



Inflammation Markers and FEF₂₅₋₇₅: A Relevant Link in Children With Asthma

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Jang and colleagues investigated the relationship between exhaled nitric oxide (FeNO) and atopy profiles in children with asthma.¹ They focused their attention mainly on the FeNO, showing that this parameter varied according to the profile of atopy. The authors also investigated lung function, including bronchial hyperreactivity. However, forced vital capacity, forced expiratory volume in 1 second (FEV1), and their ratio were considered as parameters for lung function test. On the other hand, great attention has been recently paid to the relevance in asthma pathogenesis of FEF₂₅₋₇₅ values,² a parameter considered to better reflect small airways than the gold standard parameters of airway obstruction, such as forced vital capacity and FEV1.² This may be even more relevant in childhood.³ Indeed, in allergic children with asthma, impaired FEF₂₅₋₇₅ values appear to be related to severe bronchial hyperreactivity,⁴ reversible airway obstruction⁵; perception of breathlessness and of positive response to reversibility test, assessed by visual analogue scale.^{6,7}

Since impaired FEF₂₅₋₇₅ values were reported to be significantly associated also with the presence of high fractional FeNO levels in children with allergic rhinitis and/or asthma,⁸ it is possible that this marker of small airway obstruction could be related also to the degree of allergic sensitization (and of systemic allergic inflammation).

Previously, it has been reported that children with asthma due to mite allergy showed a tightly link between allergen-specific serum IgE and markers of allergic inflammation, such as eosinophil and FeNO, but no correlations were detected with bronchial obstruction, expressed as FEV1 values.⁹ On the basis of these considerations, we evaluated a cohort of children with allergic asthma, sensitized to house dust mites, aiming at investigating possible relationship between lung function, mainly concerning FEF₂₅₋₇₅, and inflammation parameters, such as blood eosinophilia, allergen-specific serum IgE, and FeNO. Fifty-six children, 5 to 12 years of age, were studied. Blood eosinophil number inversely and moderately related with FEF₂₅₋₇₅ val-

ues ($r = -0.52$; $P < 0.0001$), as reported in Figure A. Also the relative blood eosinophil numbers, expressed as percentage on total blood leukocytes count, significantly and negatively correlated with FEF₂₅₋₇₅ values ($r = -0.4$; $P = 0.001$) (Figure B). Serum levels of house dust mites specific IgE showed negative significant correlations with FEF₂₅₋₇₅ values ($r = -0.31$; $P = 0.01$), as reported in Figure C. Finally, a significant negative relationship was demonstrated between FeNO and FEF₂₅₋₇₅ values ($r = -0.33$; $P = 0.01$) (Figure D). In contrast, a significant negative correlation with FEV1 values was detected only for blood eosinophil numbers ($r = -0.35$; $P = 0.006$). These findings further confirm the possibility to detect links between markers of allergic inflammation and airflow limitation and further support the importance of FEF₂₅₋₇₅ assessment in the evaluation of childhood allergic asthma.

REFERENCES

1. Jang WN, Park IS, Choi CH, Bauer S, Harmin S, Seo SC, et al. Relationships between exhaled nitric oxide and atopy profiles in children with asthma. *Allergy Asthma Immunol Res* 2013;5:155-61.
2. Lipworth B. Targeting the small airways asthma phenotype: if we can reach it, should we treat it? *Ann Allergy Asthma Immunol* 2013; 110:233-9.
3. Ciprandi G, Capasso M, Tosca M, Salpietro C, Salpietro A, Marseglia G, et al. A forced expiratory flow at 25-75% value <65% of predicted should be considered abnormal: a real-world, cross-sectional study. *Allergy Asthma Proc* 2012;33:e5-8.
4. Ciprandi G, Tosca MA, Capasso M. Forced expiratory flow between 25 and 75% of vital capacity might be a predictive factor for bronchial hyperreactivity in children with allergic rhinitis, asthma, or

