scientific reports



OPEN

Machine learning-based prediction of 90-day prognosis and in-hospital mortality in hemorrhagic stroke patients

Ahmad A. Abujaber¹, Ibrahem Albalkhi^{2,3}, Yahia Imam⁴, Said Yaseen⁵, Abdulqadir J. Nashwan¹, Naveed Akhtar⁴ Ibrahim M. Alkhawaldeh⁶

This study aims to predict hemorrhagic stroke outcomes, including 90-day prognosis and in-hospital mortality, using machine learning models and SHapley Additive exPlanations (SHAP) analysis. Data were collected from a national Stroke Registry from January 2014 to July 2022. Various predictive factors were considered, such as stroke severity at presentation, patient demographics, laboratory results, admission location, and other clinical features. Random forest, logistic regression, XGboost, support vector machines, and decision trees were trained and evaluated. SHAP analysis was conducted to identify key predictors. The RF model demonstrated superior performance in predicting prognosis, while LR was more effective in predicting in-hospital mortality. The National Institute of Health Stroke Score (NIHSS) and admission location were key predictors. Despite its limitations, this research underscores the importance of advancing stroke registries and emphasizes the necessity for comprehensive external validation of predictive models. Furthermore, it demonstrates the importance of initial stroke severity in influencing patient outcomes and highlights the significance of admission to stroke units in reducing poor outcomes. This may help shape interventions to enhance stroke center capacities and influence strategic policies. This study contributes towards developing more precise predictive models for hemorrhagic stroke outcomes, potentially impacting clinical practice and optimizing resource allocation significantly.

Keywords Hemorrhagic stroke, Stroke registry, Prognosis, Mortality, Machine learning

Stroke is a major public health issue, impacting 13.7 million people annually and causing 5.5 million deaths¹, as well as significant long-term disability². While ischemic strokes are more frequent³, hemorrhagic strokes are more severe, with in-hospital mortality rates of 20–30% and up to 80% of survivors requiring long-term care⁴.

Various risk factors, including diabetes, COPD, high blood pressure, smoking, diet, and physical activity, are associated with different types of strokes⁴. However, certain risk factors are more closely linked to specific stroke subtypes. Hypertension, for instance, increases the risk of severe outcomes in hemorrhagic strokes compared to ischemic strokes⁵, while diabetes is linked to poorer outcomes in ischemic strokes⁶. A study in Uganda found that individuals recovering from hemorrhagic strokes experienced more severe impairments within 30 days post-discharge compared to those recovering from ischemic strokes, as measured by the modified Rankin scale (mRS)⁷.

Hemorrhagic strokes are more common in males, accounting for 10–20% of annual stroke cases⁸. A study from 2000 to 2019 found that while both sexes with intracerebral hemorrhage (ICH) saw declining favorable outcomes, only females improved in unfavorable outcomes and mortality rates. For subarachnoid hemorrhage (SAH), both genders showed improvements in outcomes and mortality⁹. Beyond physical effects, hemorrhagic strokes also lead to psychological issues like anxiety, depression, fatigue, and sleep disturbances, particularly in patients with higher mRS scores, as shown by research from Sarah Ecker and colleagues¹⁰.

¹Nursing Department, Hamad Medical Corporation, P.O. Box 3050, Doha, Qatar. ²College of Medicine, Alfaisal University, Riyadh, Saudi Arabia. ³Department of Neuroradiology, Great Ormond Street Hospital NHS Foundation Trust, Great Ormond St, London WC1N 3JH, UK. ⁴Neurology Section, Neuroscience Institute, Hamad Medical Corporation, Doha, Qatar. ⁵School of Medicine, Jordan University of Science and Technology, Irbid, Jordan. ⁶Faculty of Medicine, Mutah University, Al-Karak, Jordan. [™]email: anashwan@hamad.ga

Modern healthcare focuses on preventing adverse outcomes by customizing patient care plans. This approach requires significant investment in improving clinicians' ability to predict outcomes, allowing personalized care that enhances patient and family quality of life¹¹. Scholars have worked extensively to predict stroke outcomes, including short- and long-term prognosis, disabilities, and mortality, using variables like patient demographics, health conditions, lab results, stroke severity, and hospital-related factors. For instance, Wang et al. identified age, hemorrhage size, location, and the presence of diabetes and hypertension as key factors in predicting a one-month hemorrhagic stroke prognosis¹². Matsuo and colleagues noted that lifestyle habits, such as smoking, impact recovery, with non-smokers more likely to have favorable outcomes¹³. Research consistently shows that hemorrhagic stroke prognosis varies by location, with strokes in the cerebral lobes generally having better outcomes than those in deeper brain structures like the thalamus¹⁴.

Some researchers have used established tools to predict stroke prognosis, including mortality. Moon and colleagues employed the Acute Physiology and Chronic Health Evaluation II (APACHE II), a tool for predicting mortality in critically ill patients, which effectively predicted both ischemic and hemorrhagic stroke mortality with an AUC of 0.80^{15} . Abdelghany and team used the modified Stroke subtype, Oxfordshire Community Stroke Project Classification, Age, and pre-stroke Rankin score (mSOAR) to predict 7-day in-hospital mortality and post-stroke disability, achieving AUC values between 0.8 and 0.82^{16} .

The use of machine learning in medical research has significantly expanded, particularly in screening, diagnosis, and prognosis. Machine learning algorithms show great promise in predicting stroke outcomes, often matching, or surpassing traditional methods in accuracy for 90-day prognosis and mortality for both ischemic and hemorrhagic strokes. Recent advancements in interpretable machine learning techniques have also improved the accuracy of predicting functional outcomes at discharge. Building on this potential, our study seeks to leverage these models to predict in-hospital mortality and 90-day post-discharge prognosis for hemorrhagic stroke patients, using the mRS as the evaluation metric, thus laying the groundwork for a more detailed methodological approach.

Methods

Study population

We collected data from the Stroke Registry at Hamad General Hospital (HGH) from January 2014 to July 2022. The dataset includes all individuals aged 18 years and older who were admitted to HGH with a primary diagnosis of stroke. Since the establishment of the stroke registry in Qatar until July 2022, a total of 15,859 patients have sought specialized stroke treatment at the hospital. This figure encompasses patients diagnosed with ischemic and hemorrhagic strokes, transient ischemic attacks (TIAs), and stroke mimics. However, our study specifically focuses on patients diagnosed with hemorrhagic strokes, with all other conditions excluded.

Baseline variables

The collected data included a wide range of patient details, covering demographics, hemodynamic measurements upon admission (e.g., heart rate (HR) and blood pressure (BP)), laboratory results, factors that contribute to stroke risk, pre-existing medical conditions, admission locations, and hospitalization outcomes (e.g., length of stay (LOS), and hospital-acquired infections (e.g., Urinary Tract Infection and Pneumonia). Stroke severity at admission was assessed using the National Institute of Health Stroke Score (NIHSS).

