

# A Study of Severe Anemia in Elderly with Special Reference to Etiology

Dr. Antarikhya Bordoloi, Jr. Resident, 1st Year<sup>#1</sup>, Dr. Sayali Suhas Damle, Jr. Resident, 3<sup>rd</sup> Year<sup>\*2</sup>,

Professor (Dr.) Tushar Kanti Biswas, MD, Professor & Head <sup>#3</sup>

<sup>#</sup> Dept. of Geriatrics, MGM Medical College & Hospital, MGM

University of Health Sciences (Deemed University), Sector I,

Kamothe, Navi Mumbai – 410209, Maharashtra, INDIA

**Abstract** - This study was a prospective, observational study conducted over two years in a tertiary care hospital in Navi Mumbai, India, to study severe Anemia in the elderly with a special reference to etiology. Owing to India's increasing elderly population, Anemia in the elderly is a common condition that is vastly overlooked. The study included 50 individuals with severe Anemia (Hb less than 8g/dL), with an age more than or equal to 60 years. Our study's age varied from 60 to 82 years, with maximum patients from 60 to 65. Easy fatigability was present in 80% of patients, followed by anorexia, tiredness, breathlessness, and pedal edema. Generalized weakness, black colored stools, jaundice, headache, dysphagia, intermittent claudication, giddiness, and lower limb weakness and paresthesia were some other symptoms experienced by these patients. The most common type of Anemia among elderly patients, as found through this study, was microcytic hypochromic, followed by normocytic normochromic, dimorphic, and macrocytic hyperchromic Anemia. Severe Anemia in the elderly had varied manifestations.

**Keywords** — Anemia, Atypical, Elderly, Etiology, Haemoglobin, Fatiguability

## I. INTRODUCTION

Anemia is defined as a hemoglobin concentration of less than 12g/dl in women and less than 13g/dl in men according to world health organization criteria. Severe Anemia in the elderly is defined as a hemoglobin concentration of less than 8 mg/dl in both women and men. The most common causes of Anemia include nutritional deficiencies (iron, vitamin B12, folate), chronic diseases, hematopoietic malignancies (Myelodysplastic syndrome), bleeding disorders, various blood disorders (Thalassemia, Sickle cell disease), autoimmune hemolysis, etc. Anemia in the elderly should not be ignored as it can disguise various underlying conditions.

Diagnosis of Anemia is made initially with complete blood count and peripheral smear examination. It is then used to determine the causative factors, serum iron studies, serum vitamin B12 levels, reticulocyte count, LDH, Liver function tests, and Renal function tests are done. If these tests are inconclusive, then a bone marrow examination can be done.

Anemia affects the quality of the life of the patient as well as the caregiver as it causes various health hazards. This study contemplates a review of clinical manifestations and causative factors of severe Anemia in the elderly.

## II. MATERIALS AND METHODS

The study was a prospective observational study in which the data was collected from elderly patients (age ≥ 60 yrs) detected having severe Anemia (both inpatient and outpatient) attending MGM Hospital for treatment from May 2018 to December 2019 in the Department of Geriatrics, MGM Medical College Hospital, Navi Mumbai.

The detailed history of symptoms such as easy fatigue and loss of energy, unusually rapid heartbeat with exercise, shortness of breath, headache, dizziness, concentration difficulty, pale skin, leg cramps, insomnia, and thorough physical examination, particularly the gastrointestinal system. All previous history of hospital admissions and investigations were recorded. Investigations done were complete blood count, serum iron studies, serum vitamin B12 levels, P.S for cell type, reticulocyte count, stool for occult blood, urine routine & microscopy, USG abdomen & pelvis, bone marrow aspiration & biopsy.

