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Data in Brief





Data Article

Data set on a study of gene expression in peripheral samples to identify biomarkers of severity of allergic and nonallergic asthma



Selene Baos ^{a,b}, David Calzada ^a, Lucía Cremades ^a, Joaquín Sastre ^{b,c}, Joaquín Quiralte ^d, Fernando Florido ^e, Carlos Lahoz ^{a,b}, Blanca Cárdaba ^{a,b,*}

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ABSTRACT

This article contains information related to the research article entitled "Biomarkers associated with disease severity in allergic and nonallergic asthma" (S. Baos, D. Calzada, L. Cremades, J. Sastre, J. Quiralte, F. Florido, C. Lahoz, B. Cárdaba, In press). Specifically, the clinical criteria stablished for selecting the study population (n=104 subjects) are described. Moreover, this article describes the criteria for selecting the 94 genes to be analyzed in PBMCs (peripheral blood mononuclear cells), it is provided a description of these genes and a Table with the genes most differentially expressed by clinical phenotypes and, finally it is detailed the experimental methodology followed for studying the protein expression of MSR1 (macrophage scavenger receptor 1), one of the genes evaluated in the research.

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E-mail address: bcardaba@fjd.es (B. Cárdaba).

^a Immunology Department, IIS- Jiménez Díaz Foundation, UAM, Madrid, Spain

^b CIBERES, CIBER of Respiratory Diseases, Spain

^c Allergy Department, Jiménez Díaz Foundation, Madrid, Spain

^d Allergy Department, Vírgen del Rocío University Hospital, Seville, Spain

^e Allergy Department, San Cecilio University Hospital, Granada, Spain

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^{*}Correspondence to: Immunology Department, IIS- Jiménez Díaz Foundation, Avda Reyes Católicos N° 2, 28040 Madrid, Spain. Fax: +34 915448246.

Specifications Table

Subject area More specific subject area	Biology Immunology, biomarkers, asthma, allergy.
Type of data	Table, text file.
How data was acquired	Bibliographic search, qRT-PCR, Western Blot.
Data format	Raw
Experimental	Subjects' diagnosis was done according to the GEMA (Spanish Guide for
factors	Asthma Management) classification. PBMCs were extracted from peripheral blood through gradient separation. RNA and protein from PBMCs were extracted with TRIzol's method.
Experimental features	Genes selected were according to the 3 criteria stated. Through qRT-qPCR, gene expression differences among clinical groups were studied. The highest statistically significant data among the three clinical phenotypes are showed. Western blot was done to determine the protein expression of one of the genes studied.
Data source location	Madrid, Spain; Seville, Spain; Granada, Spain.
Data accessibility	Data is with this article.

Value of the data

- Data presented here shows the selection and clinical criteria [1] of the study population.
- Gene selection criteria of interesting candidates to be asthma' biomarkers are provided in order to understand the validity of genes studied.
- A gene list of candidate biomarkers of asthma and allergy diseases is suggested for studying.
- A summary of the most differential genes among clinical phenotypes is showed. These data could be important for future biomarkers analyses.
- Western-blot method for MSR1 expression on protein extracted from PBMCs could be useful for future research.

1. Data

The data shown in the article give information on the criteria of patients' selection and the criteria for choosing genes to be studied as candidate biomarkers for these diseases in peripheral samples. The specific western-blot method for the analysis of MSR1 expression on protein extracted from PBMCs is provided. Table 1 provides a list of candidate genes to be validated as relevant biomarkers and Table 2 summarize the possible biomarkers that differentiate asthmatic and allergic phenotypes.

2. Experimental design, materials and methods

2.1. Subjects

The study population comprised 104 unrelated subjects, 30 healthy control (HC) subjects, 30 patients with nonallergic asthma (NA), 30 with allergic asthma (AA), and 14 nonasthmatic allergic (AR) subjects. The samples of the groups with asthma came from the asthma biobank of the CIBERES (IIS-Fundación Jiménez Díaz-UAM, Madrid). A biorepository in which were included samples from clinically well-characterized subjects, from 5 Spanish Hospitals participant of this network (*Fundación*

Table 1 List of the 94 genes studied.

