hydrogen, PT: Prothrombin time, SD: Standard deviation, HGB: Hemoglobin

the control group (32.40 $\pm$ 22.70, p=0.020). CK level was higher in the PE group (194.77 $\pm$ 650.76) compared to the control group (120.02 $\pm$ 151.06, p=0.038). Total protein level was lower in PE patients (6.23 $\pm$ 1.11) than in the control group (6.60 $\pm$ 0.83, p=0.023). Similarly, albumin level was lower in PE patients (3.39 $\pm$ 0.64) compared to the control group (3.63 $\pm$ 0.54, p=0.013). CRP level was higher in the PE group (7.51 $\pm$ 6.94) compared than the control group (6.02 $\pm$ 7.92, p=0.015). In addition, triglyceride levels were significantly higher in PE patients (136.74 $\pm$ 91.60) compared to the control group (112.69 $\pm$ 59.70, p=0.020). These findings reveal that there are significant differences in the biochemical profiles of PE patients compared to the control group. The comparison of coagulation factors in PE and the control group is given in Table 7.

On the other hand, in this study, the effect of PE on 3-month mortality was examined, revealing 34 (43.6%) patients in the PE group and 19 (24.4%) patients in the control group died within 3 months. 3-month mortality in the PE group was significantly higher than in the control group (p<0.05).

Table 6. The comparison of biochemical parameters in PE and control groups

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	Group		
	PE	Control	p value
	Mean ± SD	Mean ± SD	
Biochemistry			
Creatinine	1.18±0.58	1.52±1.63	0.234
BUN	26.72±18.63	32.40±22.70	0.020
LDH	531.89±1320.29	412.57±357.75	0.207
CK	194.77±650.76	120.02±151.06	0.038
CK-MB	37.48±73.75	31.68±35.51	0.828
Total protein	6.23±1.11	6.60±0.83	0.023
Albumin	3.39±0.64	3.63±0.54	0.013
ALT	50.03±164.60	67.16±191.92	0.766
AST	64.34±192.95	69.64±149.98	0.578
Pro-BNP	4102.11±6775.67	6472.31±10494.64	0.983
Procalcitonin	1.10±3.90	3.43±14.85	0.486
CRP	7.51±6.94	6.02±7.92	0.015
Troponin	109.54±376.94	207.31±856.01	0.632
Triglycerides	136.74±91.60	112.69±59.70	0.020
Total cholesterol	175.25±52.31	160.27±42.33	0.135
LDL-cholesterol	107.65±38.86	97.12±29.52	0.110
HDL-cholesterol	38.93±11.58	39.73±16.24	0.944

ALT: Alanine transaminase, AST: Aspartate transaminase, CK: Creatinine kinase, CK-MB: Creatinine kinase myocardial band, LDH: Lactate dehydrogenase, LDL-cholesterol: Low-density lipoprotein, HDL-cholesterol: High-density lipoprotein cholesterol, Pro-BNP: ProBrain natriuretic peptide, PE: Pulmonary embolism, SD: Standard deviation, BUN: Blood urea nitrogen, CRP: C-reactive protein

#### Al Modeling

To classify PE, a MLP artificial neural network model was created in which the variables that were statistically different between PE and the control group were used as the independent variables. Gradient descent was used as the optimization function for the model. The performance metrics of the classification model are given in Table 8.

Considering the performance metrics obtained in Table 8, the MLP classification model is quite successful in classifying PE and control groups. Figure 1 shows the graph of the variable importance values obtained from the MLP model, where the most important variables in classifying PE are displayed in order. As a result of the MLP model created according to Figure 1, Wells score and D-dimer were found to be the two important variables in predicting PE.