## III. OBSERVATION & RESULTS

The study included 50 elderly patients of Anemia with age equal to or more than 60 years, ranging from 60 to 84. Easy fatigability was the commonest symptom, present in 40 (80%) patients followed by anorexia in 30 (60%) patients, tiredness in 23 (46%) patients, breathlessness and pedal edema in 21 (42% each) patients, generalized weakness in 18 (36%), lightheadedness and jaundice in 5 (10% each), black colored stools, headache and dysphagia 4 (8% each) patients, claudication 3 (6%) patients, bleeding only in 1 (2%) patients. The associated morbidities present were Diabetes mellitus in 16 (32%) patients, Hypertension in 23 (46%) patients, ESRD in 7 (14%) patients, CVA in 5 (10%) patients. Out of the total, 50 patients, 18 (36%) patients had a history of frequent alcohol consumption, and 7 (14%) patients were using anticoagulants for different conditions. The most common type of Anemia seen in peripheral smear among elderly patients was found to be microcytic hypochromic (56%), followed by normocytic normochromic



(24%), dimorphic, and macrocytic hyperchromic Anemia (10% each). 8 (16%) patients had positive stool occult blood, and pallor was seen in 48(96%) patients. Bone marrow examination was performed in 14 (28%) patients and was found to be microplastic in 12%, megaloblastic with erythroid hyperplasia in 8%, marked erythroid hyperplasia in 4% patients. Hypercellular marrow in 2% and severely hypocellular marrow was seen in 2% of patients. On ultrasonography, hepatomegaly was seen in 10, splenomegaly in 8%, and hepatosplenomegaly in 6% of patients. The majority of the patients (30%) had 6-7 gm% Hb, followed by 26% had 5-6 gm % Hb, and 20% had more than 7gm%, 16% had 4-5gm% Hb, and 8% had less than 4gm%Hb. In the study, serum iron levels were <50µg/dl in 60%, 50-100µg/dl in 10%, 100-150µg/dl in 18%,150-200µg/dl in 4% and >250µg/dl in 8% patients. Thus, most elderly patients with severe Anemia had severe iron deficiency, i.e., serum iron levels <50µg/dl. The serum ferritin concentration was <100ng/ml in 60%, 200-300ng/ml in 22%, >400ng/ml in 8%, 100-200ng/ml in 6%and between 300-400ng/ml in 4% patients. Vitamin B 12 level was found to be <500pg/ml in 80% patients. Reticulocyte count was <1 in 42% and 1-2 in 38% of patients with more than 4 in only 4%.