In terms of ethnicity, patients were grouped into five specific categories based on their reported nationality: Qatari, Middle East and North Africa (MENA) region, South Asia region, Southeast Asia region (defined according to the United Nations geo-scheme), and all other nationalities were combined into "others" category. It is important to note the distinct classification for Qatari patients, which was implemented to facilitate meaningful comparisons. This categorization considers the unique demographic makeup of the country, where expatriates constitute a sizable portion of the population. This methodology has consistently been used in prior research examining stroke in Qatar. All relevant risk factors, including pre-existing medical conditions and smoking history, were documented during the patient's hospital stay and cross-verified by stroke registry personnel through electronic medical records. Twenty-nine variables were used for predicting the prognosis (mRS-90) and twenty-eight were used to predict the in-hospital mortality. The details are presented in Table 1.

Outcome measures

In this study, we focused on two key outcome variables: the 90-day post-hospital discharge prognosis assessed by the modified Rankin Scale (mRS-90) and in-hospital mortality among patients with hemorrhagic stroke. For the mRS-90, which was collected during the follow-up visit 90 days after discharge, we simplified it into a binary variable to enhance the model's parsimony and to facilitate the interpretation of the output. An mRS score of \leq 2 was categorized as a favorable prognosis, indicating a good prognostic outcome, while an mRS score of > 2 was categorized as an unfavorable prognosis, indicating a poorer outcome. As for in-hospital mortality, patients who passed away during their hospitalization following the stroke incident were considered deceased, while those who were discharged after their admission for hemorrhagic stroke were categorized as survivors.

Inclusion/exclusion criteria

This study encompassed all adult patients aged 18 years or older who received a diagnosis of hemorrhagic stroke. From the initial cohort of 15,859 patients, 1657 patients who were diagnosed with hemorrhagic stroke were 1657, and data on functional outcomes (mRS-90) were available for 1098 patients. Figure 1 summarizes the data inclusion/exclusion procedure.

	90-day modified Ra	nkin Score	In-hospital Mortality						
Variable	Feature Favorable (mRS≤2) Unfavorable (mRS>2)				Feature	Alive	Died	Total	
	< Mean (51.3)	313	321	634	< Mean (50.9)	820	91	911	
Age (years)	≥ Mean (51.3)	202	262	464	≥ Mean (50.9)	678	68	746	
	Mean ± SD (51.3 ± 13)-IQR 16	Mean	Mean ± SD (50.9 ± 13.1)-IQR 17					
	1: Male	447	470	917	1: Male	1248	130	1378	
Sex	2: Female	68	113					279	
	1: Qatari	54	104			199	25	224	
	2: MENA	66	68		-	820 91 678 68 137 1248 130 250 29 199 25 176 22 809 76 266 32 48 4 1337 143 150 16 11 0 1356 136 142 23 06)-IQR 0 830 14 668 145 -IQR 16 130 34 418 41 611 48 246 26 93 10 992 61 476 93 IQR 3.8 762 86 752 72 37.1)-IQR 54 750 87 737 71 24.8)-IQR 32 793 77 676 77 15.9)-IQR 21 776 85 143 7 136 6 241 9 202 52 979 115 519 44 215 61 1283 98 1056 137 442 22 1397 147 101 12 1459 150 39 9 1412 145 86 13	22	198	
Ethnicity	3: South Asian	275	303	578	3: South Asian	809	76	885	
,	4: South-East Asian	98	96	194	4: South-East Asian	266	32	298	
	5: Other	22	12			809 76 266 32 48 4 1337 143 150 16 11 0 1356 136 142 23 26)-IQR 0 830 14 668 145 -IQR 16 130 34 418 41 611 48 246 26 93 10 992 61 476 93 QR 3.8 762 86 752 72 7.1)-IQR 54 750 87 737 71 4.8)-IQR 32 793 77 676 77 5.9)-IQR 21		52	
	1: Ambulance	435	554					1480	
Mode of Arrival	2: Private vehicle	77	25					166	
111000 011111101	3: In-hospital	3	4					11	
	< Mean (0.35)	498	483		•			1492	
Modified Rankin Score (mRS) pre-stroke onset	≥ Mean (0.35)	17	100					165	
Wiodined Rainen Score (Into) pre stroke onset	Mean \pm SD (0.35 \pm 1.1			117	1 1			103	
	< Mean (11.2)	424	174	508				844	
NIHSS at admission	< Mean (11.2) ≥ Mean (11.2)	91	409			820 91 678 68 17 1248 130 250 29 199 25 176 22 809 76 266 32 48 4 1337 143 150 16 11 0 1356 136 142 23 60)-IQR ∪ 836 14 668 145 IQR 16 130 34 418 41 611 48 246 26 93 10 992 61 476 93 QR 3.8 762 86 752 72 7.1)-IQR 54 750 87 737 71 6.8)-IQR ∪ 1.8)-IQR ∪ 2.8 751 77 6.7 759 87 737 71 6.8)-IQR ∪ 1.8)-IQR ∪ 1.8 1.8 1.8 1.8 1.8 1.8 1.8 1.8 1.8 1.8		813	
NIII33 at admission	Mean ± SD (11.2 ± 8.6		409	300			143	013	
	`	1	20	70		820 91 678 68 117 1248 130 250 29 199 25 176 22 809 76 266 32 48 4 1337 143 150 16 11 0 1356 136 142 23 06)-IQR 0 830 14 668 145IQR 16 130 34 418 41 611 48 246 26 93 10 992 61 476 93 IQR 3.8 762 86 752 72 37.1)-IQR 54 750 87 737 71 24.8)-IQR 32 793 77 676 77 15.9)-IQR 21 776 85 143 7 136 6 241 9 202 52 979 115 519 44 215 61 1283 98 1056 137 442 22 1397 147 101 12 1459 150 39 9 1412 145 86 13 1332 153	164		
	1: Underweight	40	30	634			164		
D I M I I (DM)	2: Normal weight	157	157					459	
Body Mass Index (BMI)	3: Overweight	201	251					659	
	4: Obese	92	101					272	
	5: Extremely Obese	25	44		·			103	
	< Mean (8.7)	354	342					1053	
Random Blood Sugar (mmol/l)							93	569	
	Mean \pm SD (8.7 \pm 3.9)	-IQR 3.6			Mean ± SD (8.8 ± 4)-1	QR 3.8	29		
	< Mean (180)	300	275	575	< Mean (180.5)	762	86	848	
Systolic Blood Pressure (SBP) (mmHg)	≥ Mean (180)	212	308	520	≥ Mean (180.5)	752	72	797	
	Mean ± SD (180 ± 36.	5)-IQR 53			Mean ± SD (180.5 ± 3	7.1)-IQI	0 136 23 R 0 14 145 6 34 41 48 26 10 61 93 38 86 72 DR 54 87 71 DR 32 77 77 DR 21 85 7 6 6 9 9 52 115 44 61		
	< Mean (106.5)	286	276	562	< Mean (106.7)	750	87	837	
Diastolic Blood Pressure (DBP) (mmHg)	≥ Mean (106.5)	226	307	533	≥ Mean (106.7)	737	71	808	
	Mean \pm SD (106.5 \pm 2	4)-IQR 33		Mean ± SD (106.7 ± 24.8)-IQR 32					
	< Mean (83)	262	318	580	< Mean (83.35)	793	77	870	
Heart Rate (bpm)	≥ Mean (83)	246	262	508	≥ Mean (83.35)	676	77	753	
	Mean ± SD (83 ± 15.5)-IQR 19			Mean ± SD (83.35 ± 1	5.9)-IQI	R 21		
	≤3 h	225	358	583	≤3 h	776	85	861	
	3-6 h	39	65	104	3-6 h	143	7	150	
Time from onset to hospital arrival (hour)	6-24 h	51	47	98	6-24 h	136	6	142	
	>24 houra	146	37	183	>24 houra	241	9	250	
	Unidentified	54	76	130	Unidentified	820 91 678 68 17 1248 130 250 29 199 25 176 22 809 76 266 32 48 4 1337 143 150 16 11 0 1356 136 142 23 60)-IQR ∪ 830 14 668 145 IQR 16 130 34 418 41 611 48 246 26 93 10 992 61 476 93 QR 3.8 762 86 752 72 7.1)-IQR 54 750 87 737 71 4.8)-IQR 12 1576 85 143 7 15.9)-IQR 12 15776 85 143 7 15.9)-IQR 21 776 85 143 7 159 150 159 14 215 61 1283 98 1056 137 442 22 1397 147 101 12 1459 150 39 9 1412 145 86 13 1332 153	254		
	0: No	355	354	709	0: No	979	20 91 78 68 79 248 130 50 29 99 25 76 22 09 76 66 32 8 4 337 143 50 16 1 0 356 136 42 23	1094	
Diabetes Mellitus (DM)	1: Yes	160	229	389	1: Yes	519	44	563	
	0: No	79	71	150	0: No	215	61	276	
Hypertension (HTN)	1: Yes	436	512	948	1: Yes	1283	98	1381	
	0: No	346	406					1193	
Dyslipidemia	1: Yes	169	177					464	
	0: No	486	528					1544	
Prior stroke	1: Yes	29	55					113	
	0: No	508	560					1509	
Atrial Fibrillation (AF)	1: Yes	7	23					48	
	0: No	490	542					1558	
Coronary Artery Disease (CAD)	1: Yes	25	41					99	
	0: No		536	974	0: No		91 68 130 29 25 22 76 32 4 143 16 0 136 23 .0 14 145 34 41 48 26 10 61 93 86 72 R 54 87 71 R 21 R 32 77 77 R 21 85 7 6 9 52 115 44 61 98 137 22 147 12 150 9 145 13 153	1485	
Tobacco use	0: No 1: Yes	438 77	47	124	0: No 1: Yes			172	
			I T/		1. 103	1 100	U	1/4	

