

ORIGINAL ARTICLE

## Frequency, types and causes of anaemia in ischemic stroke patients admitted to medical wards of Khyber Teaching Hospital.

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**ABSTRACT... Objective:** To determine the frequency, types and causes of anaemia in ischemic stroke patients admitted to medical wards of Khyber Teaching Hospital Peshawar. **Study Design:** Cross-sectional study. **Setting:** Department of Medicine, Khyber Teaching Hospital, Peshawar. **Period:** 5th December 2023 till 4th June 2024. **Methods:** Convenient sample technique was used. A total 368 patients presenting with stroke were enrolled. The patients were evaluated for the type and cause of anemia. Hb <12.0gm/dl was considered the cut off for anemia. Data was analysed using SPSS version 24. **Results:** Mean age of the patients was 53.34 ± 13.29 years. Male to female ratio was nearly 1: 1. Hypertension was the most common comorbidity observed in 86 patients (23.4%). Normochromic normocytic anemia was the common type observed in 150 patients (40.7%). **Conclusion:** Normochromic normocytic anemia is the most common type of anemia in patients with ischemic stroke. Elderly female patients with underlying chronic disease like disease and prolonged illness are more likely having anemia.

**Key words:** Anemia Types, Hypertension, Ischemic Stroke.

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### INTRODUCTION

Stroke is the second most common cause of mortality worldwide.<sup>1</sup> Any rapidly increasing deficit of one or both sides of the body that lasts more than twenty-four hours is referred to be a stroke. Either an ischemic or hemorrhagic stroke causes the brain's supply of blood and nutrients to be reduced in that area. It is estimated that each year, about 15 million people worldwide suffer from strokes. Of these 15 million individuals, two thirds will die and a third will become disabled. Stroke is the third leading cause of death in developing nations, after cancer and coronary heart disease, respectively. 87% of stroke cases are ischemic strokes, which are the most frequent type of stroke.<sup>2,3</sup>

Stroke is the primary cause of both chronic neurologic impairment and severe functional impairment globally. Slurred speech, abrupt weakening of one side (arm, face, or leg), obscured vision in one or both eyes, problems walking suddenly, and a sudden, intense headache are common clinical signs of stroke.<sup>4</sup> A number of possible risk factors for stroke include alcohol intake, smoking, dyslipidemia, hypertension,

and—most importantly—diabetes mellitus (DM).<sup>5</sup>

According to Jo YJ et al.'s study, among patients who had an ischemic stroke, the most prevalent risk factor was found to be hypertension, smoking, ischemic heart disease, diabetes mellitus, hyperlipidemia, atrial fibrillation, carotid artery stenosis, obesity, and a history of stroke in the family in all age group whereas as per Safeer M et al., common risk factors in the younger group were obesity, smoking, and hypertension, and the older group's most common risk factors were diabetes mellitus, and hypertension.<sup>6,7</sup>

Limited research has been done on risk factors in stroke patients due to cerebral infarction in Pakistan, therefore, the current study aims to assess the prevalence of risk factors among ischemic stroke. It will help in prevention of this disease by providing awareness about the risk factors of ischemic stroke in general population. It will help physicians in better management of stroke and decrease chances of its recurrence.

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The aim of this study was to determine the frequency, types and causes of anaemia in admitted ischemic stroke patients.

## METHODS

A Cross sectional study conducted at Department of Medicine, KTH, Peshawar, from 5th December 2023 till 4th June 2024. The Sampling Technique was Non – probability Consecutive technique. The WHO sample size calculator was used to determine the sample size. Previously reported prevalence of anaemia in ischemic stroke patients= 39.6%.<sup>8</sup> The Margin of Error: d=5%, 95 % confidence level. Expected sample size was 368. All those patients with age 40 to 70 years, both genders and patients presenting with ischemic stroke were included in study. Exclusion criteria included Ischemic stroke patients having recent blood transfusions, hemorrhagic stroke and space occupying brain lesion.

Patients were enrolled from the medical units after receiving approval from the hospital ethics board (286/DME/KMC-11/5/2022). An informed consent was obtained from each patient. Age, gender, educational attainment, monthly income, diabetes, hypertension, smoking, and obesity were among the demographic data that were recorded. Both a physical examination and a thorough history were obtained. A complete blood count, a renal function test, a liver function test, serum electrolytes, a chest x-ray, a routine examination of the stool, and a routine examination of the urine were all performed as baseline investigations (BLIs). On specially created proforma, the causes, forms, and duration of ischemic stroke were recorded.

SPSS version 23 was used to examine the data that was obtained. For quantitative characteristics like age, hemoglobin level, length of ischemic stroke, and monthly income, mean +/- SD was computed. For categorical factors such as gender, medicines used, status of BLIs, types and causes of anemia, and background medical illnesses such as diabetes, hypertension, and obesity, frequency and percentages were determined. Data stratification was used to address effect modifiers such as age, gender, length of disease, diabetes, hypertension, smoking, and obesity. A p-value < 0.05 was deemed

statistically significant.