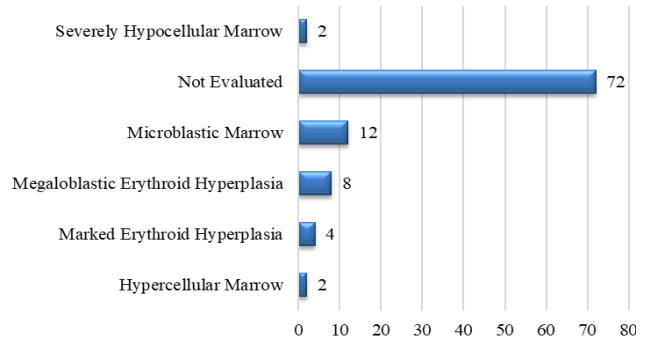


Fig. III Bone marrow examination & findings

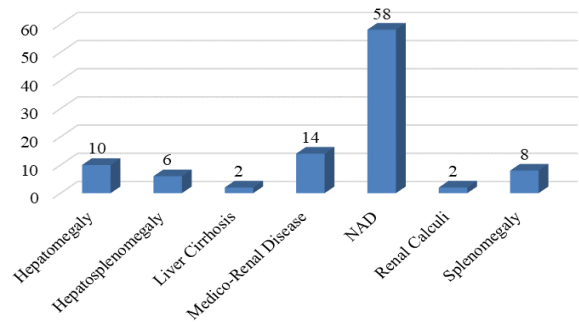


Fig. IV Findings of USG Abdomen & Pelvis

IV. FIGURES

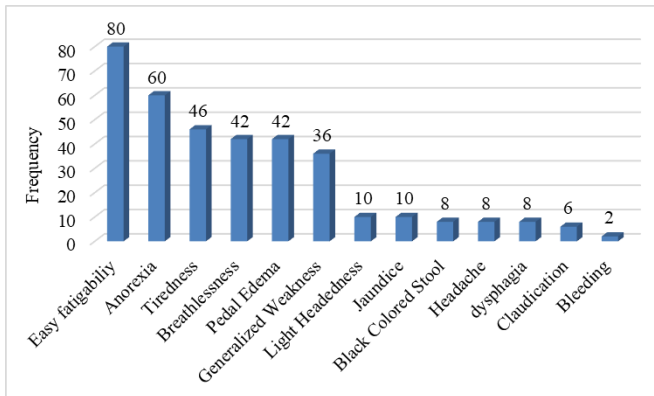


Fig. I Symptoms and their frequencies noticed during the study

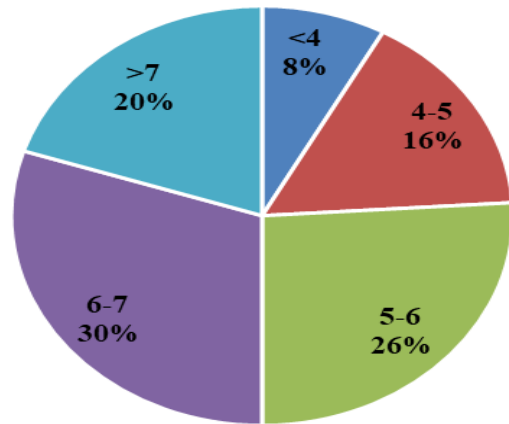


Fig. V Distribution of no. of patients with various levels of hemoglobin

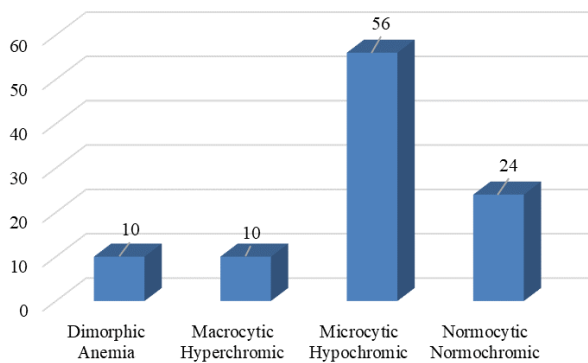


Fig. II Types of Anemia and their distribution

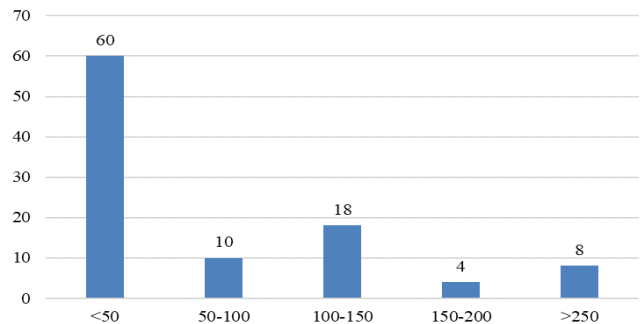


Fig. VI Distribution of levels of Serum Iron among patients with anemia

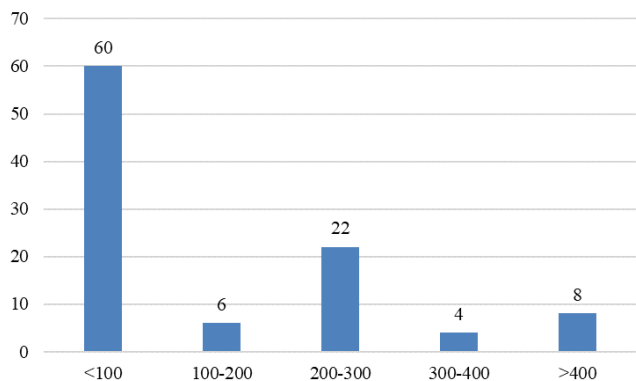


Fig. VII Distribution of levels of Serum Ferritin among patients with anemia

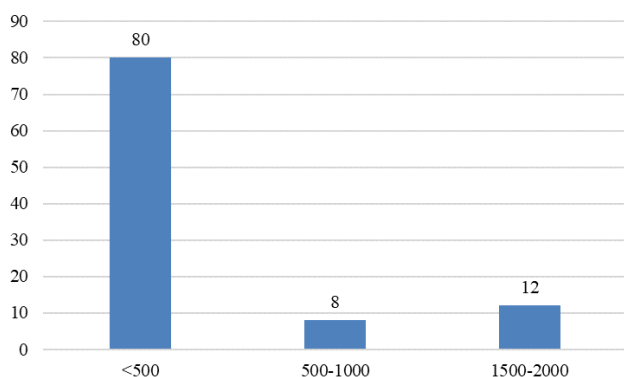


Fig. VIII Fig. VIII Distribution of levels of Vit B12 among patients with anemia

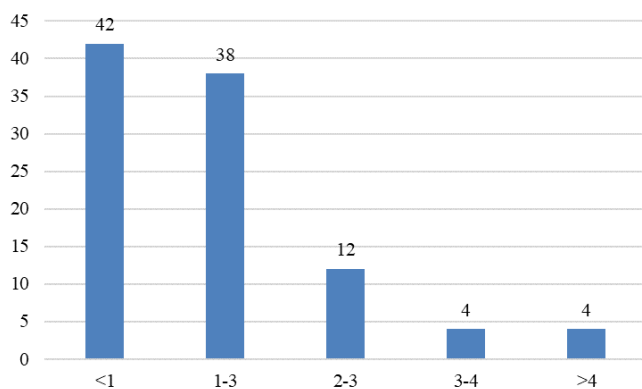


Fig. IX Distribution of levels of Retic Count among patients with Anemia

### V. DISCUSSIONS

In this prospective observational study, elderly patients of the age  $\geq 60$  years (both inpatient and outpatient) detected having severe Anemia (hemoglobin less than 8 g/dl) were studied for clinical signs and symptoms, and etiologies delineate strategies for treatment and prevention of severe Anemia.

In the study, 38% were between 60-65 years, followed by 32% between 65 and 70, with gender-wise distribution of 50% males and 50% females. Among the various clinical

features in the study, easy fatigability was present in 80%, followed by anorexia in 60%, tiredness in 46%, breathlessness & pedal edema in 42% each, generalized weakness in 36%, lightheadedness & jaundice in 10% each, black-colored stools headache & dysphagia in 8% each, claudication in 6%, bleeding 2% were also seen. The tissue hypoxia, which occurs due to the