Journal of Research in Applied and Basic Medical Sciences 2020; 6(4): 272-278





RABNS Journal of Research in Applied and Basic Medical Sciences



Relationship between Peripheral Blood Eosinophil Count and Neutrophil/Lymphocyte Ratio with Number of Attacks Leading to **Hospital Admission in Chronic Obstructive Pulmonary Disease**

Maryam Mirzaee¹, Mohammad Amin Abbasi², Abbas Fadaii^{1*}

¹ Department of Pulmonology and Intensive Care Medicine, Shahid Labbafinejad Hospital, Shahid Beheshti University of Medical Sciences, Tehran, Iran

² Firoozabadi Clinical Research Development Unit (FCRDU), Iran University of Medical Sciences (IUMS), Tehran, Iran

*Corresponding authors: Abbas Fadaii, Address: Department of Pulmonology and Intensive Care Medicine, Shahid Labbafinejad Hospital, Shahid Beheshti University of Medical Sciences, Tehran, Iran, Email: abbasfadaii@gmail.com Tel: +982151048000

Abstract

Background & Aims: Chronic obstructive pulmonary disease is an important common respiratory disorder. Determination of the prognostic factors is important to improve the outcomes and decrease the burden of problem. We aimed to determine the relationship between peripheral blood eosinophil count and neutrophil/lymphocyte ratio (NLR) with number of attacks leading to hospital admission in patients with chronic obstructive pulmonary disease.

Materials & Methods: In this cohort study, 200 consecutive patients with chronic obstructive pulmonary disease from 2016 to 2018 were enrolled including those with and without attacks leading to hospital admission and eosinophil and NLR were compared in them. Results: The results in this study demonstrated that mean eosinophil was 1.5 and 1.4 in those without and with attack without significant difference (p=0.641). The mean NLR was 4.6 and 5.9 in those without and with attack with significant difference (p=0.022).

Conclusion: Totally, according to the obtained results, it is concluded that higher NLR is related to further attacks leading to hospital admissions but the eosinophil count has no significant effect in this area.

Keywords: Eosinophil, NLR, COPD, Exacerbation

Received 28 August 2020; accepted for publication 26 November 2020

Introduction

Chronic obstructive pulmonary disease (COPD) has prevalence rate of 0.2 to 37 percent worldwide with male and early predominance leading to mortality rate of 3 to 111 cases per 100 000 subjects annually (1). It is different by race and geographic distribution (2, 3). Determination of contributing factors for disease severity in COPD cases is important to decrease the burden of disease (4, 5). Global Institute for chronic obstructive lung disease has defined COPD as partially reversible airway obstruction with progressive pattern and abnormal pulmonary pattern against dangerous gas and particles (6, 7). It is the fourth cause of death after

cardiovascular disease, cancer, and vascular disorders (8). It is the fifth disabling disease worldwide (7).

Exacerbations of this disease are related to worse quality of life and increased morbidity and mortality (8, 9). During exacerbations, there is inflammation and three percent increase of neutrophils in airways (10, 11). Corticosteroids are used for treatment of exacerbations (12) and are especially effective in cases with higher inflammation and eosiniphil percentage (13-15). Neutrophils are introduced as novel prognostic markers and predictive for disease severity and exacerbations (16). Determination of the prognostic factors is important to improve the outcomes and decrease the burden of problem. Accordingly, we aimed to determine the relationship between peripheral blood eosinophil count and neutrophil/lymphocyte ratio with the number of attacks leading to hospital admission in patients with COPD.

Methods

In this cohort study, 200 consecutive patients with COPD from 2016 to 2018 in Labafinejad Hospital, Tehran, Iran were enrolled. Inclusion criteria were age range from 20 to 100 years, FEV1/FVC ratio less than 0.7, and minimal 10 pack/year smoking history. Exclusion criteria were systemic corticosteroid use in the last three months and over a 12% increase in FEV1 after bronchodilator use.

Clinical and laboratory assessments:

The study was approved by the ethical committee in Shahid Beheshti University of Medical Sciences, **Table 1.** Demographic Data of the Study Participants (n = 200)

Tehran, Iran including those with and without severe attacks leading to hospital admission and eosinophil and NLR were compared in them. Severe attacks were defined as those needing admission and medium grade was considered as cases that only received corticosteroids with/without antibiotic. Data were gathered from medical documents and recorded in the checklists.