	90-day modified Ra	nkin Score	In-hospital Mortality					
Variable	Feature	Favorable (mRS≤2)	Unfavorable (mRS>2)	Total	Feature	Alive	Died	Total
Various on anti-platelete mes etuales	0: No	468	489	957	0: No	1312	137	1449
Known on anti-platelets pre-stroke	1: Yes	47	94	141	1: Yes	186	22	208
Known on anti-coagulants pre-stroke	0: No	506	557	1063	0: No	1457	151	1608
Known on anti-coagulains pre-stroke	1: Yes	9	26	35	1: Yes	41	8	49
Vnovn on stating pro stroke	0: No	476	483	959	0: No	1319	140	1459
Known on statins pre-stroke	1: Yes	39	100	139	1: Yes	179	19	198
	< Mean (260.7)	258	316	574	< Mean (258.3)	768	137 22 151 8 140 19 77 67 91 85 65	845
Platelets count on admission	≥ Mean (260.7)	239	254	493	≥ Mean (258.3)	673	67	740
	Mean ± SD (260.7 ± 7	8.6)-IQR 89	Mean ± SD (258.3 ± 80)-IQR 91					
	< Mean (10.9)	312	329	641	< Mean (11)	1312 137 14 186 22 20 1457 151 16 41 8 49 1319 140 14 179 19 19 768 77 84 673 67 74 879 85 96 533 65 59 1)-IQR 2.2 953 103 10 453 46 49 15.3)-IQR 4.4 719 5 72 165 42 20 1273 126 13 225 33 25 1344 152 14 154 7 16	964	
Prothrombin Time on admission (seconds)	≥ Mean (10.9)	Tean (10.9) 172 239 411 ≥ Mean (1		≥ Mean (11)	533	65	598	
	Mean ± SD (10.9 ± 6.4	1)-IQR 2.2	Mean ± SD (11 ± 6.4)-IQR 2.2					
	< Mean (27.6)	306	379	685	< Mean (27.8)	953	12 137 6 22 57 151 8 19 140 9 19 8 77 3 67 QR 91 9 85 3 65 R 2.2 33 103 3 46 1QR 4.4 112 5 42 77 126 5 33 44 152 4 7	1058
Activated Partial Thromboplastin Time (seconds)	≥ Mean (27.6)	175	185	360	≥ Mean (27.8)	1312 137 1 186 22 2 1457 151 1 41 8 4 1319 140 1 179 19 1 768 77 8 673 67 7 80)-IQR 91 879 85 9 533 65 5 -IQR 2.2 953 103 1 453 46 4 5.3)-IQR 4.4 719 5 7 614 112 7 165 42 2 1273 126 1 225 33 2 1344 152 1 154 7	499	
	Mean ± SD (27.6 ± 13	.3)-IQR 4.4	Mean ± SD (27.8 ± 15.3)-IQR 4.4					
	1: Stroke Unit	341	191	532	1: Stroke Unit	719	5	724
Admission location	2: ICU	112	342	454	2: ICU	Alive Diec 1312 137 186 22 1457 151 41 8 1319 140 179 19 19 19 19 19 19 1	112	726
	3: Other	62	50	112	3: Other	165	137 22 151 8 140 19 77 67 91 85 65 2 103 46 3.44 5 112 42 126 33 152 7	207
Hospital acquired Pneumonia	0: No	497	423	920	0: No	1273	126	1399
Hospital acquired Fliedillollia	1: Yes	18	160	178	1: Yes	225	33	258
Hospital acquired Urinary Tract Infection (UTI)	0: No	503	478	981	0: No	1312 137 1 186 22 2 1457 151 1 41 8 4 1319 140 1 179 19 1 768 77 8 673 67 7 30)-IQR 91 879 85 9 533 65 5 -IQR 2.2 953 103 1 453 46 4 5.3)-IQR 4.4 719 5 7 614 112 7 165 42 2 1273 126 1 225 33 2 1344 152 1 154 7 1	1496	
Hospital acquired Ormary Tract Infection (OTI)	1: Yes	12	105	117	1: Yes	154	137 22 151 8 140 19 77 67 91 85 65 2 103 46 4.4 5 112 42 126 33 152 7	161
	< Mean (11.2)	437	276	713				
Length of stay (LOS) (days)	≥ Mean (11.2)	Mean (11.2) 78 307 385		385				
	Mean ± SD (11.2 ± 10	.2)-IQR 11.2					137 22 151 8 140 19 77 67 91 85 65 2 103 46 4.4 5 112 42 126 33 152 7	
90-day mRS	0: Favorable (≤ 2) 1: Unfavorable (> 2) 515 583 109							
In-hospital Mortality					0: Alive 2: Died in-hospital	1498	159	1657

Table 1. Statistical characteristics of the collected stroke dataset.