decreased oxygen-carrying capacity, is responsible for the clinical signs and symptoms. In 78% of the patients, no other non-specific symptoms of severe Anemia were observed. In the study, 32% of patients had diabetes mellitus, 46% had hypertension, and 14% had ESRD, 10% presented with CVA. In the study, 36% of the patients had a history of frequent alcohol consumption. Excessive alcohol consumption causes bone marrow suppression, which ultimately affects all three lineage precursors. It indirectly causes nutritional deficiencies, mainly vitamin B12 deficiency. In the study, 96% had pallor with a peripheral smear showing 56% microcytic hypochromic Anemia, followed by 24% normocytic normochromic anemia, 10% each of dimorphic & macrocytic hyperchromic Anemia. Stool occult blood allows for an increase in early-stage cancer detection showed 16% positivity in the study. Bone marrow examination is the most reliable test for diagnosing Anemia. Diseases that show hypoplasia and marrow infiltration are demonstrated well in bone marrow aspiration and biopsy. It should be considered in unclear macrocytic Anemia, especially when additional cytopenias are seen and other possible causes of leukopenia or thrombocytopenia have been excluded. In 28% of Of patients, bone marrow examination showed microplastics in 12%, megaloblastic with erythroid hyperplasia in 8%, and marked erythroid hyperplasia in 4% of patients. The abdomen's ultrasonography showed hepatomegaly in 10%, splenomegaly in 8% & hepatosplenomegaly in 6% of patients. Liver cirrhosis was seen in 2%, medico-renal disease in 14%, and renal calculi in 2% of patients. In the study majority of patients(30%) had 6-7 gm% Hb with serum iron levels  $<50\mu\text{g/dl}$  in 60% , vitamin B12  $<500\text{pg/ml}$  in 80% and reticulocyte count  $<1$  in 42% & 1-2 in 38% of patients.