Data analysis was done among 200 cases in two groups by SPSS version 16.0 software. The utilized tests were Kolmogorov-Smirnov, Independent-Sample-T, and ROC analysis. The p-value less than 0.05 was considered statistically significant.

Results

In the present study, 160 male and 40 female patients were enrolled. The mean was 50.23 ± 11.14 . Other demographic information can be found in Table 1.

The results in this study demonstrated that leukocyte count was same across the groups (Figure 1, p=0.507). As shown in Figure 2, the mean eosinophil was 1.5 and 1.4 in those without and with attack without a significant difference (p=0.641).

The mean neutrophil count was significantly (p=0.020) higher in exacerbations (Figure 3). The mean lymphocyte count was significantly (p=0.014) lower in exacerbations (Figure 4). The mean NLR was 4.6 and 5.9 in those without and with attack with significant difference (p=0.022). The AUC was 60% and the sensitivity and specificity were 60% by cut-off point of 3.8.

Characteristic	Total
Age, year	50.23±11.14
Men, n (%)	160 (80%)
Smoking, n (%)	200(100%)
GOLD ^a Stage 2 (%)	120 (60%)
GOLD Stage 3 (%)	80(40%)
Comorbidity	
HTN, n (%)	80(40%)
DM, n(%)	60(30%)
CHF, n (%)	50(43.5)
IHD, $n(\%)$	70 (35%)

a: GOLD: Global Initiative for Obstructive Lung Disease; HTN: hypertension DM: diabetes mellitus; CHF: chronic heart failure and IHD: ischemic heart renal disease

b: Variables are expressed as mean (standard deviation) and categorical data are expressed as number (percentage)