Handling missing data and class imbalance

Multiple Imputations using the Chained Equations (MICE) technique to generate data imputations³⁰ were employed in cases of missing data. The missing data included activated partial thromboplastin time (APTT) at 6.2%, prothrombin time (PT) at 5.7%; platelet counts at 4.3%, heart rate at 2.1%, blood sugar levels at 2.1%, and blood pressure at 0.7%. The cohort exhibited a mortality rate of 9.6%, resulting in concerns about class imbalance. Class weighting to mitigate the imbalance was utilized. Specifically, class weights inversely proportional to class frequencies were assigned, giving more weight to the minority class (patients who died) to enhance their influence during training. It is important to note that the mRS-90 (favorable vs. unfavorable) had an even distribution of classes.

Model training and evaluation

The dataset was divided into a training set (80%) and a validation set (20%) using stratified random sampling. We constructed and fine-tuned our models using the training data and then assessed their performance using the validation data. We trained five different machine learning models, including extreme gradient boosting (XGBoost), random forest (RF), support vector machine (SVM), logistic regression (LR), and decision trees (DT).

To gauge the effectiveness of these models, we employed multiple classification metrics, which included accuracy, precision, specificity, recall, F1-score, area under the receiver operating characteristic curve (AUC), Matthew's correlation coefficient (MCC), log loss, and Brier score. These metrics offer insights into how well the models can accurately classify positive and negative instances while considering the class imbalance. The model with the highest F1-score will be selected as the primary model for subsequent external and temporal validation.

We also utilized the SHAP (SHapley Additive exPlanations) library, a powerful tool for interpreting machine learning models' predictions. This tool generates individual-level feature importance scores known as SHAP values, which quantify the contribution of each feature to a specific prediction outcome.

Ethics and inclusion statement

This study adhered to ethical standards by including local researchers throughout all phases of the research process, including study design, implementation, data ownership, intellectual property, and authorship of

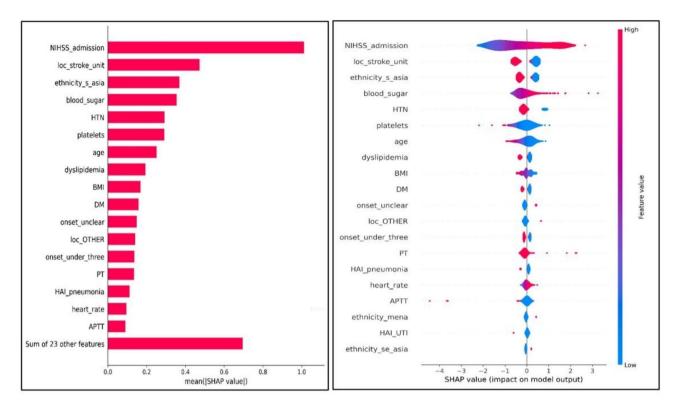


Fig. 1. Data inclusion and exclusion procedure.

publications. The involvement of local partners ensured that the research remained locally relevant and addressed context-specific concerns. Roles and responsibilities among all collaborators were agreed upon at the outset of the project, and capacity-building initiatives for local researchers were discussed and incorporated into the research framework. The research adhered to all applicable local regulations and would not have been restricted or prohibited in the setting of the researchers. No specific exceptions were required for this study, as it was in complete agreement with local stakeholders. The research received approval from the Institutional Research Board (IRB) at Hamad Medical Corporation, Qatar, with reference number MRC-01-22-594, in compliance with ethical guidelines. No aspect of the research posed a risk of stigmatization, incrimination, or discrimination for the participants. Provisions were made to ensure the safety and well-being of all participants. Additionally, risk management plans were implemented to protect the researchers' health, safety, and security. No biological materials, cultural artifacts, or associated traditional knowledge were transferred out of the country. Finally, local and regional research relevant to the study was duly considered and appropriately cited to reflect the contributions of the regional academic community.

Results Baseline characteristics

Table 1 presents the study population's characteristics for both outcome measures. Obviously, 53% of the patients presented with an unfavorable mRS of 90 days post-discharge, while 9.5% had in-hospital mortality.

Model evaluation

Five distinct models were trained for separate predictions. Overall, the machine learning models performed better in predicting functional outcomes, as measured by the mRS-90, compared to its performance in predicting mortality. Specifically, the Random Forest (RF) model demonstrated an impressive F1-Score of 0.815, indicating a strong balance between precision and recall. It also achieved a very good AUC of 0.871 and a low log loss of 0.140, underscoring its high predictive accuracy.

Conversely, when predicting mortality, the Logistic Regression (LR) model showed reasonable accuracy and discrimination power (AUC) of 0.819 and 0.859, respectively. However, it had a moderate F1-score of 0.44, indicating suboptimal precision and recall. This suggests that the inclusion of crucial predictive variables may need improvement.

In summary, based on the various evaluation metrics, we selected the Random Forest (RF) model for functional outcome prediction (mRS-90) and the Logistic Regression (LR) model for predicting in-hospital mortality as the primary choices for further SHAP analysis. Table 2 detail the models' performance metrics. Figures 2 and 3 illustrate the discrimination power for the two selected models.

Model	Accuracy	Precision	Specificity	Recall	F1-Score	AUC	MCC	Balanced Accuracy	Log Loss	Cohen's Kappa	Gini Coefficient	Brier Score
A: mode	A: models performance for predicting functional outcome (mRS-90)											
RF	0.791	0.795	0.737	0.835	0.815	0.871	0.576	0.786	0.440	0.575	0.742	0.140
XGB	0.777	0.777	0.707	0.835	0.805	0.850	0.548	0.771	0.605	0.546	0.699	0.173
SVM	0.782	0.835	0.818	0.752	0.791	0.882	0.567	0.785	0.443	0.564	0.764	0.141
DT	0.764	0.776	0.717	0.802	0.789	0.759	0.521	0.759	8.519	0.521	0.519	0.236
LR	0.759	0.815	0.798	0.727	0.769	0.854	0.523	0.763	0.492	0.519	0.708	0.157
B: mode	B: models performance for predicting in-hospital mortality											
LR	0.819	0.304	0.818	0.828	0.444	0.859	0.428	0.823	0.415	0.363	0.717	0.133
SVM	0.807	0.284	0.809	0.793	0.418	0.848	0.396	0.801	0.228	0.332	0.695	0.067
XGB	0.904	0.421	0.964	0.276	0.333	0.859	0.291	0.620	0.311	0.284	0.718	0.080
DT	0.861	0.185	0.927	0.172	0.179	0.550	0.103	0.550	4.994	0.103	0.100	0.139
RF	0.910	0.333	0.993	0.034	0.063	0.893	0.083	0.514	0.207	0.047	0.786	0.063

Table 2. Models performance for predicting functional outcome (mRS-90) and predicting in-hospital mortality. RF: Random Forest, XGB: XGBoost, SVM: Support Vector Machines, DT: Decision Tree, LR: Logistic Regression

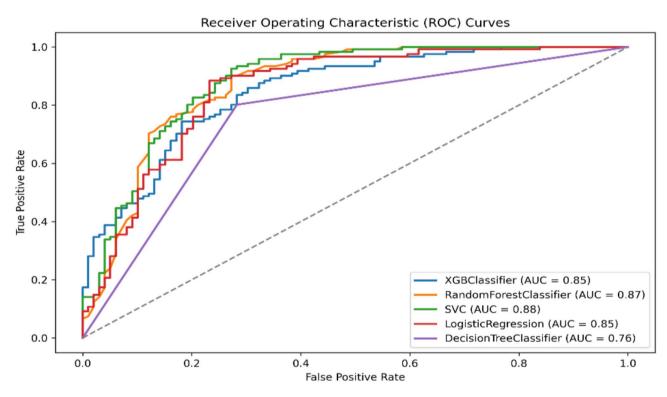


Fig. 2. Area Under the Curve (AUC) for functional outcome (mRS-90)- All models.