## RESULTS

The mean age of our study population was 53.34 ± 13.29 years with ranged of 30 to 80 years. The mean BMI was 24.05 ± 2.6 Kg/m<sup>2</sup> [Table-I].

**TABLE-I**

**Demographic characteristics of study participants**

| Parameters                   | Frequency             | Percent |      |
|------------------------------|-----------------------|---------|------|
| Gender                       | Male                  | 189     | 51.4 |
|                              | Female                | 179     | 48.6 |
| Age in years                 | ≤55                   | 183     | 49.7 |
|                              | >55                   | 185     | 50.3 |
| Diabetes                     | Yes                   | 29      | 7.9  |
|                              | No                    | 339     | 92.1 |
| Hypertension                 | Yes                   | 86      | 23.4 |
|                              | No                    | 282     | 76.6 |
| Smoking                      | Yes                   | 54      | 14.7 |
|                              | No                    | 314     | 85.3 |
| Obesity                      | Yes                   | 147     | 39.9 |
|                              | No                    | 221     | 60.1 |
| Disease duration (in months) | ≤3                    | 217     | 59.0 |
|                              | >3                    | 151     | 41.0 |
| Anemia Type                  | Microcytic            | 144     | 39.1 |
|                              | Normocytic            | 150     | 40.8 |
|                              | Macrocytic            | 74      | 20.1 |
| Causes Of Anemia             | Iron Deficiency       | 157     | 42.7 |
|                              | Sickel Cell Disease   | 52      | 14.1 |
|                              | Folate/B12 Deficiency | 159     | 43.2 |

Type of anemia on basis of cell size and cause of anemia with age and gender as shown in Table-II. No significant association of gender and age with types and causes of anemia.

Stratification of type of anemia and cause of anemia with obesity and duration of stroke as shown in Table-III. There is significant association of causes of anemia compared obesity and duration of stroke (P 0.000).

TABLE-II

Cross tabulation of patients` gender and age with types and causes of anemia

| Risk Factors    | Gender                |            |            | Age (Years) |            |            |       |
|-----------------|-----------------------|------------|------------|-------------|------------|------------|-------|
|                 | Male                  | Female     | P value    | ≤55         | >55        | P-Value    |       |
| Types of anemia | Microcytic            | 70 (48.6%) | 74 (51.4%) | 0.564       | 66 (45.8%) | 78 (54.2%) | 0.154 |
|                 | Normocytic            | 82(54.7%)  | 68 (45.3%) |             | 73 (48.7%) | 77 (51.3%) |       |
|                 | Macrocytic            | 37 (50.0%) | 37 (50.0%) |             | 44 (59.5%) | 30 (40.5%) |       |
| Causes          | Iron Deficiency       | 89 (56.7%) | 68 (43.3%) | 0.206       | 77 (49.0%) | 80 (51.0%) | 0.974 |
|                 | Sickle Cell Disease   | 24 (46.2%) | 28 (53.8%) |             | 26 (50.0%) | 26 (50.0%) |       |
|                 | B12/folate deficiency | 76 (47.8%) | 83 (52.2%) |             | 80 (50.3%) | 79 (49.7%) |       |

TABLE-III

Cross tabulation of obesity and stroke duration with types and causes of anemia

| Risk Factors    | Disease Duration (Months) |             |            | Obesity |            |             |       |
|-----------------|---------------------------|-------------|------------|---------|------------|-------------|-------|
|                 | 3 or Below                | More Than 3 | P-Value    | Yes     | No         | P-Value     |       |
| Types of anemia | Microcytic                | 81 (56.3%)  | 63 (43.8%) | 0.336   | 51 (35.4%) | 93 (64.6%)  | 0.292 |
|                 | Normocytic                | 95 (63.3%)  | 55 (36.7%) |         | 62 (41.3%) | 88 (58.7%)  |       |
|                 | Macrocytic                | 41 (55.4%)  | 33 (44.6%) |         | 34 (45.9%) | 40 (54.1%)  |       |
| Causes          | Iron Deficiency           | 72 (45.9%)  | 85 (54.1%) | 0.000   | 65 (41.4%) | 92 (58.6%)  | 0.000 |
|                 | Sickle Cell Disease       | 37 (71.2%)  | 15 (28.8%) |         | 38 (73.1%) | 14 (26.9%)  |       |
|                 | B12/folate deficiency     | 108 (67.9%) | 51 (32.1%) |         | 44 (27.7%) | 115 (72.3%) |       |

