

## Risk factors and outcomes of stroke in a tertiary hospital in Hadhramout Governorate, Yemen

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### ABSTRACT

**Background:** Although stroke is a major problem in Hadhramout Governorate, there is a scarcity of reliable information on risk factors of stroke and predictors of in-hospital mortality. The aim of this study was to explore the risk factors and outcomes of stroke patients admitted to Ibn Sina Hospital, Hadhramout, Yemen, and to identify the predictors of in-hospital mortality. **Methods and Materials:** This retrospective cross-sectional study was conducted in Ibn Sina Hospital in Mukalla district, Hadhramout Governorate, over a 4-month period (from January 1, 2021, to April 30, 2021). **Results:** During the study period, we recruited 100 cases of stroke, of whom 77 (77%) were male and 23 (23%) were female. Their mean age was  $65.42 \pm 12.78$  years. Hypertension was the most common risk factor, occurring in 81 (81%) patients, while no risk factors were identified in 5 (5%) cases (Cryptogenic). Ischemic stroke was found in 70 (70%) cases, and hemorrhagic stroke was noticed in 30 (30%) patients. The in-hospital mortality was 29 (29%), and the univariate analysis found male sex, hypertension, and hemorrhagic stroke as probable predictors of in-hospital mortality. Only hemorrhagic stroke (adjusted odds ratio [OR]=2.053 and 95% confidence interval [CI]=0.822–1.599;  $p < 0.001$ ) and hypertension (adjusted OR= 1.677; 95% CI=0.555–1.495;  $p = 0.011$ ) were found to be independent predictors of mortality by multivariate logistic regression analysis. **Conclusion:** Stroke is a major problem in Hadhramout Governorate with ischemic stroke being more than hemorrhagic. Men were more likely to have a stroke than women and majority of cases were elderly. Hypertension, diabetes mellitus, smoking, and dyslipidemia were the most commonly identified risk factors that were significantly associated with stroke. Hemorrhagic stroke and presence of hypertension were found to be risk factors for in-hospital mortality; therefore, hypertension should be well-controlled to reduce in-hospital mortality.

**Key words:** Hadhramout Governorate, Hemorrhagic stroke, In-hospital mortality, Ischemic stroke, Risk factors

Stroke is the second leading cause of death worldwide after ischemic heart disease. Each year, there are 15 million strokes and 5.8 million stroke-related deaths, with more than 65% of these deaths occurring in developing countries [1-3]. In addition, stroke is the third most prevalent cause of disability, because it makes victims dependent on society, which affects their personal, social and economic life. Stroke poses a serious threat not only to public health but also to the socioeconomic development of society. The annual cost of treating stroke victims in the United States of America is approximately \$34.3 billion [4,5]. The risk factors and lifestyles differ in various countries, requiring investigation of stroke risk factors in each country to identify the pattern of the disease and issue the appropriate educational programs, to reduce the burden of stroke and minimize the high

costs of rehabilitation and other unfavorable effects of stroke on the population [6,7].

Stroke is becoming an important cause of premature death and disability in low-income countries like Yemen. The latest figures from the World Health Organization (WHO) show that 13,570 stroke deaths accounting for 8.72% of all deaths occurred in Yemen in 2020. The age-adjusted death rate is 127.49/100,000 population, ranking Yemen 32<sup>nd</sup> in the world [8]. Part of the reason for this is that the civil war has led to a lack of well-run health-care facilities.

In Yemen, there is a scarcity of reliable information on stroke risk factors and outcome predictors, as only a few studies have addressed these issues [9-12]. We conducted this study to explore the risk factors and outcomes of stroke patients admitted to Ibn Sina hospital, Hadhramout, Yemen, and to identify the predictors of in-hospital mortality.

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## PATIENTS AND METHODS

### Study Design, Population, and Setting

A retrospective cross-sectional study was conducted in Ibn Sina Hospital in Mukalla, Hadhramout Governorate, over a 4-month period (January 1, 2021–April 30, 2021). Ibn Sina Hospital is the largest government hospital in Mukalla, in which medical and health institute students get their training and is a tertiary hospital that serves about 4,000,000 people [10]. Mukalla is a seaport and the capital city of Yemen's largest governorate, Hadhramout. The city lies in the southern part of the Arabian Peninsula on the Gulf of Aden, extending along the shores of the Arabian Sea, about 480 km (300 miles) east of Aden.