SHAP analysis

The SHAP analysis yielded invaluable insights into our predictive models for both mRS-90 and mortality. It revealed that, in descending order of importance for predicting mRS-90, the most influential factors were admission NIHSS, length of stay, admission location, age, and hospital-acquired pneumonia. In contrast, the primary predictors for mortality included NIHSS, admission location, ethnicity, blood sugar, HTN, and platelet count. For visual representations, refer to Figs. 4 and 5.

Discussion

This study examined the effectiveness of five machine learning models and harnessed SHAP analysis to uncover the primary predictors for hemorrhagic stroke outcomes, specifically in-hospital mortality and the 90-day post-hospital discharge prognosis measured by mRS-90. The results underlined the superior performance of the RF model in predicting prognosis and the LR model in predicting in-hospital mortality, surpassing other models in accuracy. SHAP analysis on both models illuminated the pivotal variables that strengthened their predictive capabilities.

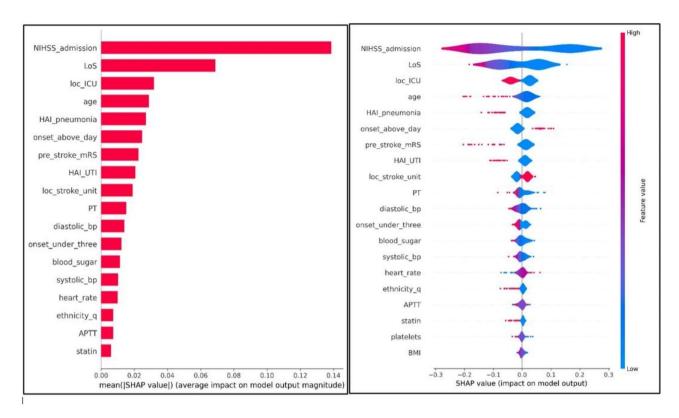


Fig. 3. Area Under the Curve (AUC) for In-hospital mortality - All models.

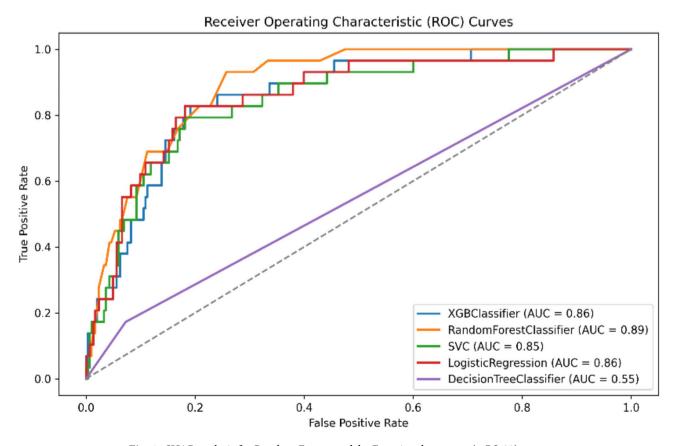


Fig. 4. SHAP analysis for Random Forest model—Functional outcome (mRS-90).

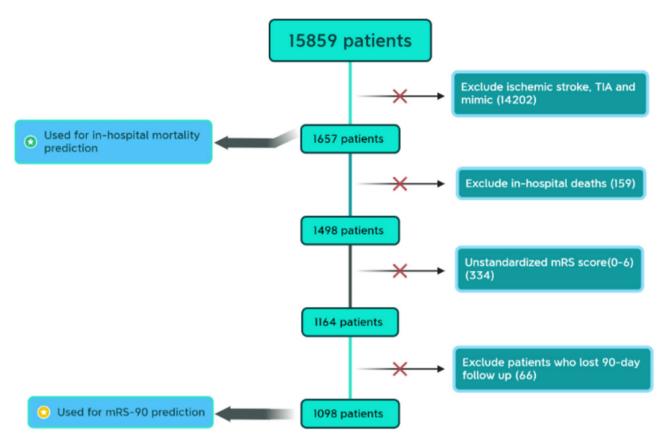


Fig. 5. SHAPA analysis for Logistic Regression model—in-hospital mortality.

For predicting mRS-90, the influential variables included stroke severity (NIHSS upon admission), length of hospital stay (LOS), admission location, age, and hospital-acquired pneumonia. Conversely, for predicting in-hospital mortality in hemorrhagic stroke patients, the crucial factors encompassed NIHSS upon admission, admission location, patient ethnicity, blood sugar levels upon admission (RBS), hypertension (HTN), and admission platelet count. Remarkably, other variables exhibited lower SHAP values, indicating minimal contribution to the model's predictive outcomes. Notably, except for LOS and hospital-acquired pneumonia, all these influential factors can be assessed early upon admission, making them valuable early indicators of prognosis. Consequently, this research can improve clinicians' ability to promptly predict prognosis and mortality and customize care strategies to address mortality risk. Additionally, it can assist families in forming realistic expectations regarding anticipated outcomes, enhancing their involvement in the care delivery plan¹⁷.

Predictors of hemorrhagic stroke prognosis (mRS-90)

Stroke severity, measured by the NIHSS, was identified as the most significant predictor of stroke prognosis, including mortality and functional disability across all timeframes $^{18-20}$. In this study, the mean NIHSS score was 11.2 ± 8.6 . Patients with unfavorable mRS-90 outcomes had significantly higher NIHSS scores than those with favorable outcomes (15.6 vs. 6.2, p < 0.05). This highlights the NIHSS's critical role in early stroke outcome prediction and its importance in guiding personalized preventive care plans 21 .

The LOS plays a role in predicting stroke outcomes. Our study demonstrated that LOS significantly enhances the model's predictive performance (Fig. 4). Generally, a longer LOS is often associated with in-hospital complications, such as hospital-acquired infections, which can potentially result in poor prognostic outcomes and may lead to mortality²². The mean LOS in our study was 11.2 ± 10.2 days. Approximately 80% of patients with LOS exceeding the mean value experienced unfavorable mRS-90 outcomes, whereas only 39% of those with a shorter LOS did so (p-value <0.05). Additionally, the mean LOS for patients in the unfavorable outcome group was significantly longer than those with favorable mRS-90 outcomes (15 vs. 6.9 days, p-value <0.05). It is important to note that LOS can be influenced by various factors, including stroke severity and in-hospital complications^{23,24}. A secondary analysis revealed no statistically significant correlation between LOS and stroke severity (NIHSS). Still, it did find a significant association between LOS and the development of hospital-acquired infections, such as pneumonia (p-value <0.05).