TABLE-IV

Stratification of anemia types and causes with comorbidities (Diabetes and Hypertension)

| Risk Factors    | Diabetes              |            |              | Hypertension |            |             |       |
|-----------------|-----------------------|------------|--------------|--------------|------------|-------------|-------|
|                 | Yes                   | No         | P value      | Yes          | No         | P-Value     |       |
| Types of anemia | Microcytic            | 19 (13.2%) | 125 (86.8%)  | 0.000        | 49 (34.0%) | 95 (66.0%)  | 0.000 |
|                 | Normocytic            | 0 (0.0%)   | 150 (100.0%) |              | 17 (11.3%) | 133 (88.7%) |       |
|                 | Macrocytic            | 10 (13.5%) | 64 (86.5%)   |              | 20 (27.0%) | 54 (73.0%)  |       |
| Causes          | Iron Deficiency       | 9 (5.7%)   | 148 (94.3%)  | 0.419        | 32 (20.4%) | 125 (79.6%) | 0.135 |
|                 | Sickle Cell Disease   | 5 (9.6%)   | 47 (90.4%)   |              | 9 (17.3%)  | 43 (82.7%)  |       |
|                 | B12/folate deficiency | 15 (9.4%)  | 144 (90.6%)  |              | 45 (28.3%) | 114 (71.7%) |       |

Cross tabulation between type of anemia, causes of anemia with obesity and duration of stroke as shown in Table-III. There is significant association of types of anemia compared obesity and duration of stroke (P 0.000), whereas no significant association compared to causes of anemia.

## DISCUSSION

The most prevalent morphological kind of anemia was normocytic normochromic (40.8%) blood image, which was followed by microcytic hypochromic anemia. In contrast, the study's

macrocytic normochromic anemia was the least morphologically significant kind of anemia. This study's normocytic normochromic blood picture of anemia is not surprising because it has been documented in several earlier studies conducted in Benin, Pakistan, India, and other countries.<sup>9-11</sup>

Additionally, compared to those of normal weight, hospitalized obese patients had a twofold increased risk of developing anemia. The outcome was consistent with research done in Austria, Pakistan, Iran, and the United States of America.<sup>12-14</sup>

Low-grade systemic inflammation, seen in obese individuals, is the main mechanism tying obesity and anemia together.<sup>15</sup> Patients with obesity have significantly higher serum levels of hepcidin and interleukin-6, which lowers their iron intake since their liver and splenic macrophages sequester iron and they absorb less of it.<sup>15</sup> Malnourished patients had a 2.59-fold higher incidence of anemia than normal admitted patients. The outcome was consistent with research from India and the Netherlands.<sup>11,16</sup> Malnutrition is associated with poor treatment tolerance, reduced quality of life, higher healthcare expenses, poverty, and medical conditions like intestinal obstruction and malabsorption syndrome.<sup>11,17</sup>

Patients with diabetes mellitus at the time of admission were 3.20 times more likely to have anemia. The following causes could be the cause of this: Anemia is frequently caused by bleeding, especially in the gastrointestinal tract in patients with cirrhosis and chronic liver disease.<sup>18</sup> A chronic liver disease may result in anemia from bleeding because of thrombocytopenia, decreased platelet activity, or a deficiency in liver produced clotting factors.<sup>19</sup> Additionally, hepcidin, the main regulator of iron homeostasis, is secreted by the liver making iron deficiency prevalent in individuals with advanced chronic liver disease (CLD).<sup>20</sup>

Compared to COPD patients, admitted patients with hypertension had an 11.2-fold increased risk of anemia. This may be because individuals with chronic kidney disease (CKD) are more likely to experience anemia due to the fact that their failing kidneys generate less erythropoietin.<sup>11,22</sup> Nearly half of individuals with CKD have normocytic normochromic anemia, which is caused by peritubular fibroblasts in the kidneys producing less erythropoietin than usual.<sup>22</sup>

Anemia was more common in patients with longer illness durations than in COPD patients who were hospitalized. The primary causes of anemia due malaria are rapid hemolysis of parasitized red blood cells.<sup>23</sup> This may be the result of infections raising hepcidin levels, which also exacerbate anemia. The prevalence of anemia in HIV-positive people rises as the illness progresses, leading to changes

in iron metabolism, increased hepcidin, anemia of inflammation, and a persistent acute phase response.<sup>24</sup>

## CONCLUSION

Normochromic normocytic anemia is the most common type of anemia in patients admitted with ischemic stroke in our population followed by microcytic anemia while macrocytic anemia is the least common type. Elderly female with previous comorbidities like diabetes mellitus and prolonged disease duration are more likely develop anemia. Iron deficiency is most consistent cause of anemia which is more likely in patients with underlying chronic disease.

## CONFLICT OF INTEREST

The authors declare no conflict of interest.

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#### AUTHORSHIP AND CONTRIBUTION DECLARATION

|   |  |
|---|--|
| 1 | <b>Mohammad Haroon:</b> Conceptualization, writing, supervision. |
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| 3 | <b>Irfan Ullah:</b> Statistical analysis.                        |
| 4 | <b>Shah Umam:</b> Methodology.                                   |
| 5 | <b>Osama Ali Khan:</b> Writing.                                  |
| 6 | <b>Ejaz Khan:</b> Data entry.                                    |