

# Patterns of geriatric anemia: A hospital-based observational study in North India

Dheeraj Sharma<sup>1</sup>, Vikas Suri<sup>1</sup>, Ashok K. Pannu<sup>1</sup>, Savita V. Attri<sup>2</sup>,  
Neelam Varma<sup>3</sup>, Rakesh Kochhar<sup>4</sup>, Subhash Varma<sup>1</sup>, Savita Kumari<sup>1</sup>

Departments of <sup>1</sup>Internal Medicine, <sup>2</sup>Paediatric Biochemistry, <sup>3</sup>Hematology and <sup>4</sup>Adult Gastroenterology, Post Graduate Institute of Medical Education and Research, Chandigarh, Punjab, India

## ABSTRACT

**Background:** Geriatric anemia is a global health problem because of its high prevalence and associated significant morbidity and mortality. **Aim:** The objectives of this study were to estimate the pattern of anemia in the elderly patients and the underlying etiology of anemia. **Research Design and Methods:** This was a hospital-based prospective observational study, conducted in patients aged 60 years and above at PGIMER, Chandigarh, a tertiary care center of North India. Anemia is defined as hemoglobin level less than 13 g/dl in men and 12 g/dl in women. **Results:** Among the 105 older patients with anemia, the mean value of hemoglobin was  $8.8 \pm 2.3$  g/dl. The etiological distribution of anemia was iron deficiency in 26 patients (24.8%), chronic disease in 24 patients (22.9%), hematological disorders in 21 (20%), chronic kidney disease in 13 (12.4%), multifactorial in 8 (7.6%), vitamin B12 deficiency in 2 (1.9%), folate deficiency in 1 (0.9%), and hypothyroidism in 1 patient (0.9%). No etiology could be found in 9 patients (8.6%). 57.6% of the iron-deficient patients had upper gastrointestinal lesions and 30.7% had a nutritional cause. Common chronic diseases causing anemia were malignancy (36.6%) and liver disease (29.1%). The myelodysplastic syndrome was the commonest hematological disorder. 53.35% of the patients had normocytic anemia, 40% had microcytic anemia, and 6.6% had macrocytic anemia. **Conclusions:** In most of the cases, anemia in the elderly had a treatable cause. Thus, a thorough investigation including gastrointestinal endoscopy is warranted. Unexplained progressive or unresponsive anemia requires bone marrow examination.

**Keywords:** Anemia, elderly, geriatrics

## Introduction

Anemia is a global health problem in older adult population because of the high prevalence and associated significant morbidity and mortality.<sup>[1,2]</sup> The reported prevalence of anemia in the elderly is 2.9%–51% and correlates with advanced age and multiple related conditions, including iron deficiency, inflammatory conditions, malignancy, and low serum erythropoietin.<sup>[2–4]</sup> The simultaneous occurrence of multiple causes of anemia is common in elderly.

Elderly patients with anemia are heterogeneous in terms of clinical history, coexisting medical conditions, and concomitant medication use than young adults. In elderly, anemia is associated with poor performance status, increased frailty, dementia, depression, reduced mobility, increased risk of falls, and poor quality of life.<sup>[5–12]</sup> Anemia portends worse prognosis in elderly patients with cardiovascular and other chronic illnesses. Studies have reported a survival benefit with the treatment of geriatric anemia.<sup>[13,14]</sup> An intensive effort should always be made to reach an etiological diagnosis for better management of these patients.

In the present study, we aimed to prospectively investigate the pattern of anemia in the elderly Indian patients and the underlying etiology of anemia.

**Address for correspondence:** Dr. Savita Kumari,  
4<sup>th</sup> Floor, F Block, Department of Internal Medicine, PGIMER,  
Chandigarh, Punjab, India.  
E-mail: savita349@yahoo.com

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## Materials and Methods

### Study population

This was a hospital-based prospective observational study, conducted in a tertiary care center of North India, from July 2014 to December 2015.

