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# Comparison of therapeutic response between asthma, COPD, and ACOS patients by evaluation of spirometric findings

Mohammad Esmaeil Hejazi<sup>1</sup>, Zahra Pakzad<sup>1\*</sup>, Horieh Shojaan<sup>1</sup>, Niusha Kalami<sup>1</sup>, Veghar Hejazi<sup>2</sup>, Tahereh Vaezi<sup>2</sup>

<sup>1</sup>Tuberculosis and Lung Diseases Research Center, Tabriz University of Medical Sciences, Tabriz, Iran <sup>2</sup>Faculty of Medicine, Mashhad University of Medical Sciences, Mashhad, Iran

\*Corresponding author: Zahra Pakzad, Tuberculosis and Lung Diseases Research Center, Tabriz University of Medical Sciences, Tabriz, Iran. Email: pakzadzahra585@gmail.com

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## ABSTRACT

Asthma and Chronic obstructive pulmonary disease (COPD ) both are a common public health problem that affects a large portion of population. Nearly 20% of patients with obstructive lung disease have features of both asthma and COPD called ACOS that GOLD\_GINA guidelines defines as persistent airflow limitation with several features of asthma and several features of COPD. Yet there is a little data available about diagnosis and treatment of this entity and current study aimed to compare therapeutic response between asthma, COPD and Asthma-COPD overlap syndrome (ACOS) subjects through spirometric data. In the present cross-sectional study, 30 known patients with mild to moderate asthma, 30 known patients with mild to moderate COPD and 30 known patients with mild to moderate ACOS according to GOLD\_GINA guidelines were enrolled. We assessed post bronchodilator the ratio of the forced expiratory volume in the first one second to the forced vital capacity of the lungs (fev1/fvc) in all patients. Then they took standard treatment for 2 months and after this period spirometry was repeated. Spirometric data's changes was

compared between the three groups by SPSS26 statistical software. Fev1 changes in response to treatment did not differ significantly between three groups (p > 0.05) but fev1/fvc changes differed significantly and this parameter in asthma was more than ACOS and in COPD was least. (In asthma, spirometric symbolized therapeutic response is more significant than ACOS, and in ACOS, it is more important than COPD in terms of fev1/fvc changes) and there was not any difference between the three groups regarding to FEV1 changes.

Keywords: Therapeutic response; Asthma; COPD; ACOS; spirometry

## INTRODUCTION

Asthma and COPD are prevalent chronic respiratory diseases driven by complex interactions between intrinsic and extrinsic factors that result in a wide spectrum of clinical presentations.<sup>1–4</sup> Despite clear differences in their aetiology and pathophysiology,<sup>5,6</sup> differentiating asthma and COPD at a diagnostic level is often challenging, particularly as analyses of patient spirometry data and inflammatory markers frequently identify obstructive airway disease phenotypes that encompass a mix of asthma and COPD features.<sup>7,8</sup> Presently, this patient group is referred to as patients with asthma-COPD overlap (ACO).<sup>9,10</sup> Individuals with ACO have increased disease severity,11 poorer quality of life12 and incur higher healthcare costs when compared with patients with asthma or COPD alone.13 GOLD GINA guidelines defines as persistent airflow limitation with several features of asthma and several features of COPD. Considering the defect of enough data about its mechanism and phenotypes exact definition is impossible actually.<sup>14</sup> Asthma is considered as a risk factor of COPD development. (Severe and lasted asthma with airways remodeling develops COPD).<sup>15–17</sup> Patients with ACOS have more disease progression rate, morbidity, mortality, respiratory symptoms and exacerbation attacks than patients with asthma or COPD alone.18-21 Some features of ACOS are listed in the following flowchart<sup>22</sup>:



ACOS also may affect smokers without previous or current asthma that develops as chronic airflow obstruction in eosinophilic background.<sup>23</sup> Patients with COPD and very positive bronchodilator test (>400 mL)is very possible to have some features of asthma and categorized as ACOS.<sup>24</sup> Differentiating ACOS from COPD has been considered important, as it affects the choice of therapy, i.e. the use of inhaled glucocorticoids. Differentiating ACOS from asthma, however, has received less attention, even though there are options of targeted treatment for COPD, such as long-acting muscarinic antagonists (LAMA) and roflumilast. Furthermore, ACOS is presently a phenotype with heterogenic and poorly defined clinical features. For this reason, there is an urgent need for the identification of specific characteristics and biomarkers of ACOS.<sup>25</sup> Regarding

to review of available data, most of studies about obstructive pulmonary disease excluded ACOS subjects, so little information about this entity is available and current study aimed to compare therapeutic response between asthma and COPD and ACOS subjects through spirometric data.

#### **METHOD**

This cross-sectional study included 30 known patients with asthma, 30 known patients with COPD and 30 known patients with ACOS according to GOLD GINA criteria, who refer to pulmonology clinics at Tabriz. The patients were included if they had good cooperation and mild to moderate severity of disease according to GOLD GINA guidelines. A patient with effective treatment for malignancy, infection, or cardiovascular disease was excluded.