### VI. CONCLUSIONS

Anemia is seen commonly in the elderly with varied etiologies presenting with vast clinical manifestations. Therefore, early detection and thorough workup are necessary to prevent complications and diagnose the underlying conditions.

Among the elderly population, iron deficiency anemia is very common, although vitamin B12 deficiency is rare. Nutritional deficiencies contribute to the various presenting symptoms such as easy fatigability, anorexia, tiredness, breathlessness, and pedal oedema, which remain the most common symptoms. Anemia can have atypical presentations with giddiness, lower limb weakness in elderly populations. Frank or occult bleeding through the gastrointestinal tract is a serious etiological factor in which exclusion of underlying

gastrointestinal malignancy is important.

Detailed personal history about nutrition, substance abuse, drugs, bowel characteristics, and bleeding manifestations should be taken.

Bone marrow examination is not necessary for all patients due to the presence of specialized blood tests.

The study established the etiological factors and the presenting symptoms in the elderly population by including the laboratory investigations and the different blood parameters.

#### ACKNOWLEDGMENT

It had been a sheer privilege to have worked under my teacher's endearing guidance, mentor, and guide Dr. T K Biswas, MD, General Medicine, Professor and Head, Department of Geriatrics, MGM Medical College and Hospital, Navi Mumbai and the co-author for this study. I also express my sincere gratitude and appreciation to Dr. Sayali Damle, Jr. Resident, 3rd Year, Dept. of Geriatrics, MGM Medical College & Hospital, and my co-author for this study for her immense contribution and continued guidance for completing this study on time successfully. I also would like to thank all the patients and their attendants who gave consent for their participation in this study, without whom this study would not have been possible.