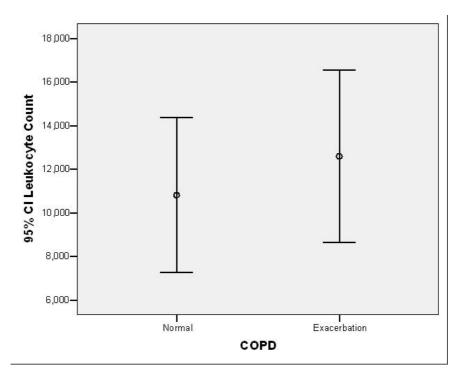


Figure 1: Leukocyte count according to exacerbation

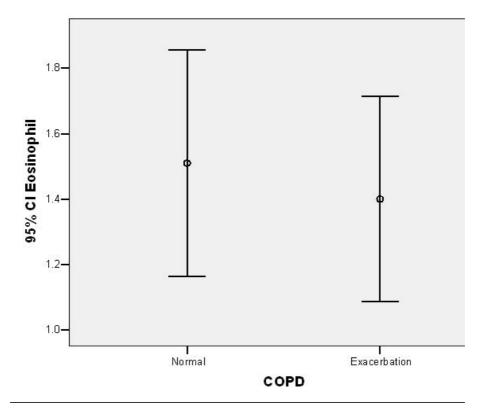


Figure 2: Eosinophil count according to exacerbation

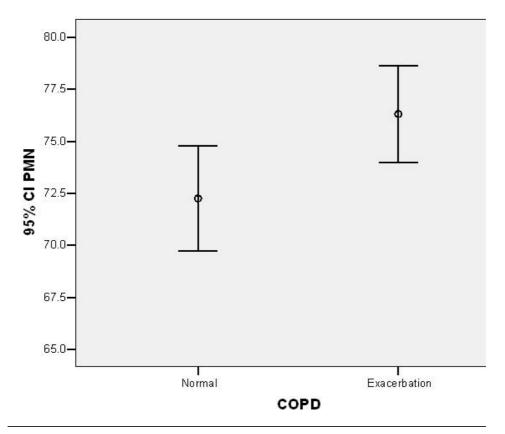


Figure 3. Neutrophil count according to exacerbation

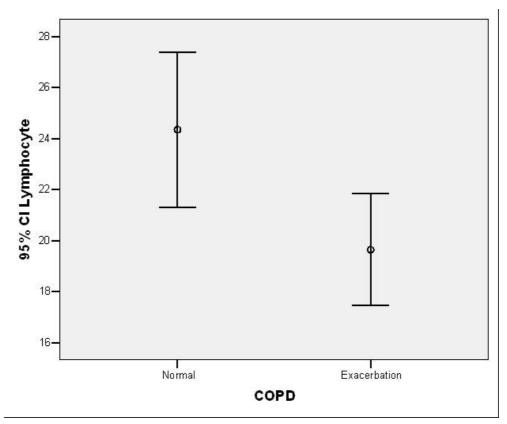


Figure 4. Lymphocyte count according to exacerbation

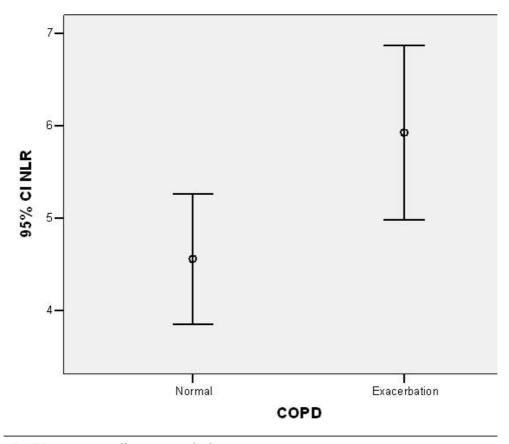


Figure 5. NLR count according to exacerbation

Discussion

This study was done to determine the importance and value of CBC factors especially eosinophil and NLR in prediction of exacerbations. In our study the eosinophil was same between those with and without exacerbations but significantly higher NLR was seen in acute exacerbations. In the current study, the sensitivity and specificity were 60% with cut-off point of 3.8.

Krogh et al. (15) reported that eosinophil over 3.4 was related to exacerbations and the amount was 1.15 and 1.8 for increased risk in severe and medium attacks. The difference in our study was 0.1 that showed no difference. Song et al. (17) assessed 467 cases and found that there is no significant association between exacerbation and eosinophil as well as FEV1. It was in congruence with our study.

Pascoe et al. (18) assessed the possible role of eosinophil for prediction of therapeutic response to inhalational corticosteroids and there was significant role. However, the treatment response was not assessed in our study but may be assessed in future studies. Ye et al. (19) similarly reported that higher NLR is related to higher severity of COPD with significant prognostic role. Also in their study the mortality rate was related to NLR.

Teng et al. (20) assessed 698 cases with COPD and found that AUC was 74 percent with sensitivity and specificity of 61 and 75 percent. The sensitivity was same in our study but the AUC and specificity were lower in our study that may be due to the smaller sample population. Liu et al. (21) reported AUC of 74 percent for NLR with cut-off point of 4.2 and the specificity and sensitivity were 71 and 74 percent, respectively. The cut-off point was near 3.8 points and the sensitivity and specificity were lower in our study. El-Gazzar (22) reported significantly higher NLR in exacerbations that is in line with our findings.

Conclusion

Totally, according to the obtained results, it is concluded that higher NLR is related to further attacks leading to hospital admissions but the eosinophil count has no significant effect in this area. However further studies with a larger sample size and multi-center sampling are required to attain more definite results. Also, evaluation of the possible role of the other hematological factors is recommended for COPD exacerbations.

Conflict of interest

The authors have no conflict of interest.

References

- Rycroft CE, Heyes A, Lanza L, Becker K. Epidemiology of chronic obstructive pulmonary disease: a literature review. Int J Chron Obstruct Pulmon Dis 2012; 7:457-94.
- Mannino DM, Buist AS. Global burden of COPD: risk factors, prevalence, and future trends. Lancet 2007; 370(9589):765-73.
- Martin A, Badrick E, Mathur R, Hull S. Effect of ethnicity on the prevalence, severity, and management of COPD in general practice. Br J Gen Pract 2012;62(595):e76-81.
- 4. Divo M, Cote C, de Torres JP, Casanova C, Marin JM, Pinto-Plata V, et al. Comorbidities and risk of mortality in patients with chronic obstructive pulmonary disease. Am J Respir Crit Care Med 2012;186(2):155-61.
- Soriano JB, Zielinski J, Price D. Screening for and early detection of chronic obstructive pulmonary disease. Lancet 2009;374(9691):721-32.
- Lozano R, Naghavi M, Foreman K, Lim S, Shibuya K, Aboyans V, et al. Global and regional mortality from 235 causes of death for 20 age groups in 1990 and 2010: a systematic analysis for the Global Burden of Disease Study 2010. Lancet 2012;380:2095–2128.
- Kasper D, Braunwald E, Hauser S, Longo D, Larry Jameson J, Fauci A. Harrison's Principle of Internal Medicine. 16th ed. McGraw- Hill, 2005. P. 1547-51.
- Steven D, Shapiro MD, Gordon L, Snider MD, Stephen I, Rennard M. Chronic Brobchitis and Emphysema. In: Murray JF, Nadel J. Text book of respiratory medicine. 4th ed. Philadelphia: Elsevier Saunders; 2005.P. 1116-27.
- Seemungal TA, Donaldson GC, Paul EA, Bestall JC, Jeffries DJ, Wedzicha JA. Effect of exacerbation on quality of life in patients with chronic obstructive