Table 7. The comparison of coagulation factors in PE and control group					
	Group				
	PE	Control	p value		
	Mean ± SD	Mean ± SD			
Fibrinopeptide A	28.26±27.16	20.29±15.92	0.129		
Protein S	79.44±89.64	84.74±114.92	0.917		
Coagulation factor 5	98.99±59.62	99.30±74.59	0.996		
Vitamin K	312.67±334.77	237.04±182.20	0.862		
Plasminogen	835.50±1354.78	613.63±357.08	0.337		
PE: Pulmonary embolism, SD: Standard deviation					

#### Discussion

The mean age of the PE group was 68.48±16.4 years; the female rate was 57.7%, and the most common presenting complaint was dyspnea. Palpitation was a significant symptom in the PE group. Diabetes mellitus and DVT were significantly higher in the PE group, whereas hypertension and coronary artery disease were significantly higher in the control group. Among the vital signs, tachycardia and low mean systolic and diastolic blood pressure were significant in PE. We found both Wells and Geneva scores to be significant in the diagnosis of PE. When we compared the scores, the specificity and sensitivity of the Wells score were higher compared to another scoring method. An increasing shock index is significant for the diagnosis of PE. Low PaCO<sub>2</sub> in blood gas was a significant finding in patients with PE. Elevated D-dimer, elevated CRP, triglycerides, and low BUN, total protein, and albumin were significant in the diagnosis of PE. In addition, fibrinopeptide A, factor 5, protein S, vitamin K, and plasminogen from the thrombophilia panel were not significantly associated with PE.

Table 8. Performance metrics and 95% confidence intervals for a model to classify PE				
Performance metrics	Value	95% confidence interval		
Accuracy	0.96	0.88-1.00		
Sensitivity	1.00	0.78-1.00		
Specificity	0.89	0.52-0.99		
F1-score	0.97	0.90-1.00		
MCC	0.91	0.80-1.00		
G-mean	0.97	0.90-1.00		
MCC: Matthews's correlation coefficient, PE: Pulmonary embolism				

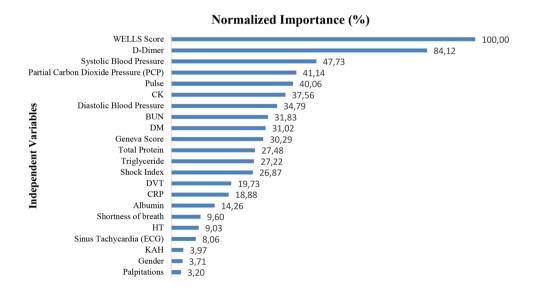


Figure 1. The variable significance based on MLP estimation model

MLP: Multilayer perceptron, CK: Creatine kinase, BUN: Blood urea nitrogen, DM: Diabetes mellitus, DVT: Deep vein thrombosis, CRP: C-reactive protein, HT: Hypertension, KAH: Coronary artery disease

When the mean age of PE patients was analyzed, it was found that Wells et al. (9) had a mean age of 50.5±18.4 years, van der Hulle et al. (11) had a mean age of  $53\pm18$  years, Le Gal et al. (15) had a mean age of 60.6±19.4 years, Roy et al. (22) had a mean age of 52±18.5 years, and Penaloza et al. (23) had a mean age of 63.9 years. In the present study, the mean age of the PE group was 68.48±16.4 years, and the mean age of the control group was 70.44±13.4 years. No significant difference was found between them. The female rate was 62.7% in Wells et al (9), 62% in van der Hulle et al. (11), 58.2% in Le Gal et al. (15), 60.8% in Roy et al. (22), and 62% in Penaloza et al. (23). In this study, consistent with the literature, the percentage of females in the PE group was found to be 57.7%, while the percentage of males was 42.3%. Many studies have demonstrated the relationship between tachycardia and PE (24-26). The significantly higher rate of tachycardia in the PE group is compatible with the literature.

PE is most often a complication of DVT. According to the literature, the rate of DVT in patients diagnosed with PE varies between 21% and 37%. Even in patients with suspected PE, if the lower extremity Doppler USG is positive, anticoagulant treatment can be started without the need for further examination (27). In the current study, the detection rate of DVT in the PE group was found to be significantly higher than in the control group. As a result, the relationship between PE and DVT is similar to that described in the literature.