#### REFERENCES

- [1] World Health Organization. *Deficiency Anemia: Assessment, Prevention, and Control*. Geneva. World Health Organization; 200.
- [2] Haemoglobin Concentration for the Diagnosis of Anemia and Assessment of Severity, the World Health Organization
- [3] Red Blood Cells and Bleeding Disorders, Robbins Basic Pathology, 8th Edition, Chapter 14, 659-662
- [4] Nancy C. Andrews, Iron Deficiency and Related Disorders, Wintrob's Clinical Haematology, 12th Edition, Chapter 27, 812.
- [5] Green R, Charlton R, Seftel H, et al., Body Iron Excretion in Men: A Collaborative Study, *AM J Med*, 1968; 45:336-353
- [6] Hallberg L, Hogdahl AM, Nilsson L, Rybo G, Menstrual Blood Loss – A Population Study, Variation at Different Ages and Attempts to Define Normality. *Acta Obstet Gynecol Scand*, 1966 ; 45 : 320 – 351.
- [7] Hallberg L, Rossander-Hulten L, Iron Requirements in Menstruating Women, *Am J Clin Nutr* 1991; 54:1047 – 1058
- [8] Beutler E, Iron Therapy in Chronically Fatigued Non-Anemic Women, *Ann Intern Med* 1960; 52: 378.
- [9] Robert T, Means Jr., Bertil Glader, Anemia General Considerations, Wintrob's Clinical Haematology, 12th Edition, Chapter 26, 787
- [10] Herbert B, Experimental Nutritional Folate Deficiency in Men, *Trans Assoc AM Physicians* 1962; 75: 307 – 320.
- [11] Nath BJ, Lindenbaum J, Persistence of Neutrophil Hyper segmentation During Recovery from Megaloblastic Granulopoiesis, *Ann Intern Med* 1979; 90 (5): 757 – 760.
- [12] Lindenbaum J, Heaton EB, Savage DG, et al., Neuropsychiatric Disorders Caused by Cobalamine Deficiency in the Absence of Anemia or Macrocytosis, *N Engl J Med* 1988; 318: 1720 – 1728.
- [13] Carmel R, Pernicious Anemia: The Expected Findings of Very Low Serum Cobalamine Levels, Anemia and Macrocytosis are often Lacking, *Arch Intern Med* 1988; 148: 1712 – 1714.
- [14] Heaton EB, Savage DG, Brust JC, et al., Neurologic Aspects of Cobalamine Deficiency, *Med* 1991; 70: 229 – 245.
- [15] Savage D, Lindenbaum J, Relapses After Interruption of Cyanocobalamine Therapy in Patients with Pernicious Anemia, *Am J Med* 1983; 74: 765 – 772.
- [16] Magnus EN, Cobalamine and Unsaturated Trans Cobalamine Values in Pernicious Anemia: Relation to Treatment, *Scand J Haematol* 1986; 36: 457 – 465.
- [17] Crosby W, Akeroyd JH, The Limit of Haemoglobin Synthesis in Hereditary Anemia, *Am J Med* 1952; 13: 273 – 283.
- [18] Ham T, Hemoglobinuria, *Am J Med* 1955; 18: 990 – 1006.
- [19] Dawson AA, Ojston D, Fullerton HW, Evaluation of Diagnostic Significance of Certain Symptoms and Physical Signs in Anemic Patients, *BMJ* 1969; 3 (668): 436 – 439.
- [20] Bartels E, Anemia as the Cause of Severe Congestive Heart Failure: Report of a Case, *Annals of Internal Medicine* 1937; 11: 400 – 404.
- [21] Pickering G, Wayne EJ, Observations on Angina Pectoris and Intermittent Claudication in Anemia, *Clinical Science* 1933; 1: 305 – 325.
- [22] Varat MA, Adolf RJ, Fowler NO, Cardiovascular Effects of Anemia, *American Heart Journal* 1972; 83 (3): 415 – 426.
- [23] Marshall R, A Review of Lesions on the Optic Fundus in Various Diseases of the Blood, *Blood* 1959; 14: 882 – 891.
- [24] Aisen ML, et al., Retinal Abnormalities Associated with Anemia, *American Academy of Ophthalmology* 1983; 101 (7): 1049 – 1052.
- [25] Merin S, Freund M, Retinopathy in Severe Anemia, *American Journal of Ophthalmology* 1968; 66 (6): 1102 – 1106.
- [26] Schaber J, Blumberg AG, Papilledema Associated with Blood Loss Anemia, *Annals of Internal Medicine* 1961; 55: 1004 – 1007.
- [27] Robert T, Means Jr., Bertil Glader, Anemia General Considerations, Wintrob's Clinical Haematology, 12th Edition, Chapter 26, 801.
- [28] Consensus Conference, Peri-operative Red Blood Cell Transfusion, *JAMA* 1988; 260 (18): 2700 – 2703.
- [29] Practice Strategies for Elective Red Blood Cell Transfusion, American College of Physicians, *Ann Intern Med* 1992; 116 (5): 403 – 406.