### Case definition

Older patients aged 60 years and above presenting to the medical outpatient department or medicine ward of Internal Medicine with clinically suspected and laboratory-confirmed anemia were included. World Health Organization (WHO) criteria, hemoglobin values less than 12 g/dl in women and less than 13 g/dl in men, were used to define anemia.<sup>[15]</sup> We excluded patients who had received a blood transfusion during the last 12 weeks and patients on chemotherapy or radiotherapy.

Standard protocol approvals and patient consent: The Institutional Ethics Committee approved the study. We obtained written informed consent from all study participants.

### Data collection

The number of patients enrolled during the study period determined the sample size. On enrolment, a clinical research form including a medical and dietary history, sociodemographic data, and physical examination was completed, and samples of blood, urine, and stool were collected. If the clinical condition permitted it, a bone marrow aspirate was obtained under local anesthesia. All conditions were managed according to standard protocols.

### Laboratory assays

A complete blood count, including red blood cell (RBC) count, i.e. hemoglobin, hematocrit, reticulocyte count; RBC indices, i.e. mean cell volume (MCV), mean cell hemoglobin (MCH), MCH concentration, red cell distribution width; white blood cell count (cell differential and nuclear segmentation of neutrophils); platelet count; and cell morphology, i.e. cell size, hemoglobin content, anisocytosis, poikilocytosis, and polychromasia, was performed. Serum iron study including serum iron, total iron binding capacity, and serum ferritin was done for all patients. Serum levels of folate and vitamin B<sub>12</sub> were done in patients with dimorphic and macrocytic anemia or in patients with normocytic or microcytic blood picture in which no other cause of anemia was found. Hemoglobin electrophoresis was done in patients with reduced MCV or MCH and normal iron studies. Bone marrow studies (aspiration and/or biopsy) were carried out on patients with blood smear showing immature white cells or nucleated red cells, indeterminate status of iron stores and unexplained progressive or unresponsive anemia. Renal function tests including blood urea and serum creatinine were done for all. Additional investigations or diagnostic procedures (including radiological imaging, gastrointestinal (GI) endoscopy, tissue biopsy, stool analysis, thyroid function test, etc.) were performed when judged to be appropriate.

### Statistical analysis

We used Statistical Package for the Social Sciences (SPSS), version 16 for Windows, for data analysis, and recorded discrete data as frequency and percentage, and symmetrically distributed continuous data as mean  $\pm$  SD. The study subjects were categorized into three groups according to age (of age 60–69 years, 70–79 years, and 80 years and above). Similarly, study groups were categorized into different types of anemia by MCV values, such as normocytic, microcytic, and macrocytic anemia. Differences in the mean values were analyzed using the ANOVA test. Differences in the type of anemia by age groups, gender, and other variables were analyzed using Chi-square test.  $P < 0.05$  was considered statistically significant.

## Results

### Baseline characteristics

We enrolled 120 patients aged 60 years and above with anemia according to WHO definition. Fifteen patients were lost to follow-up and were excluded. The mean age was  $68.1 \pm 7.8$  years, ranging from 60 to 92 years. The male to female ratio was 1.6:1 and there was an almost equal distribution of rural (50.5%) and urban population (49.5%).

### Clinical and laboratory spectrum of anemia

Easy fatigability was the most common complaint, seen in 86.7% of patients followed by weakness (80%), shortness of breath (45.7%), swelling of feet (22.9%), headache (19%), vertigo (18.1%), palpitation (14.3%), bleeding (12.4%), and tinnitus (6.7%). Vertigo was more common in older patients aged 80 years and above and was found in 28.6% of cases of this age group.

On physical examination, pallor was the most common finding, noted in 84.8% of patients; peripheral edema (21.9%) and glossitis (9.5%) were next common findings.

The mean value of hemoglobin was  $8.8 \pm 2.3$  g/dl with a range of 3.9–13 g/dl. Older age group of 80 years and above had lower mean hemoglobin (7.5 g/dl).