Upon examining studies comparing the diagnostic accuracy of the most commonly used scoring systems (KOTS, Wells, and modified Geneva), it was found that the diagnostic accuracy of the Wells score was higher than that of the modified Geneva and simplified Geneva. In the studies of Shen et al. (17) and Wong et al. (28), the specificity and sensitivity of the Wells score were found to be significantly higher than the modified Geneva score. In the current study, it was found that high Wells and Geneva scores were significant in the diagnosis of PE. In this respect, the study aligns with previous literature in its methodological approach.

Thrombophilia is a inherited risk factor for VTE. Factor V Leiden deficiency and protein C deficiency are two additional common causes. Depending on the characteristics of the population selected in studies, the thrombophilia detection rate is between 10-50% (29). In this study, we measured Factor V Leiden, fibrinopeptide a, protein s, vitamin K, and plasminogen levels in accordance with the thrombophilia panel. In our study, no significant difference was found in these parameters between the PE group and the control group. The reason why thrombophilia is not significant, unlike in the literature, is what we think is due to the high average age of the patient population in the study. Additionally, it would be appropriate to perform a thrombophilia

examination by waiting 3-6 weeks after the diagnosis of PE, but this situation could not be achieved.

D-dimer is the fibrin breakdown product resulting from the destruction of thrombus formed during thrombolytic events (30). With acute PE, D-dimer level increases. Studies have shown that high D-dimer levels have HS, but low specificity in VTE. It has a high negative predictive value as it excludes.

VTE events with >95% sensitivity in ambulatory patients and in patients with low or medium COTS, unless the latter have any comorbidities. While sensitivity with high positive predictive value has low specificity, sensitivity with high negative predictive value has high specificity. it is more meaningful in excluding PE rather than making a diagnosis (31). In a meta-analysis study, in the preliminary diagnosis of PE, the D-dimer test was found to be high in 94% of the patients and normal in 6%. In our study, consistent with the literature, the positive predictive value of D-dimer in the diagnosis of PE was found to be significantly high.

There are publications on the use of BUN levels for predicting mortality in PE-patients. In a study conducted in our country, the relationship between a BUN value of 34.5 mg/dL at the time of diagnosis and mortality in patients diagnosed with acute PE, and treated aggressively with t-PA, was found to be significant with 85% sensitivity and 91% specificity (32). In another study, the ratio of BUN to serum albumin (B/A) was investigated to predict the mortality of patients hospitalized in the intensive care unit with a diagnosis of PE. This study showed that as the B/A ratio increases, the intensive care mortality of PE patients also increases (33). In the study, the BUN value at the time of admission was found to be significantly higher in the control group. Since the current study attempts to diagnose rather than predict mortality, there are no similar studies in the literature. More studies are needed on this subject.

Aujesky et al. (34) investigated the benefits of using CRP in combination with KOTS in diagnosing PE and concluded that while a CRP value >5 mg/dL was significant in excluding PE when combined with low KOTS, CRP alone could not exclude PE. Roumen-Klappe et al. (35), also reported that CRP increased in PE. A study comparing D-dimer and CRP levels in the diagnosis and exclusion of PE found that a standard CRP test using a cut-off level of 5 mg/dL can be used alone or in combination with KOTS to safely exclude PE (36). In the current study, the CRP value was significantly higher in the PE group compared to the control group. Considering similar studies in the literature, we think that elevated CRP, combined with COTS at medium to high risk, may be meaningful. However, we believe that studies with a larger number of patients are needed on this subject.