We assessed demographic (age, sex, ...) and spirometric (post bronchodilator fev1 and fev1/fvc) parameters in all patients. The severity of asthma and ACOS was defined based on GINA guideline and severity of COPD was defined based on GOLD guideline. Spirometry was performed for all patients, then they took standard treatment regimen for 2 months according to GOLD GINA, and after this period spirometry was repeated. Changes of fev1 and fev1/fvc was recorded.

#### STATISTICAL ANALYSIS

Data analysis was conducted by Statistical Package for Social Sciences (SPSS) software version 26. Continous data were presented as mean and standard deviation and the categorical and nominal data were reported as number and percentages.

A p-value of less than 0.05 was considered significant.

#### RESULTS

A total of 90 obstructive pulmonary disease with the mean age of  $57.74 \pm 14.32$  were enrolled Asthma = Reference group.

in this study and 12 person of them didn't complete follow-up. About 66.7% of patients were male and 73.1% had moderate severity according to GOLD GINA guidelines.

Table 1 reports the FEV1/FVC changesFEV1/ FVC increases significantly more in Asmathic patients following treatment than in ACOS patients and least in COPD patients.

#### DISCUSSION

The aim of this study was to evaluate the differences of therapeutic response to standard treatment on base of spirometric finding in three groups of patients include asthma, COPD and ACOS. Totally Fev1 changes in response to treatment did not differ significantly between three groups (p > 0.05)but fev1/fvc changes differed significantly and this parameter in asthma was more than ACOS and in COPD was least. Considering FEV1/FVC changes, the spirometric symbolized therapeutic response in asthma was higher than that of ACOS, and the same was true for ACOS than COPD. ACOS may represent approximately 25% of COPD patients and nearly 20% of asthma patients. In primary COPD diagnosed patients identification of ACOS has therapeutic implication because the asthmatic component should be treated with inhaled corticosteroids that is only recommendation in COPD patients who suffer from recurrent exacerbations or have fev1/fvc<50%.<sup>26,27</sup> So, what is the harm in treating a patient with COPD and suspected (but unconfirmed) or historical (and no longer clinically relevant) asthma with an Inhaled corticosteroids

TABLE 1. FEV1/FVC changes.

Variable		Coefficient	Std. error	p
Age		-0.49	0.093	0.602
Sex		-2.047	2.750	0.459
Disease	COPD	-7.558	3.097	0.017
	ACOS	-6.959	3.112	0.028



FIG 1. Increasment of FEV1/FVC percentage.



FIG 2. FEV1 changes.

(ICS)? It has been suggested that the main reason for diagnosing concomitant asthma in patients with COPD is to identify those who are likely to have a better response to ICS.<sup>28</sup> According to defect of enough information about diagnosis and therapeutic response of ACOS, current study aimed to compare therapeutic response between asthma, COPD and ACOS by evaluation of changes of fev1 and fev1/ fvc. This type of research has never been attempted before, based on all available data. In a previous

Variable		Coefficient	Std. error	p
Age		-1.061	6.436	0.870
Sex		216.747	190.080	0.258
Disease	COPD	-306.705	214.026	0.156
	ACOS	-280.902	215.094	0.196

**TABLE 2.**FEV1 changes.

Asthma = Reference group.

study evaluating the reversibility of the airway after 12 years of active treatment for asthma, patients with ACOS had significantly higher fev1 and fvc results. Our results indicate that changes in fev1 in response to treatment did not differ significantly between three groups. But changes of fev1/fvc differed significantly between 3 groups (p < 0.05), and this parameter was greater in asthma than ACOS and in ACOS was more than COPD.

In previous study that performed by Tommola et al at Sinajoki respiratory department between 1999 and 2013 aimed to compare ACOS and adult onset asthma showed that reversibility in response to treatment in ACOS was more than asthmatic subjects, that is incompatible with our results.<sup>29</sup> Some previous studies compared decline in pulmonary function between asthma, COPD and ACOS and have not observed significant differences in the rate of decline of fev1 between three groups.<sup>27</sup>

#### CONCLUSION

Our results indicated the significant higher level of therapeutic response in asthma compared to ACOS and this parameter in ACOS was significantly more than COPD on the base on spirometric findings. There for use of ICS in treatment of ACOS might be effective for clinical improvement in comparison to COPD patients, however there is no definite available data about ACOS. For a patient with COPD, a diagnosis of concomitant asthma had better thoroughly considered and based on an individualized assessment – in order to weigh up the benefits and risks of treating the individual with ICS. However, considering the limitation of the study, use of another tools for monitoring disease include the assessment of frequency and severity of symptoms, inflammatory markers and airway hyper responsiveness, and cohort investigations are needed with a higher sample size to approve this preliminary results.

#### **CONFLICT OF INTEREST**

The authors declare that they have no conflict of interest.

#### ETHICAL

All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee (include name of committee + reference number) and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards.

Informed consent was obtained from all individual participants included in the study.

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