pulmonary disease. Am J Respir Crit Care Med 1998;157:1418–22.

- Saha S, Brightling CE. Eosinophilic airway inflammation in COPD. Int J Chron Obstruct Pulmon Dis 2006;1:39– 47.
- Bafadhel M, McKenna S, Terry S, Mistry V, Reid C, Haldar P, et al. Acute exacerbations of chronic obstructive pulmonary disease: identification of biologic clusters and their biomarkers. Am J Respir Crit Care Med 2011;184: 662–71.
- Bafadhel M, McKenna S, Terry S, Mistry V, Pancholi M, Venge P, et al. Blood eosinophils to direct corticosteroid treatment of exacerbations of chronic obstructive pulmonary disease: a randomized placebocontrolled trial. Am J Respir Crit Care Med 2012; 186:48–55.
- Pascoe S, Locantore N, Dransfield MT, Barnes NC, Pavord ID. Blood eosinophil counts, exacerbations, and response to the addition of inhaled fluticasone furoate to vilanterol in patients with chronic obstructive pulmonary disease: a secondary analysis of data from two parallel randomised controlled trials. Lancet Respir Med 2015; 3:435–42.
- Siddiqui SH, Guasconi A, Vestbo J, Jones P, Agusti A, Paggiaro P, et al. Blood eosinophils: a biomarker of response to extrafine beclomethasone/formoterol in chronic obstructive pulmonary disease. Am J Respir Crit Care Med 2015; 192:523–5.
- Vedel-Krogh S, Nielsen SF, Lange P, Vestbo J, Nordestgaard BG. Blood Eosinophils and Exacerbations in Chronic Obstructive Pulmonary Disease. The Copenhagen General Population Study. Am J Respir Crit Care Med. 2016;193(9):965-74.
- Çoban Ağca M, Aksoy E, Duman D, Özmen İ, Yıldırım E. Does eosinophilia and neutrophil to lymphocyte ratio affect hospital re-admission in cases of COPD exacerbation? Tuberk Toraks 2017;65(4):282-90.
- Song JH, Lee CH, Kim JW, Lee WY, Jung JY, Park JH, et al. Clinical implications of blood eosinophil count in patients with non-asthma-COPD overlapsyndrome COPD. Int J Chron Obstruct Pulmon Dis 2017; 12:2455-64.
- Pascoe S, Locantore N, Dransfield MT, Barnes NC, Pavord ID. Blood eosinophil counts, exacerbations, and

response to the addition of inhaled fluticasone furoate to vilanterol in patients with chronic obstructive pulmonary disease: a secondary analysisof data from two parallel randomised controlled trials. Lancet Respir Med 2015;3(6):435-42.

- Ye Z, Ai X, MD, Liao Z, You C, Cheng Y. The prognostic values of neutrophil to lymphocyte ratio for outcomes in chronic obstructive pulmonary disease. Medicine (Baltimore) 2019; 98(28): e16371.
- 20. Teng F, Ye H, Xue T. Predictive value of neutrophil to lymphocyte ratio in patients with acute exacerbation of

chronic obstructive pulmonary disease. PLoS One 2018; 13(9): e0204377.

- Liu J, Liu J, Zou Y. Relationship between neutrophil– lymphocyte ratio and short-term prognosis in the chronic obstructive pulmonary patients with acute exacerbationBiosci Rep 2019; 39(5): BSR20190675.
- El-Gazzar AG, Kamel MH, Elbahnasy OKM, El-Naggar ME. Prognostic value of platelet and neutrophil to lymphocyte ratio in COPD patients. Expert Rev Respir Med 2020;14(1):111-6.