When the studies investigating the use of AI in the diagnosis of PE were examined, Müller-Peltzer et al.'s (37) study found common false positives originating from soft tissue and pulmonary vein in diagnosing PE with AI. In their study, Li et al. (38), Douillet et al. (39) stated that AI with ML algorithms will be a future tool to guide the physician regarding suspected acute PE. With the modeling obtained, it was observed that PE was classified with very high success and possible risk factors were obtained. According to the variable importance values obtained by modeling, Wells score and D-dimer were identified as the most important risk factors. With the current study, it has been shown that AI can be used in PE prediction, in line with the literature. In terms of better evaluation of the results of our study and the usability of AI in the clinic, we think that further studies with a larger number of patients are needed.

## **Study Limitations**

The first limitation of our study is that the accuracy of the presented AI model was not tested prospectively. The second limitation is that the patients diagnosed with PE or alternative diagnoses were not examined in the survey. The contribution to the survey of the AI model that emerged as a result of the study could not be examined. Another limitation is that it is not known whether the clinicians who collected the qualitative data of the study were trained extensively about PE. Another limitation is that the study was conducted in a single center and with a limited number of patients. Therefore, it is recommended that the study be repeated in a multicenter study with a larger number of patients and conducted by clinicians who have standard knowledge about PE and its exclusion. That the AI model be studied prospectively in diagnosis and that the patients diagnosed be followed up in the survey.

### Conclusion

As a result, in the diagnosis of PE, the importance of evaluating patients' anamnesis, physical examination, laboratory and imaging findings well and using scores has been determined. However, it has been determined that AI can be used before imaging methods are requested in the diagnosis of PE.

#### **Ethics**

**Ethics Committee Approval:** The appropriateness of this study was approved by the İnönü University Scientific Research and Publication Ethics Committee with (decision number: 2022/45, date: 20.04.2022).

**Informed Consent:** In this study, 156 patients admitted to the Department of Emergency Medicine of İnönü University Faculty

of Medicine Turgut Özal Medical Center from 13.10.2022-14.10.2024, with PE symptoms, were prospectively analyzed.

#### **Footnotes**

#### **Authorship Contributions**

Surgical and Medical Practices: M.S., M.G.T., Concept: M.S., M.G.T., H.Y., Design: M.S., M.G.T., Z.K., Ş.Y., Data Collection or Processing: M.S., H.Y., Z.K., Ş.Y., Analysis or Interpretation: M.G.T., H.Y., Z.K., Ş.Y., Literature Search: M.S., M.G.T., Writing: M.S., M.G.T.

**Conflict of Interest:** The authors declare that they have no conflicts of interest.

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#### References

- Turetz M, Sideris AT, Friedman OA, Triphathi N, Horowitz JM. Epidemiology, pathophysiology, and natural history of pulmonary embolism. Semin Interv Radiol. 2018;35:92-8.
- Pruszczyk P, Torbicki A, Kuch-Wocial A, Szulc M, Pacho R. Diagnostic value of transoesophageal echocardiography in suspected haemodynamically significant pulmonary embolism. Heart. 2001;85:628-34.
- 3. Jiménez D, de Miguel-Díez J, Guijarro R, Trujillo-Santos J, Otero R, et al. Trends in the management and outcomes of acute pulmonary embolism: analysis from the RIETE registry. J Am Coll Cardiol. 2016;67:162-70.
- Barco S, Woersching AL, Spyropoulos AC, Piovella F, Mahan CE. European Union-28: an annualised cost-of-illness model for venous thromboembolism. Thromb Haemost. 2016;115:800-8.
- 5. Kadel PB. Postpartum pulmonary embolism and outcome, experience at a tertiary centre. Int J Multidiscip Res Anal. 2021;4:82-5.
- Türkdoğan Tunalı F, Ertekin E, Zencir C, Yazıcı O, Tunçyürek Ö, Çanakçı SE. The role of right ventricular volume in the diagnosis of pulmonary embolism and morbidity prediction. J Surg Med. 2021;5:799-802.
- Hogg K, Brown G, Dunning J, Wright J, Carley S, Foex B, Mackway-Jones K. Diagnosis of pulmonary embolism with CT pulmonary angiography: a systematic review. Emerg Med J. 2006;23:172-8.
- Shonyela FS, Yang S, Liu B, Jiao J. Postoperative acute pulmonary embolism following pulmonary resections. Ann Thorac Cardiovasc Surg. 2015;21:409-17
- Wells PS, Anderson DR, Rodger M, Stiell I, Dreyer JF, Barnes D, et al. Excluding pulmonary embolism at the bedside without diagnostic imaging: management of patients with suspected pulmonary embolism presenting to the emergency department by using a simple clinical model and d-dimer. Ann Intern Med. 2001;135:98-107.
- Wolf SJ, McCubbin TR, Feldhaus KM, Faragher JP, Adcock DM. Prospective validation of wells criteria in the evaluation of patients with suspected pulmonary embolism. Ann Emerg Med. 2004;44:503-10.
- van der Hulle T, Cheung WY, Kooij S, Beenen LFM, van Bemmel T, van Es J, et al. Simplified diagnostic management of suspected pulmonary embolism (the YEARS study): a prospective, multicentre, cohort study. Lancet. 2017;390:289-297. Epub 2017 May 23. Erratum in: Lancet. 2017;390(10091).
- Kline JA, Mitchell AM, Kabrhel C, Richman PB, Courtney DM. Clinical criteria to prevent unnecessary diagnostic testing in emergency department patients with suspected pulmonary embolism. J Thromb Haemost. 2004;2:1247-55.