### Definitions, Inclusion and Exclusion Criteria

Patients were included in the study if they had experienced an acute or a sub-acute stroke confirmed by computed tomography (CT) scans and/or magnetic resonance imaging (MRI) scan. They also had to meet the criteria of the American Heart Association/American Stroke Association, which define an ischemic stroke as “an episode of neurological dysfunction caused by focal cerebral, spinal, or retinal infarction” and a hemorrhagic stroke as “rapidly developing clinical signs of neurological dysfunction attributable to a focal collection of blood within the brain parenchyma or ventricular system that is not caused by trauma [13]. Patients presented with transient ischemic attack, subdural, or epidural hematoma and symptoms caused by recent trauma were excluded from the study.

We defined the main risk factors as follows. Hypertension was considered positive if the patient is using medication for hypertension or if three consecutive blood pressure estimation readings were above 140/90 mm Hg. Diabetes mellitus was diagnosed if there was a history of diabetes mellitus or if the patient was receiving drugs for diabetes or their random blood sugar levels were above 200 mg (11.1 mmol/L) with symptoms, or if fasting blood sugar scored more than 126 mg (7 mmol/L) or HbA1c >6.5% for more than two readings during hospitalization. Hyperlipidemia was considered positive if lipid profile results show total cholesterol >200 mg/dL and/or triglyceride >150 mg/dL [14]. Heart disease was positive if history, clinical examination, available investigations on admission, or requested investigations during hospitalization yielded one or more of the following: Rheumatic heart disease, ischemic heart disease, atrial fibrillation, and cardiomyopathy. Habits of smoking and khat chewing were positive if patients were using them daily for a duration of at least more than a year.

### Data Collection and Source of Data

We obtained the list of the stroke patients from the medical record. We recruited the following variables on a special data collection sheet developed for the sake of the study: Age, gender, risk factors such as hypertension, diabetes mellitus, smoking, dyslipidemia,

smoking, khat chewing, history of previous attack, and the family history of stroke. Furthermore, we looked for coronary artery disease, and atrial fibrillation. The course of the disease during hospitalization and outcome was recorded as alive on discharge or died during hospitalization (in-hospital mortality).

### Ethical Approval and Data Analysis

The study was approved by the administration of Ibn Sina Hospital. Data were reported as means±standard deviation (SD) for quantitative variables, while qualitative variables were described as numbers and percentages. Univariate analysis was performed to determine the probable risk factors for in-hospital mortality. Variables with  $p < 0.10$  in the univariate analysis were entered in the multiple logistic regression to identify the independent risk factors for in-hospital mortality at  $p < 0.05$ . Data analysis was performed with SPSS software (v 23; IBM Corp, Armonk, NY, USA).

## RESULTS

During the study period, we recruited 100 consecutive stroke cases, of which 77 (77%) were male and 23 (23%) were female. Their mean age was  $65.42 \pm 12.78$  years (range: 35-100 years). Arterial hypertension was the most common risk factor, occurring in 81 (81%) patients, followed by diabetes mellitus in 58 (58%) patients, smoking in 40 (40%) cases, and dyslipidemia in 39 (39%). Family history was positive for stroke in 31 (31%) patients and history of previous stroke was present in 30 (30%) patients. Risk factors were not identified (Cryptogenic) in 5 (5%) cases (Table 1). COVID-19 preceded stroke in two cases with cryptogenic stroke.

Ischemic stroke was identified in 70 (70%) cases, while hemorrhagic stroke was found in 30 (30%) patients. None of the patients were treated with a thrombolytic therapy. The in-hospital mortality was 29 (29%), and all who survived ischemic strokes were prescribed aspirin and statin on discharge. The univariate analysis found the following variables to be probable predictors of in-hospital mortality: Male sex, hypertension, and hemorrhagic stroke (Table 2). Only hemorrhagic stroke (adjusted odds ratio [OR]=2.053; and 95% confidence interval [CI]=0.822–1.599;  $p < 0.001$ ) and hypertension (adjusted OR=1.677; 95% CI=0.555–1.495;  $p = 0.011$ ) were found to be independent predictors of mortality by multivariate logistic regression analysis (Table 3).

## DISCUSSION

In Yemen, stroke is a devastating problem, especially after the civil war, which destroyed the fragile pre-existing infrastructures, including the health sector. Only around 51% of the hospitals and health-care facilities are in full working condition [15], while many Yemeni healthcare workers emigrate from the country in search of better salaries and safer conditions elsewhere [16], leaving the population with limited health manpower and little room to respond to stroke or other public health events.