The choice of admission location significantly impacted stroke prognosis, aligning with previous studies suggesting that management in specialized stroke units can lead to reduced mortality, reduced length of stay, and improved patient outcomes^{25,26}. Specifically, this research revealed that patients admitted to critical care units exhibited a higher incidence of unfavorable mRS-90 outcomes than those admitted to the stroke units or other units (such as general wards) (75.3% vs. 36% and 44.6%, respectively). This discrepancy can be attributed

to the initial severity of stroke at admission, as measured by NIHSS, which necessitates closer monitoring and specialized care in critical care settings. Notably, patients admitted to critical care units had significantly higher mean NIHSS scores than those admitted to the stroke units or other units (16.7 vs. 7.3 and 7.2, respectively, *p*-value < 0.05).

Interestingly, although the mean NIHSS scores for patients admitted to stroke units and other units were statistically similar (7.3 vs. 7.2, *p*-value > 0.05), the unfavorable mRS-90 outcomes were notably higher for patients admitted to other units (44.6% vs. 36%, *p*-value < 0.05). This observation corroborates existing research indicating that stroke unit management improves outcomes, including reduced mortality and decreased dependence²⁵. Evans and colleagues correlated that to the superior quality of specialized care delivered in the stroke units compared to the general wards, which includes superior monitoring and better management for fever, oxygenation, nutrition, and aspiration prevention²⁵. Among patients admitted to the general wards, 5.4% developed hospital-acquired pneumonia, and 7.1% developed UTI compared to 4.3% and 4.5%, respectively, for those admitted to stroke units, corroborating the previous literature discussion about improved care outcomes.

This finding shed light on the need to address the capacity challenges in stroke units. When these specialized units cannot accommodate patients, individuals are often placed in general wards, which may provide a different level of care. Consequently, the study advocates exploring expanding stroke unit capacity to enhance patient outcomes, accelerate their recovery and return to the community, and ultimately improve their quality of life.

As observed in previous research, there is a well-documented association between age and an increased likelihood of unfavorable stroke outcomes ¹⁹. Older individuals are known to be more susceptible to developing comorbidities that contribute to adverse outcomes and are linked to poorer disease prognosis²⁷. Our study's mean age was 51.3 ± 13 years, which could be considered relatively lower when assessed in regional and global scales²⁸. Our supplementary analysis revealed that 56.5% of patients older than the mean age experienced unfavorable mRS-90 outcomes, compared to 50.6% of others (p-value < 0.05). Notably, the mean age of patients with unfavorable mRS-90 outcomes was significantly higher than those with favorable mRS-90 outcomes (51.12 vs. 48.8) with a p-value < 0.05. This finding emphasizes the importance of clinicians considering personalized preventive measures for older patients presenting with hemorrhagic stroke.

Additionally, the occurrence of hospital-acquired pneumonia emerged as a robust predictor of stroke patient prognosis. Stroke patients are known to be particularly vulnerable to infections during their hospital stay, which can significantly worsen their functional outcomes²⁹. The study findings indicated that approximately 90% of patients who contracted hospital-acquired pneumonia experienced unfavorable mRS-90 outcomes, compared to 46% of those who did not develop hospital-acquired pneumonia (*p*-value < 0.05). This stresses the importance for clinicians to pay heightened attention to implementing evidence-based preventive measures, such as regular oral care, postural adjustments to prevent aspiration, dysphagia screening, and other strategies outlined by Grossmann and colleagues³⁰.

Predictors of the hemorrhagic stroke in-hospital mortality

Mortality is inherently regarded as an unfavorable outcome in the context of mRS. Consequently, it is logical to anticipate that factors contributing to a poor prognosis, as assessed by mRS-90, may overlap with those influencing mortality, such as NIHSS, in this study. The logistic regression (LR) model pinpointed stroke severity (NIHSS), admission location, patient ethnicity, blood sugar levels at admission (RBS), history of hypertension (HTN), and platelet count at admission as the primary predictors for in-hospital mortality in cases of hemorrhagic stroke."

Like its significance in predicting mRS-90 outcomes, NIHSS emerges as a pivotal predictor of mortality. In our study, the mean NIHSS score stood at 12 ± 8.8 . Notably, patients who deceased post hemorrhagic stroke exhibited a significantly higher mean NIHSS score compared to those who survived (21.3 vs. 11, p-value < 0.05). This sheds light on the importance of the severity of presentation following a hemorrhagic stroke event as an early indicator of adverse outcomes, prompting clinicians to consider early personalized preventive interventions³¹.

The location of admission significantly impacts stroke prognosis and mortality rates. Turner et al. found that admission to a stroke unit is associated with lower mortality and a higher likelihood of being discharged home within a year 26 . Patients in critical care units had a hospital mortality rate of 15%, while those in stroke units had rates above 0.7%, and other units saw a rate of 20.3% (p-value < 0.05). Despite lower severity scores (NIHSS 9.9 vs. 17.1), patients in general wards had the highest mortality, suggesting critical care may offer more appropriate treatment. The NIHSS score for other units was higher than in stroke units (9.6 vs. 5.5, p-value < 0.05), indicating a need to expand stroke unit capacity. Literature supports that specialized stroke units improve outcomes, though patients in critical care generally face poorer prognoses 32 .

Consistent with prior research findings, this study reaffirms the role of ethnicity in influencing stroke outcomes. Notably, our study reveals that patients from South Asian ethnicity exhibit a lower in-hospital mortality rate compared to those from MENA and other South-Asian ethnicities (8.6% vs. 11.2%, p-value < 0.05). Conversely, Qatari patients exhibit the highest mortality rate at 11.2%. Importantly, it is worth highlighting that there is no significant difference in NIHSS scores among the five ethnic groups. However, it is noteworthy that Qatari patients have the highest mean age, 65 years, compared to 59, 47, 47, and 50 years for MENA, South Asian, South-East Asian, and other ethnicities, respectively. In a broader context, this phenomenon can be understood by considering the demographic composition of Qatar. The Qatari population mirrors global age distribution patterns, reflecting a typical society. However, the situation changes when we shift our focus to expatriates. Among expatriates, there is a noticeable difference in age distribution. Most expatriates are younger and of working age, primarily residing in Qatar for employment purposes. This may explain why the mean age of Qatari patients is 64.9 ± 15 years compared to 49 ± 11 years for non-Qatari patients.

The fourth element that boosted the model's predictive precision is the initial Random Blood Sugar (RBS) level. This study aligns with earlier research by establishing a connection between elevated RBS levels upon admission and an adverse prognosis^{33,34}. It is reported that acute hyperglycemia increases brain damage, enlarges

infarct sizes, and reduces cerebral blood volume. It also worsens stroke effects by intensifying reperfusion injury, oxidative stress, and inflammation. Further, acute hyperglycemia increases the risk of platelet aggregation, complicating recovery 35,36 . In our study, the mean RBS level measured 8.8 ± 4 mmol/l, and notably, the mean RBS level among patients who did not survive their hospital stay was significantly higher than that of the group that did survive (10.6 vs. 8.6, with a *p*-value < 0.05).

This study revealed a paradox where patients with a known history of HTN had significantly lower in-hospital mortality rates following hemorrhagic stroke compared to those without a prior HTN diagnosis (7.1% vs. 22.1%, p < 0.05), contradicting previous literature³⁷. Patients with HTN also had lower NIHSS scores, which might be due to regular antihypertensive treatment reducing stroke severity. This could be linked to the idea that systolic blood pressure variability increases stroke mortality risk^{38,39}, with antihypertensive medications reducing these fluctuations. The study's limitations, including missing data on the hemorrhage site and size, suggest further research to understand these findings fully.