- Kline JA, Courtney DM, Kabrhel C, Moore CL, Smithline HA, Plewa MC, et al. Prospective multicenter evaluation of the pulmonary embolism rule-out criteria. J Thromb Haemost. 2008;6:772-80.
- Penaloza A, Verschuren F, Meyer G, Quentin-Georget S, Soulie C, Thys F, et al. Comparison of the unstructured clinician gestalt, the wells score, and the revised Geneva score to estimate pretest probability for suspected pulmonary embolism. Ann Emerg Med. 2013;62:117-124.e2.
- Le Gal G, Righini M, Roy PM, Sanchez O, Aujesky D, Bounameaux H, et al. Prediction of pulmonary embolism in the emergency department: the revised Geneva score. Ann Intern Med. 2006;144:165-71.
- Jian LQ, Zhau TK, Misni MN, Wee CS. A single centre annual audit on computed tomography pulmonary angiogram: demographic, clinical scoring system, patients' outcome. Borneo | Med Sci. 2020;14:11-9.
- 17. Shen JH, Chen HL, Chen JR, Xing JL, Gu P, Zhu BF. Comparison of the wells score with the revised Geneva score for assessing suspected pulmonary embolism: a systematic review and meta-analysis. J Thromb Thrombolysis. 2016;41:482-92.
- 18. Jiang F, Jiang Y, Zhi H, Dong Y, Li H, Ma S, et al. Artificial intelligence in healthcare: past, present and future. Stroke Vasc Neurol. 2017;2:230-43.
- Rana M, Bhushan M. Machine learning and deep learning approach for medical image analysis: diagnosis to detection. Multimed Tools Appl. 2022:1-39.
- Ahmad MN, Abdallah SA, Abbasi SA, Abdallah AM. Student perspectives on the integration of artificial intelligence into healthcare services. Digit Health. 2023;9:20552076231174095.
- Tavares J. Application of artificial intelligence in healthcare: the need for more interpretable artificial intelligence. Acta Med Port. 2024;37:411-4.
- Roy PM, Friou E, Germeau B, Douillet D, Kline JA, Righini M, et al. Derivation and Validation of a 4-level clinical pretest probability score for suspected pulmonary embolism to safely decrease imaging testing. JAMA Cardiol. 2021;6:669-77.
- Penaloza A, Verschuren F, Dambrine S, Zech F, Thys F, Roy PM. Performance
  of the pulmonary embolism rule-out criteria (the PERC rule) combined
  with low clinical probability in high prevalence population. Thromb Res.
  2012;129:e189-93.
- Hobohm L, Becattini C, Ebner M, Lerchbaumer MH, Casazza F, Hasenfuß G, et al. Definition of tachycardia for risk stratification of pulmonary embolism. Eur J Intern Med. 2020;82:76-82.
- Keller K, Beule J, Coldewey M, Dippold W, Balzer JO. Heart rate in pulmonary embolism. Intern Emerg Med. 2015;10:663-9.
- Bach AG, Bandzauner R, Nansalmaa B, Schurig N, Meyer HJ, Taute BM, et al. Timing of pulmonary embolism diagnosis in the emergency department. Thromb Res. 2016;137:53-7.