- [30] Practice Guidelines for Blood Component Therapy: A Report by the American Society of Anaesthesiology 1996; 84 (3): 732 – 747.
- [31] Expert Working Group: Guidelines for Red Blood Cell and Plasma Transfusion for Adults and Children, *CAN Med Assoc. J* 1997 (11 Suppl.); S1 – S24.
- [32] Welch HG, Meehan KR, Goodnough LT, Prudent Strategies for Elective Red Blood Cell Transfusion, *Ann Intern Med* 1992; 116 (5): 393 – 402.
- [33] Hebert PC, Wells G, Blajchman MA, et al., A Multicenter, Randomized, Controlled Clinical Trial of Transfusion Requirements in Critical Care, Transfusions Requirements in Critical Care Investigators, Canadian Critical Care Trials Groups, *N Engl. J Med* 1999; 340 (6): 409 – 417.
- [34] Vincent JL, Baron JF, Reinhart K, et al. Anemia and Blood Transfusion in Critically Ill Patients, *JAMA* 2002; 288 (12): 1499-1507.
- [35] Wu WC, Rathore SS, Wang Y, et al. Blood Transfusion in elderly patients with acute myocardial infarction, *N Engl. J Med* 2001; 345 (17): 1230 – 1236.
- [36] Ross S, Jeter E. Emergency Surgery: Trauma and Massive Transfusion. In: Petz LD, Kleinman S, Swisher SN, et al., eds. *Clinical Practice of Transfusion Medicine*, 3rd ed. New York: Churchill Livingstone, 1996: 563 – 579.
- [37] Carson JL, Duff A, Berlin JA, et al. Perioperative Blood Transfusion and Postoperative Mortality, *JAMA* 1998; 279 (3): 199 – 205.
- [38] Groopman J.E., Itri KM, Chemotherapy-induced Anemia in Adults: Incidence and Treatment, *J Natl Cancer Inst* 1999; 91 (19):1616 – 1634
- [39] Petz LD, Blood Transfusion in Acquired Hemolytic Anemias, In Petz LD, Kleinman S, Swisher SN, et al., eds. *Clinical Practice of Transfusion Medicine*, 3rd ed. New York: Churchill Livingstone, 1996: 469 – 499.
- [40] Petz LD, Garratty G, Blood Transfusion in Autoimmune Hemolytic Anemia. In: *Immune Hemolytic Anemias*, 2nd ed. Philadelphia: Churchill Livingstone, 2004: 375 – 400.
- [41] Grebe G, Martizez-Torres C, Layrisse M, Effect of Meals and Ascorbic Acid on the Absorption of a Therapeutic Dose of Iron as Ferrous and Ferric Salts, *Curr Ther Res Clin Exp* 1975; 17: 382 – 397.
- [42] Crosby WH, The Rationale for Treating Iron Deficiency Anemia. *Arch Intern Med* 1984; 144: 471 – 472.
- [43] Cacciola E, Witt D, Toler W, et al., Ascorbic Acid Deficiency may be a Cause of Refractoriness to Iron-Therapy in the Treatment of Iron-Deficiency Anemia, *Haematologica* 1994; 79: 96-97.
- [44] Nath BJ, Lindenbaum J, Persistence of Neutrophil Hyper-segmentation during Recovery from Megaloblastic Granulopoiesis, *Ann Intern Med* 1979; 90: 757 – 760
- [45] Marshall A Litchman, Thomas J Kipps, Folate, Cobalamine and Megaloblastic Anemias, Williams Hematology 8th edition (2010), Chapter 41.

- [46] Temple RM, Deary IJ, Winney RJ, Recombinant Erythropoietin Improves Cognitive Function in Patients Maintained in Chronic Ambulatory Peritoneal Dialysis, *Nephrol Dial Transplant* 1995; 10: 1733 – 1738.
- [47] Mayer G, Thum J, Cada Em, et al., Working Capacity is increased following recombinant human erythropoietin treatment, *Kind Int.* 1988; 34: 525 – 528.
- [48] Lago M, Perez-Garcia R, De VMSG, et al., Efficiency of Once-Weekly Subcutaneous Administration of Recombinant Human Erythropoietin Versus Three times a Week Administration in Hemodialysis Patients, *Nephron* 1996; 72: 723 – 724.
- [49] Eschbach JW, Abdulhadi MH, Browne JK, et al., Recombinant Human Erythropoietin in Anemic Patients with End-Stage Renal Disease: Results of a Phase III Multicenter Trial, *Ann Intern Med* 1989; 111: 992 – 1000.
- [50] Jones MA, Kingswood HC, Dallyn PE, et al., Changes in diurnal blood pressure variation and red cell and plasma volumes in patients with renal failure who develop erythropoietin-induced hypertension, *Clin Nephrol* 1995; 44: 193 – 200.