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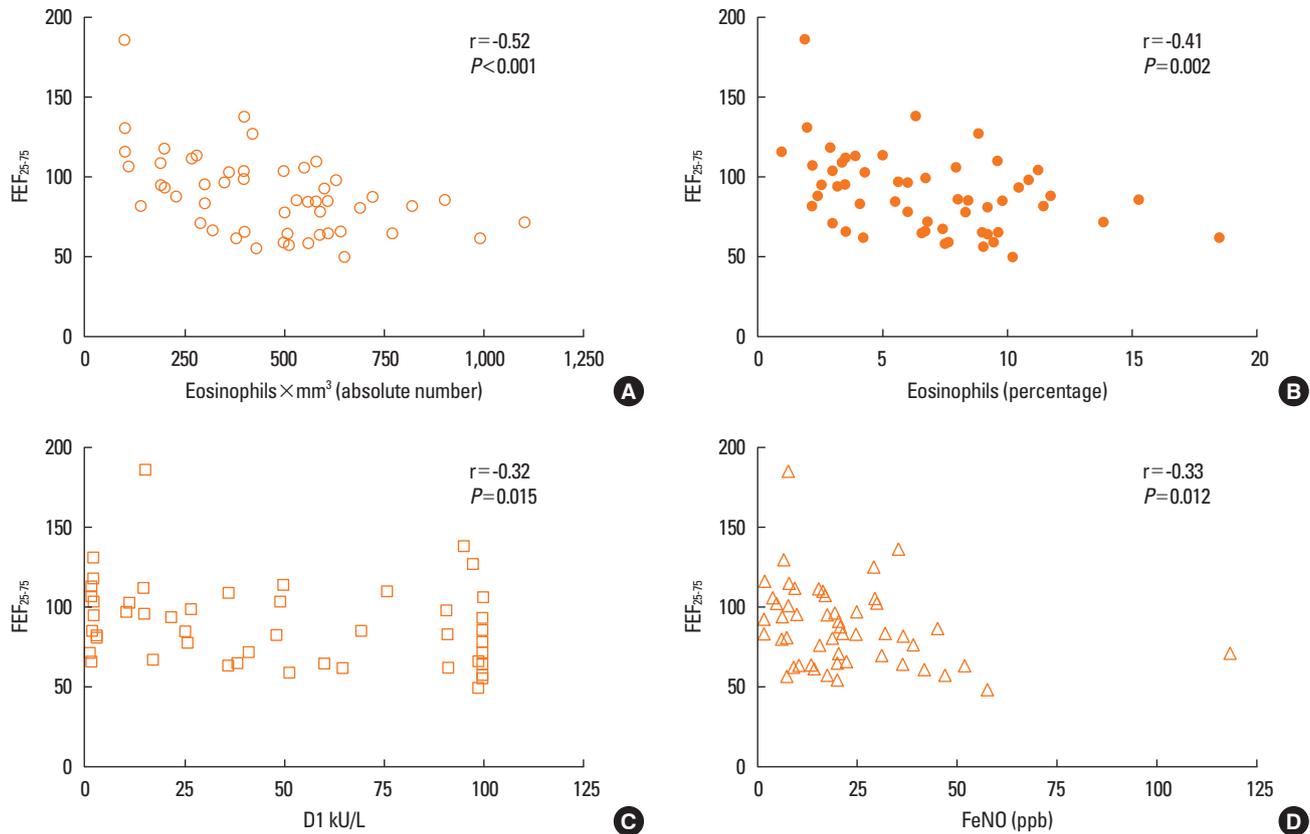


Figure. Relationship between FEF₂₅₋₇₅ values and different inflammatory/allergy markers: blood eosinophils (as absolute number (A), as relative number (B), serum IgE to house dust mites (C), and FeNO (D).

both. *Allergy Asthma Proc* 2011;32:e22-8.

- Simon MR, Chinchilli VM, Phillips BR, Sorkness CA, Lemanske RF Jr, Szefer SJ, et al. Forced expiratory flow between 25% and 75% of vital capacity and FEV₁/forced vital capacity ratio in relation to clinical and physiological parameters in asthmatic children with normal FEV₁ values. *J Allergy Clin Immunol* 2010;126:527-34.e1-8.
- Tosca MA, Silvestri M, Olcese R, Pistorio A, Rossi GA, Ciprandi G. Breathlessness perception assessed by visual analogue scale and lung function in children with asthma: a real-life study. *Pediatr Allergy Immunol* 2012;23:537-42.
- Tosca MA, Silvestri M, Rossi GA, Ciprandi G. Perception of bronchodilation assessed by Visual Analogue Scale in children with asthma. *Allergol Immunopathol (Madr)*. Forthcoming 2012.
- Ciprandi G, Capasso M. Association of childhood perennial allergic rhinitis with subclinical airflow limitation. *Clin Exp Allergy* 2010;40:398-402.
- Silvestri M, Pistorio A, Battistini E, Rossi GA. IgE in childhood asthma: relevance of demographic characteristics and polysensitisation. *Arch Dis Child* 2010;95:979-84.