- 27. Fields JM, Davis J, Girson L, Au A, Potts J, Morgan CJ, et al. Transthoracic echocardiography for diagnosing pulmonary embolism: a systematic review and meta-analysis. J Am Soc Echocardiogr. 2017;30:714-723.e4.
- 28. Wong DD, Ramaseshan G, Mendelson RM. Comparison of the Wells and Revised Geneva scores for the diagnosis of pulmonary embolism: an Australian experience. Intern Med J. 2011;41:258-63.
- Monreal M, Campo RD, Papadakis E. Thrombophilia and venous thromboembolism: RIETE experience. Best Pract Res Clin Haematol. 2012;25:285-94.
- Stein PD, Terrin ML, Hales CA, Palevsky HI, Saltzman HA, Thompson BT, et al. Clinical, laboratory, roentgenographic, and electrocardiographic findings in patients with acute pulmonary embolism and no pre-existing cardiac or pulmonary disease. Chest. 1991;100:598-603.
- 31. Stein PD, Hull RD, Patel KC, Olson RE, Ghali WA, Brant R, et al. D-dimer for the exclusion of acute venous thrombosis and pulmonary embolism: a systematic review. Ann Intern Med. 2004;140:589-602.
- 32. Tatlisu MA, Kaya A, Keskin M, Avsar S, Bozbay M, Tatlisu K, et al. The association of blood urea nitrogen levels with mortality in acute pulmonary embolism. J Crit Care. 2017;39:248-53.
- 33. Fang J, Xu B. Blood urea nitrogen to serum albumin ratio independently predicts mortality in critically ill patients with acute pulmonary embolism. Clin Appl Thromb Hemost. 2021;27:10760296211010241.
- 34. Aujesky D, Hayoz D, Yersin B, Perrier A, Barghouth G, Schnyder P, et al. Exclusion of pulmonary embolism using C-reactive protein and D-dimer. A prospective comparison. Thromb Haemost. 2003;90:1198-203.
- 35. Roumen-Klappe EM, den Heijer M, van Uum SH, van der Ven-Jongekrijg J, van der Graaf F, Wollersheim H. Inflammatory response in the acute phase of deep vein thrombosis. J Vasc Surg. 2002;35:701-6.
- 36. Steeghs N, Goekoop RJ, Niessen RW, Jonkers GJ, Dik H, Huisman MV. C-reactive protein and D-dimer with clinical probability score in the exclusion of pulmonary embolism. Br J Haematol. 2005;130:614-9.
- 37. Müller-Peltzer K, Kretzschmar L, Negrão de Figueiredo G, Crispin A, Stahl R, et al. Present limitations of artificial intelligence in the emergency setting performance study of a commercial, computer-aided detection algorithm for pulmonary embolism. Rofo. 2021;193:1436-44.
- 38. Li X, Wang X, Yang X, Lin Y, Huang Z. Preliminary study on artificial intelligence diagnosis of pulmonary embolism based on computer in-depth study. Ann Transl Med. 2021;9:838.
- Douillet D, Roy PM, Penaloza A. Suspected acute pulmonary embolism: gestalt, scoring systems, and artificial intelligence. Semin Respir Crit Care Med. 2021;42:176-82.