**Table 1: Demographic and clinical characteristics of the 100 patients involved**

Variables	Frequency (%)
Sex	
Male	77 (77)
Female	23 (23)
Age±SD (range) years	65.42±12.78 (35–100)
Risk factors	
Hypertension	81 (81)
Diabetes mellitus	58 (58)
Smoking	40 (40)
Dyslipidemia	39 (39)
Family history of stroke	31 (31)
History of stroke	30 (30)
Khat chewing	16 (16)
Chronic kidney disease	8 (8)
Coronary artery disease	4 (4)
Atrial fibrillation	3 (3)
Cryptogenic (no risk factors)	5 (5)
Number of risk factors	
1	14 (14)
>1	81 (81)
Type of stroke	
Ischemic	70 (70)
Hemorrhagic	30 (30)
Intracerebral hemorrhage	28 (93.3)
Subarachnoid hemorrhage	2 (6.7)
Outcome	
Alive	71 (71)
Died	29 (29)

SD: Standard deviation

**Table 2: Univariate analysis of predictors of in-hospital mortality**

Variables	Unadjusted odds ratio (95% confidence interval)	p-value
Gender – Male	1.422 (0.228–2.809)	0.10
Age	0.677 (0.255–2.799)	0.434
Hypertension	1.055 (0.365–1.071)	<0.001
Dyslipidemia	1.138 (0.092–12.063)	0.920
Smoking	1.003 (0.991–4.015)	0.632
Diabetes mellitus	1.010 (0.992–2.027)	0.286
Family history of stroke	1.467 (1.264–2.701)	0.566
History of stroke	1.032 (0.333–3.299)	0.954
Khat chewing	0.932 (0.906–0.959)	0.993
Chronic kidney disease	1.406 (0.899–3.777)	0.286
Coronary artery disease	1.916 (0.745–4.925)	0.177
Atrial fibrillation	0.431 (0.160–4.159)	0.955
Cryptogenic (no risk factors)	1.807 (0.846–9.311)	0.823
Ischemic stroke	1.945 (0.698–5.417)	0.203
Hemorrhagic stroke	2.053 (0.822–1.599)	<0.001

Yemen's civil war and subsequent humanitarian crises have diverted the attention of the Yemeni authorities away from health

**Table 3: Multivariate analysis of predictors of in-hospital mortality**

Variables	Adjusted odds ratio (95% confidence interval)	p-value
Male sex	0.988 (1.228–6.809)	0.655
Hypertension	1.677 (0.555–1.495)	0.011
Hemorrhagic stroke	1.558 (0.365–1.996)	<0.001

problems such as stroke. The biggest challenge for Yemeni's health care system, which is to provide care, that is, affordable and accessible to its population is hampered by a lack of humanpower and inadequate resources such as diagnostics, pathology services and stock-outs. Regarding the stroke management in Yemen, the country lacks well-run public health-care facilities such as rehabilitation facilities and stroke units with MRI/CT capabilities. Furthermore, given the state of the economy, the government is unable to provide thrombolysis therapy, which is the gold standard of care for acute ischemic stroke. As a result, most people rely on the private sector for health care, which provides almost 80% of health services. However, the private health sectors are providing some of the facilities such as CT scan and MRI at high prices which are out of reach for *most people*.

The true stroke incidence and prevalence in Yemen is yet unknown, due to lack of stroke data banks and stroke registries. Moreover, with the highly privatized nature of health-care service, most patients are unable to access private health facilities due to financial constraints. Therefore, many people, especially the elderly, are left at home without being diagnosed or counted.

In the present study, the mean age of stroke patients was 65.42 years, which is consistent with a study by Dey *et al.* [17], who reported a mean age of 68.41 years, and Al-Shenqiti [7], who reported a mean age of 67.52 years, but higher than that mentioned by Nour *et al.*, who reported a mean age of 59.61 years [18].

Increased burden of stroke among aged population can be attributed to an increase in risk factors such as cardiovascular disease, hypertension, diabetes mellitus, and ischemic heart disease [17]. As noted, about 77% of the patients involved in this study were male, which is consistent with studies from Saudi Arabia [7], India [17,19], Ethiopia [20], and Kenya [21]. The global trend toward male dominance is not fully understood but could be partly explained by the fact that hypertension, diabetes mellitus, smoking, and alcoholism are more common among male patients.

The results of this study are consistent with the previous local studies [9-12], which showed that ischemic stroke was the most frequent form, followed by hemorrhagic stroke. This is consistent with studies from Saudi Arabia [7], India [17,19], Ethiopia [20], and Kenya [21], but unlike a study conducted at St. Paul Hospital, which found that hemorrhagic strokes made up 61.3% of cases, making them more frequent than ischemic strokes [22].