Anemia characterization based on RBC indices and peripheral smear revealed that normocytic anemia was the most common type, seen in 53.3% of patients followed by microcytic anemia (40%) and macrocytic anemia (6.6%). 56.1% patients were pure vegetarians and microcytic anemia was more common in this group (45.8%). Chronic alcohol consumption was noted in 37.1% elderly patients. Normocytic anemia was present in 53.8% of all alcohol consumers, microcytic anemia in 35.9%, and macrocytic anemia in 10.3%.

### Etiological classification

Among the 105 patients, the etiological distribution of anemia was iron deficiency anemia (IDA) in 26 patients (24.8%), anemia in chronic disease (ACD) in 24 patients (22.9%), hematological disorders in 21 (20%), chronic kidney disease (CKD) in

13 (12.4%), mixed in 8 (7.6%), vitamin B12 deficiency in 2 (1.9%), folate deficiency in 1 (0.9%), and hypothyroidism in 1 patient (0.9%). No etiology for anemia could be found in nine (8.6%) patients [Table 1].

Among the 26 cases of IDA, upper GI lesions were observed in 15 patients (57.6%), which included chronic gastritis in 8 (30.7%), peptic ulcer disease with bleed in 3 (11.5%), esophagitis with hiatus hernia in 2 (7.7%), carcinoma esophagus in 1 (3.8%), and celiac disease in 1 (3.8%). Lower GI lesions were seen in four patients, i.e. adenocarcinoma, terminal ileal ulcers, and ulcer cecum. GI malignancy was found in two (7.7%) of IDA patients. Nutritional IDA was seen in five patients (30.7%).

In our study, ACD was found in 24 patients (22.8%). Common underlying chronic diseases were malignancy in 8 (36.6%) and

chronic liver disease in 7 (29.1%) patients. Other causes of chronic inflammation noticed were pulmonary tuberculosis, rheumatoid arthritis, reactive arthritis, left ventricular failure, angiodysplasia, pulmonary aspergillosis, and liver abscess.

In our study, 21 cases were due to hematological disorders. Out of these, 16 (76.2%) patients were having malignant hematological disorders including myelodysplastic syndrome (MDS) in 4 (19%), multiple myeloma in 2 (9.5%), chronic lymphoproliferative disease in 3 (14.2%), chronic myeloproliferative disease in 3 (14.2%), Non-Hodgkin disease in 3 (14.2%), and Hodgkin disease in 1 (4.7%). Five patients (23.8%) were having non-neoplastic hematological disorders which included beta-thalassemia trait in three (14.2%) and aplastic anemia in two (9.5%).

## Discussion

Using WHO criteria for anemia, 105 patients of age 60 years and above were included, and underwent complete clinical evaluation and laboratory investigations. Anemia was evaluated in a manner similar to that in younger adults. MCV was used to classify anemia into microcytic, normocytic, and macrocytic anemia. Normocytic anemia was the most common type in our study, seen in more than half of cases, similar to previous studies.<sup>[15,16]</sup>

Further evaluation for underlying etiology showed that IDA was the most prevalent anemia, just outnumbered the ACD (24.8% vs 22.9%) [Table 2].<sup>[3,17-21]</sup>

Upper GI endoscopy performed in patients with IDA with no obvious cause showed that chronic upper GI blood loss, including occult blood from chronic gastritis, peptic ulcer disease, esophagitis, and carcinoma esophagus constitute 53.8% of IDA cases. Insufficient dietary intake is still a common cause of iron deficiency in developing countries like India, and was seen in 19.2% of our IDA cases.<sup>[22]</sup>

In older adults, inflammatory conditions are frequently linked with anemia. A recent study found inflammatory disorders in 70% cases of anemia in 191 hospitalized elderly patients, of which 60% had concomitant CKD.<sup>[23]</sup> Our study found ACD in 22.9% cases and CKD in another 12.4% cases of anemia. In ACD, pro-inflammatory cytokines suppress erythropoiesis and shorten RBC survival.<sup>[24]</sup> In older age, subtle pro-inflammatory changes may lead to clinically significant anemia.<sup>[25]</sup>

