

MONTELUKAST INDUCES BETTER CONTROL OF SYMPTOMS AND MANAGEMENT OF LUNG FUNCTION, AND DECREASED INFLAMMATION IN WOMEN COMPARED WITH MEN

Giuseppe Spaziano¹, Renata Esposito¹, Domenico Giannattasio², Francesco Ferrigno², Angela Liparulo¹, Antonietta Rossi³, Fiorentina Roviezzo³, Liberato Berrino¹, Mario Polverino², Francesca Polverino⁴, Bruno D'Agostino¹

¹Department of Experimental Medicine, Section of Pharmacology "L. Donatelli", University of Campania "L. Vanvitelli", Naples - Italy, ²Pulmonary and Critical Care Medicine, Ospedale Scarlato, Scafati - Italy, ³Department of Pharmacy, School of Medicine, University of Naples Federico II, Naples - Italy, ⁴Asthma and Airway Disease Research Center, University of Arizona, Medicine, Tucson - USA

Introduction: Sex differences exist in the prevalence of asthma and allergic diseases, partially due to the effects of sex hormones on the development of allergic manifestations. Women are more prone to suffer allergic asthma, experience difficulties in controlling asthma symptoms, and show adverse responses to drugs. In particular, women tend to respond better to anti-leukotriene (LT) drugs than men, partially due to sex differences in endogenous LTs biosynthesis. However, the effects of anti-LT drugs on lung function, symptoms, and pulmonary and systemic inflammation in adult asthmatic women vs. men are unknown. The goal of this study was to characterize the effects of an anti-LT drug, montelukast (MS), in a randomized case-control pilot study of asthmatic adult females and males.

Materials and methods: Fortyone age- and sex-matched asthmatic subjects were randomly assigned to low-dose of inhaled corticosteroids (ICS) in combination with Montelukast (n=21 of which 11 females and 10 males), or low-dose ICS in combination with long-acting beta-agonists (LABA, n=20 of which 10 females and 10 males). The optimal control of the symptoms was achieved in both groups according to the Global Initiative for Asthma (GINA) guidelines. At enrollment, and after 13 weeks from the beginning of the therapy, pulmonary function tests and asthma control tests (ACT) were performed, and the fraction of exhaled nitric oxide (FeNO) and blood eosinophils levels were measured.

Results: Female subjects treated with ICS+MS had better control of the asthmatic symptoms (defined as higher ACT score, $P<0.001$) and improved pulmonary function (with higher forced expiratory volume in 1 second [FEV_{1} , $P<0.05$], forced vital capacity [FVC, $P<0.05$], compared with males in the same treatment group. Interestingly, ICS+MS-treated females had significantly lower levels of eosinophils ($P<0.01$), and FeNO ($P<0.01$) compared with males in the same treatment group. No differences between females and males treated with ICS + LABA were observed in any of these parameters evaluated.

Discussion and conclusion: MS treatment in women leads to better control of symptoms, better management of lung function, and decreased inflammation levels compared with MS treatment in men. These findings, if confirmed in larger cohorts, might be a step forward towards a gender-tailored approach to therapy with anti-LT drugs in asthma.