Previous studies have established a notable inverse relationship between hemorrhagic stroke mortality and mean platelet count 40,41 . In accordance with this existing literature, our study underscores the potential of a lower platelet count as an early indicator of hemorrhagic stroke mortality. Our secondary analysis unveiled that the mean platelet counts among patients who did not survive was significantly lower than that of the group that did survive (248.6 vs. 259.3, p-value < 0.05).

This study has revealed an intriguing finding: the severity of the initial presentation in cases of hemorrhagic stroke is a paramount predictor for prognosis and can potentially function as an early indicator. Among all the factors considered, the NIHSS exhibited the highest mean SHAP value (refer to Figs. 4 and 5). Importantly, NIHSS and admission location emerged as shared predictors for hemorrhagic stroke outcomes, demonstrating their potential as early indicators to guide clinicians in devising preventive care plans.

As we look to the future, it becomes evident that thorough validation of the study's prediction models is essential to demonstrate their utility in clinical decision-making. Despite the inherent limitations, this research holds promise by shedding light on critical areas for improvement. Notably, it underlines the importance of enhancing the stroke registry by incorporating vital variables such as hemorrhage size, location, and imaging data. Furthermore, this study may pave the way for the development of localized predictive models, facilitating early prognostication, which are crucial elements in creating personalized care plans and supporting the endeavors of precision medicine.

Limitations

This study presents several notable limitations that warrant acknowledgment. Firstly, it relies on retrospective data from a solitary medical center, potentially introducing bias to selection and constraining the generalizability of findings to broader populations. Furthermore, the absence of crucial variables in the stroke registry, including hemorrhage size, location, imaging data, and socioeconomic factors, limits the depth of analysis and overlooks potential influential factors in stroke outcomes.

Utilizing historical data may only partially encompass evolving treatment strategies and shifting patient demographics, potentially affecting the model's applicability in contemporary clinical contexts. Moreover, the prediction window for in-hospital mortality and the 90-day functional status measured by mRS may need to adequately capture longer-term outcomes, potentially limiting its utility in assessing extended prognostic trends.

Also, the dataset exhibits class imbalance, with a pronounced underrepresentation of mortality cases. This imbalance introduces complexity during model development and evaluation, potentially leading to biased model performance despite mitigation efforts. Consequently, it affects result clarity and raises concerns about the machine learning model's utility in guiding clinical decision-making. While machine learning techniques were applied to predict hemorrhagic stroke outcomes, these models necessitate comprehensive external validation to confirm their robustness and applicability.

However, although the models exhibit promise, demonstrating moderate precision and recall rates, especially in predicting in-hospital mortality, there remains room for improvement. Utilizing machine learning models introduces inherent intricacies, potentially complicating their interpretation within clinical settings and integration into practical healthcare workflows.

Lastly, while SHAP analysis identifies critical predictors, this study does not establish definitive causal relationships between these variables and stroke mortality. Thus, while this research advances our understanding of stroke mortality prediction, it is essential to consider its implications within the context of these limitations. Despite these constraints, this study provides valuable insights into predicting hemorrhagic stroke outcomes. It underscores the importance of enhancing data collection within stroke registries to enhance predictive model accuracy and relevance. Further research using more extensive and diverse datasets is imperative to ensure the robustness and refinement of the presented predictive models.

Conclusion

This study explores the prediction of hemorrhagic stroke outcomes using machine learning models and SHAP analysis to identify critical factors. Notably, the severity of hemorrhagic stroke at presentation is a crucial predictor of outcomes, including mortality. The study underscores the significant role of admission location in shaping stroke outcomes, emphasizing the impact of specialized and critical care units in providing superior care. This highlights the need for policy discussions on expanding stroke unit capacity to improve outcomes and reduce the economic burden of long-term disability. However, the study faces limitations, including its retrospective nature, single-center data reliance, and missing variables in the stroke registry, which introduces selection bias and limit generalizability. Additionally, the class imbalance in the dataset poses challenges in model development. Nevertheless, this research underlines improving data collection in stroke registries. It offers promises for more accurate predictive models for hemorrhagic stroke outcomes, mainly since it encompasses the entire population

of patients seeking specialized care. Future research should prioritize larger, more diverse datasets and a deeper exploration of variables to improve clinical practice. Additionally, adopting a multi-centric approach in future studies will enhance generalizability and contribute to advancing personalized stroke care.

Data availability

The datasets generated and/or analyzed during the current study are available from the corresponding author on reasonable request and subject to appropriate ethical approvals.