As noted in this study, hypertension, which accounts for 81% of cases, was the major risk factor associated with stroke. This finding is consistent with earlier studies conducted in hospital settings in our country [9-12] and other Arab and African countries [7,18,20-25]. The degree of mental stress brought on by

poverty and instability, particularly after the civil war, is probably closely correlated with the frequency of hypertension. The fact that many of our patients had poor blood pressure management caused by non-adherence to treatment due to cost and drug availability certainly increased the significance of hypertension control. As noted in this study, diabetes mellitus ranked second on the list of risk factors as did other hospital studies around the world [7,17-26].

The present study showed that smoking was a significant risk factor, occurring in 40% of all stroke cases, which is high compared to previous local studies conducted earlier in the same hospital [11,12], which reported smoking in 21% and 26% of cases, respectively. The difference in figures between two periods reflects the changes in the lifestyle of the population in this governorate. Dyslipidemia was found in 39% of the cases, which is high as compared with previous local studies [9-12], but consistent with a study by Qawasmeh *et al.* [24] who reported a frequency of 37% and lower than that reported by Khan *et al.*, who observed a frequency of 52.5% [23].

In the present study, khat chewing was reported in 16% of cases, which is lower than in a prior study [9]. Chewing khat is a common habit in Yemen. It is estimated that up to 90% of adult men and 73% of adult women chew khat for 3–4 h daily in this country. In addition, 15–20% of children under the age of 12 are also daily consumers [27]. Khat is a stimulant made from the leaves of the plant *Catha edulis*. Some reports from Yemen [28,29] observed a correlation between khat chewing habit and cardiovascular diseases. Although these studies have significant limitations, they highlight the necessity for additional large-scale trustworthy studies to confirm the relevance of khat chewing as a risk factor for cardiovascular disease, including stroke.

As previously mentioned, 5% of the cases in this study had no risk factors. Interestingly, two of these instances had ischemic stroke preceded by prior COVID-19, implying that COVID-19 may play a role in stroke pathogenesis. A review of the literature revealed conflicting results, while Qureshi *et al.* [30] concluded that acute ischemic stroke was uncommon in patients with COVID-19 and typically occurs in the presence of other cardiovascular risk factors, other reports found that the risk of stroke and heart attack increases sharply in the weeks following a COVID-19 diagnosis [31-33], suggesting an interplay of vascular risk factors and immune responses to the SARS-CoV-2 virus as a mechanism. A recent meta-analysis [34] concluded that when a patient is admitted with COVID-19, the risk of ischemic stroke must be considered. In general, the design of the present study was descriptive making it impossible to draw meaningful inferences about the relationship of these factors with stroke.

In-hospital mortality ranges between 2.8% and 46% [9,17,18]. In-hospital mortality in the present study was 29%, which falls within the global range. Many risk factors for in-hospital mortality were identified. Russell *et al.* [35] found the following variables as independent risk factors for in-hospital mortality: hypertension, previous stroke, Glasgow coma scale <8, clinical

diagnosis in the absence of imaging, hemorrhagic stroke, and aspiration pneumonia. While Kamabu *et al.* [36] found hemorrhagic stroke, hospital arrival delay of more than 24 h, poor compliance to anti-diabetic medication, recurrent stroke, poor compliance to antihypertensive therapy, and the advanced age >75 years as independent predictors of in-hospital mortality. Hemorrhagic stroke and hypertension were revealed to be independent predictors of in-hospital mortality after stroke in the current study.

This study has several limitations. First, it is a retrospective hospital-based study, where patient follow-up was not feasible as we mainly depend on the patient's medical records. Second, the sample size was small, although the original study was designed to include all stroke cases in 2021. However, in the retrospective review of medical records, only 100 medical records were consecutive and contained enough information to be included in the study. Third, denial of alcohol consumption can be incorrect, because it is a social and religious taboo. Finally, cardioembolic work-ups were incomplete. Future studies could include a more complete and rigorous evaluation of possible cardiovascular etiologies by having cardiologists as a part of the investigation team.

## CONCLUSION

Stroke is a major problem in Hadhramout Governorate with ischemic stroke being more than hemorrhagic. Men were more likely to have a stroke than women and majority of cases were elderly. Hypertension, diabetes mellitus, smoking, and dyslipidemia were the most commonly identified risk factors that were significantly associated with stroke. Hemorrhagic stroke and presence of hypertension were found as risk factors for in-hospital mortality; therefore, hypertension should be well-controlled to reduce in-hospital mortality. The identification of additional risk factors for stroke is crucial to the improvement of population health.

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