**Table 1: Etiological classification of anemia in the 105 older patients**

Causes	Number (%)
Iron deficiency anemia	26 (24.8%)
Chronic gastritis	8
Upper gastrointestinal ulcer	3
Nutritional	5
Gastrointestinal malignancy	2
Others	8
Anemia of chronic disease	24 (22.9%)
Solid tumor malignancy	8
Chronic liver disease	7
Chronic infection	4
Chronic inflammatory condition	2
Others	3
Chronic kidney disease	13 (12.4%)
Hematological disorder	21 (20%)
Myelodysplastic syndrome	4
Multiple myeloma	2
Chronic lymphoproliferative disease	3
Chronic myeloproliferative disease	3
Non-Hodgkin lymphoma	3
Hodgkin Lymphoma	1
Aplastic anemia	2
Beta thalassemia	3
Folate/vitamin B12 deficiency	3 (2.8%)
Hypothyroidism	1 (0.9%)
Multi-factorial cause	8 (7.6%)
Unexplained cause	9 (8.6%)

**Table 2: Major causes of anemia in geriatrics: Comparison of different studies**

	Guralnik et al. 2004	Tettamanti et al. 2010	Bhasin et al. 2011	Artz et al. 2011	Petrosyan et al. 2012	Our study 2015
Anemia of chronic disease	33.6	17	27	10	62.1	22.9
Iron deficiency anemia	16	16	30	25	30.5	24.8
Chronic kidney disease	12	15	21	3.5	17.8	12.4
Vitamin B12/folate deficiency	14.3	10	5	3.4	38.9	2.9
Hematological malignancy	NA	7.4	6	NA	6.3	15.2
Unexplained	24	26	2	44	8.4	8.6

Note: NA – not available

Benign hematological causes of anemia like hemolytic anemia, thalassemia, aplastic anemia, etc. are seen more frequently in younger age group. Conversely, malignant hematological disorders like MDS, myeloma, chronic lymphoproliferative disorder, chronic myeloproliferative disorder, and lymphoma are common in older age. Our sample included 15.2% of elderly patients with malignant hematological disorders and anemia, a higher prevalence than previous studies. MDS should be a diagnostic consideration when other cell lineages are involved. Multiple myeloma is also an important cause of anemia in the elderly. A high index of suspicion is required for this possibility when anemia is associated with back pain, hypercalcemia, or renal failure.

In our study, 7.6% of elderly patients had more than one cause for their anemia. Multi-factorial etiology is common among elderly, and some studies found more than one cause of anemia in more than half of the cases.<sup>[17]</sup> Alcohol consumption and medications are important contributors to anemia in this age group. In our study, 37.1% of patients were found to be chronic alcohol consumers.

The cause of anemia remained unexplained only in 8.6% of cases. Previous studies found no cause of anemia in at least one-fourth of elderly patients.<sup>[3,17,20]</sup> Postulated underlying mechanisms for unexplained anemia in older adults are early myelodysplasia, malnutrition, hypogonadism, low glomerular filtration rate and relative erythropoietin deficiency.<sup>[26]</sup>

The limitation of our study was a hospital-based enrolment of cases. Asymptomatic patients or patients with mild anemia may not have reported to the health center. Further population-based studies are needed to ascertain the prevalence and etiology of anemia in the older population of India and to elucidate the impact of disease on this age group.

## Conclusion

Given the rising numbers of older adults in India, family physicians require greater attention to evaluate and manage common treatable conditions such as anemia, which may lead to increased morbidity and mortality in this particular population. This prospective study shows that in most of the cases, anemia in the elderly has a treatable cause. Thus, a thorough investigation including GI endoscopy is warranted. Unexplained progressive or unresponsive anemia requires bone marrow examination.

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Nil.

## Conflicts of interest

There are no conflicts of interest.

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