Received: 17 February 2024; Accepted: 17 February 2025

Published online: 09 May 2025

References

- Zhang, R. et al. Global burden of ischemic stroke in young adults in 204 countries and territories. Neurology 100 (4), e422-e34 (2023).
- GBD 2016 Stroke Collaborators. Global, regional, and national burden of stroke, 1990–2016: A systematic analysis for the global burden of Disease Study 2016. Lancet Neurol. 18 (5), 439–458 (2019).
- 3. Feigin, V. L., Lawes, C. M., Bennett, D. A., Barker-Collo, S. L. & Parag, V. Worldwide stroke incidence and early case fatality reported in 56 population-based studies: A systematic review. *Lancet Neurol.* 8 (4), 355–369 (2009).
- Chen, H. S., Hsieh, C. F., Chau, T. T., Yang, C. D. & Chen, Y. W. Risk factors of in-hospital mortality of intracerebral hemorrhage and comparison of ICH scores in a Taiwanese population. Eur. Neurol. 66 (1), 59–63 (2011).
- O'Donnell, M. J. et al. Risk factors for ischaemic and intracerebral haemorrhagic stroke in 22 countries (the INTERSTROKE study): A case-control study. Lancet 376 (9735), 112–123 (2010).
- 6. Feigin, V. L. et al. Risk factors for subarachnoid hemorrhage: An updated systematic review of epidemiological studies. Stroke 36 (12), 2773–2780 (2005).
- 7. Olum, S. et al. Stroke mortality outcomes in Uganda. J. Stroke Cerebrovasc. Dis. 30 (5), 105661 (2021).
- 8. Unnithan, A. K. A. & Mehta, P. Hemorrhagic Stroke. StatPearls (StatPearls Publishing Treasure Island (FL), (2022).
- 9. Toyoda, K. et al. Twenty-year change in severity and outcome of ischemic and hemorrhagic strokes. *JAMA Neurol.* **79** (1), 61–69 (2022).
- 10. Ecker, S. et al. Psychological outcome after hemorrhagic stroke is related to functional status. J. Stroke Cerebrovasc. Dis. 31 (8), 106492 (2022).
- 11. Johnson, K. et al. Precision medicine, AI, and the future of personalized health care. Clin. Transl. Sci. 14 (1), 86-93 (2021).
- 12. Wang, H. L. et al. Automatic machine-learning-based outcome prediction in patients with primary intracerebral hemorrhage. *Front. Neurol.* **10**, 910 (2019).
- 13. Matsuo, R. et al. Smoking status and functional outcomes after acute ischemic stroke. Stroke 51 (3), 846-852 (2020).
- 14. Sreekrishnan, A. et al. Intracerebral Hemorrhage Location and functional outcomes of patients: A systematic literature review and Meta-analysis. *Neurocrit Care*. **25** (3), 384–391 (2016).
- 15. Moon, B. H. et al. Use of APACHE II and SAPS II to predict mortality for hemorrhagic and ischemic stroke patients. J. Clin. Neurosci. 22 (1), 111–115 (2015).
- Abdelghany, H., Elsayed, M., Elmeligy, A. & Hatem, G. Prediction of acute cerebrovascular stroke disability using mSOAR score (stroke subtype, Oxfordshire Community Stroke Project, age, mRS and NIHSS). Egypt. J. Neurol. Psychiatry Neurosurg. 59 (1), 21 (2023).
- 17. Karamchandani, R. R., Prasad, T., Strong, D., Rhoten, J. B. & Asimos, A. W. A tool to improve stroke outcome prediction: The Charlotte large artery occlusion endovascular therapy outcome score. *J. Stroke Cerebrovasc. Dis.* 31 (5), 106393 (2022).
- 18. Abedi, V. et al. Predicting short and long-term mortality after acute ischemic stroke using EHR. J. Neurol. Sci. 427, 117560 (2021).
- 19. Li, H. et al. Predicting mortality in acute ischaemic stroke treated with mechanical thrombectomy: Analysis of a multicentre prospective registry. *BMJ open.* 11 (4), e043415 (2021).
- 20. Mattishent, K. et al. Prognostic tools for early mortality in hemorrhagic stroke: Systematic review and Meta-analysis. JCN 11 (4), 339–348 (2015).
- 21. Saposnik, G., Guzik, A. K., Reeves, M., Ovbiagele, B. & Johnston, S. C. Stroke prognostication using age and NIH stroke scale: SPAN-100. *Neurology* 80 (1), 21–28 (2013).
- Ingeman, A., Andersen, G., Hundborg, H. H., Svendsen, M. L. & Johnsen, S. P. In-Hospital Medical complications, length of Stay, and Mortality among Stroke Unit patients. Stroke 42 (11), 3214–3218 (2011).
- 23. García-Rudolph, A. et al. Predicting length of stay in patients admitted to stroke rehabilitation with severe and moderate levels of functional impairments. *Med. (Baltim).* **99** (43), e22423 (2020).
- 24. Kurtz, P., Peres, I. T., Soares, M., Salluh, J. I. F. & Bozza, F. A. Hospital length of Stay and 30-Day mortality prediction in stroke: A machine learning analysis of 17,000 ICU admissions in Brazil. *Neurocrit Care.* 37 (Suppl 2), 313–321 (2022).
- 25. Evans, A. et al. Can differences in management processes explain different outcomes between stroke unit and stroke-team care? *Lancet* 358 (9293), 1586–1592 (2001).
- 26. Turner, M. et al. The impact of stroke unit care on outcome in a Scottish stroke population, taking into account case mix and selection bias. *J. Neurol. Neurosurg. Psychiatry.* **86** (3), 314–318 (2015).
- 27. Fu, C. Y. et al. Age itself or age-associated comorbidities? A nationwide analysis of outcomes of geriatric trauma. Eur. J. Trauma. Emerg. Surg. 48 (4), 2873–2880 (2022).
- 28. Jallow, E. et al. Current status of stroke in Qatar: Including data from the BRAINS study. JRSM Cardiovasc. Disease. 8, 2048004019869160 (2019).
- 29. Grieten, J. et al. Hospital-acquired infections after acute ischaemic stroke and its association with healthcare-related costs and functional outcome. *Acta Neurol. Belgica.* 122 (5), 1281–1287 (2022).
- 30. Grossmann, I. et al. Stroke and pneumonia: Mechanisms, risk factors, management, and prevention. Cureus 13 (11), e19912 (2021).
- 31. Nazarova, M. et al. Multimodal assessment of the motor system in patients with chronic ischemic stroke. *Stroke* **52** (1), 241–249 (2021).
- 32. Carval, T. et al. Outcomes of patients admitted to the ICU for acute stroke: A retrospective cohort. *BMC Anesthesiol.* **22** (1), 235 (2022).
- 33. Wang, Y. et al. Blood glucose level affects prognosis of patients who received intravenous thrombolysis after acute ischemic stroke? A meta-analysis. Front. Endocrinol. (Lausanne). 14, 1120779 (2023).
- 34. Chen, R., Ovbiagele, B. & Feng, W. Diabetes and stroke: Epidemiology, pathophysiology, pharmaceuticals and outcomes. *Am. J. Med. Sci.* **351** (4), 380–386 (2016).
- 35. Weir, C. J., Murray, G. D., Dyker, A. G. & Lees, K. R. Is hyperglycaemia an independent predictor of poor outcome after acute stroke? Results of a long-term follow up study. *Bmj* 314 (7090), 1303–1306 (1997).

- Quast, M. J. et al. Perfusion deficit parallels exacerbation of cerebral ischemia/reperfusion injury in hyperglycemic rats. J. Cereb. Blood Flow. Metab. 17 (5), 553–559 (1997).
- 37. Wajngarten, M. & Silva, G. S. Hypertension and stroke: Update on treatment. Eur. Cardiol. 14 (2), 111-115 (2019).
- 38. Pringle, E. et al. Systolic blood pressure variability as a risk factor for stroke and cardiovascular mortality in the elderly hypertensive population. *I. Hypertens.* **21**(12), (2003).
- 39. Gosmanova, E. O. et al. Association of systolic blood pressure variability with mortality, coronary heart disease, stroke, and renal disease. *J. Am. Coll. Cardiol.* **68** (13), 1375–1386 (2016).
- 40. Du, J. et al. Association of mean platelet volume and platelet count with the development and prognosis of ischemic and hemorrhagic stroke. *Int. J. Lab. Hematol.* **38** (3), 233–239 (2016).
- Mayda-Domaç, F., Mısırlı, H. & Yılmaz, M. Prognostic role of mean platelet volume and platelet count in ischemic and hemorrhagic stroke. J. Stroke Cerebrovasc. Dis. 19 (1), 66–72 (2010).

Acknowledgements

Open Access funding is provided by the Qatar National Library.

Author contributions

Conceptualization: AAA, YI, IA, IMAFormal analysis: AAA, IA, IMAData Curation, Methodology, Writing – original draft, Writing – review & editing: AAA, IA, YI, AJN, SY, NA, IMAAll authors read and approved the final manuscript.

Funding

The Medical Research Center at Hamad Medical Corporation, Qatar (Grant No. MRC-01-22-594), funded this study.

Declarations

Competing interest

The authors declare no competing interests.

Ethical approval

The Medical Research Center of Hamad Medical Corporation IRB approved this project (MRC-01-22-594). The research complies with the ethical principles outlined in the Helsinki Declaration of 1964 and its subsequent modifications and related ethical norms. The IRB waived the need for informed consent at the Medical Research Center of Hamad Medical Corporation.

Additional information

Correspondence and requests for materials should be addressed to A.J.N.

Reprints and permissions information is available at www.nature.com/reprints.

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

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 https://creativecommons.org/licenses/by/4.0/.

© The Author(s) 2025