Gene symbol	Gene name	Selection criteria	Detector	
DAM33	ADAM metallopeptidase domain 33	3	ADAM33-Hs00905552_m	
DRB1	adrenoceptor beta 1	2	ADRB1-Hs02330048_s1	
KT1	v-akt murine thymoma viral oncogene homolog 1	2	AKT1-Hs00178289_m1	
LOX15	arachidonate 15-lipoxygenase	1	ALOX15-Hs00993765_g1	
LOX5	arachidonate 5-lipoxygenase	2	ALOX5-Hs01095330_m1	
IPAF1	apoptotic peptidase activating factor 1	2	APAF1-Hs00559441_m1	
BAX	BCL2-associated X protein	2	BAX-Hs00180269_m1	
C3AR1	complement component 3a receptor 1	2	C3AR1-Hs00269693	
CCL11	chemokine (C-C motif) ligand 11	3	CCL11-Hs00237013_m1	
CL-17	chemokine (C-C motif) ligand 17	3	CCL17-Hs00171074_m1	
CCL5	chemokine (C-C motif) ligand 5	3	CCL5-Hs00982282_m1	
D40	CD40 molecule, TNF receptor superfamily mem-	2	CD40-Hs01002913_g1	
	ber 5			
CD48	CD48 molecule	2	CD48-Hs00914738_m1	
CD86	CD86 molecule	2	CD86-Hs01567026_m1	
CHI3L1	chitinase 3-like 1 (cartilage glycoprotein-39)	1	CHI3L1-Hs00609691_m1	
LCA1	chloride channel accessory 1	1	CLCA1-Hs00976287_m1	
PA3	carboxypeptidase A3 (mast cell)	1	CPA3-Hs00157019_m1	
RTAP	cartilage associated protein	2	CRTAP-Hs00197261_m1	
CTSC	cathepsin C	1	CTSC-Hs00175188_m1	
TSG	cathepsin G	1	CTSG-Hs01113415_g1	
CX3CR1	chemokine (C-X3-C motif) receptor 1	1	CX3CR1-Hs01922583_s1	
OUSP1	dual specificity phosphatase 1	1	DUSP1-Hs00610256_g1	
RNASE3	ribonuclease, RNase A family, 3	3	RNASE3-Hs01923184_s1	
EIF5A	eukaryotic translation initiation factor 5A	1	EIF5A-Hs00744729_s1	
ОХРЗ	forkhead box P3	3	FOXP3-Hs01085834_m1	
PR3	formyl peptide receptor 3	2	FPR3-Hs00266666_s1	
GADD45B	growth arrest and DNA-damage-inducible, beta	1	GADD45B-Hs04188837_g	
GPX3	glutathione peroxidase 3 (plasma)	1	GPX3-Hs01078668_m1	
SZMH	granzyme H (cathepsin G-like 2, protein h-CCPX)	2	GZMH-Hs00277212_m1	
HLA-DQB1	major histocompatibility complex, class II, DQ	1, 2	HLA-DQB1-Hs03054971_	
ILI-DQD1	beta 1	1, 2	1121 2021 11303034371	
HLA-DRB1	major histocompatibility complex, class II, DR beta 1	2	HLA-DRB1-Hs99999917_1	
FNG	interferon, gamma	3	IFNG-Hs00989291_m1	
L-10	interleukin 10	2	IL10-Hs00961622_m1	
L13	interleukin 13	1	IL13-Hs00174379_m1	
L-17	interleukin 17A	3	IL17A-Hs00174383_m1	
L1R1	interleukin 1 receptor, type I	1	IL1R1-Hs00991002_m1	
L1R2	interleukin 1 receptor, type II	1	IL1R2-Hs01030384_m1	
L-1R2 L-2	interleukin 2	3	IL2-Hs00174114_m1	
L-25 L-25	interleukin 25	3	IL25-Hs03044841_m1	
L-23 L2RB	interleukin 2 receptor, beta	1	IL2RB-Hs01081697_m1	
	* '		IL33-Hs00369211_m1	
L33 L-4	interleukin 33 interleukin 4	1 3	IL33-HS00369211_m1 IL4-Hs00174122_m1	
	interleukin 4 interleukin 4 receptor	3	_	
L4R	interleukin 4 receptor interleukin 5		IL4R-Hs00166237_m1	
L5 16		1	IL5-Hs01548712_g1	
L6	interleukin 6 (interferon, beta 2)	1, 2	IL6-Hs00985639_m1	
L8	interleukin 8	1	IL8-Hs00174103_m1	
L-9 DAV2	interleukin 9	3	IL9-Hs00914237_m1	
RAK3	interleukin-1 receptor-associated kinase 3	3	IRAK3-Hs00936103_m1	
TGAL	integrin, alpha L (antigen CD11A (p180), lym- phocyte	2	ITGAL-Hs00158218_m1	
TCD7	function-associated antigen 1; alpha polypeptide)	ว	ITCD7 Uc01ECE7E01	
TGB7	integrin, beta 7	2	ITGB7-Hs01565750_m1	
TGB8	integrin, beta 8	2	ITGB8-Hs00174456_m1	
CK	lymphocyte-specific protein tyrosine kinasep	2	LCK-Hs00178427_m1	
GALS3	lectin, galactoside-binding, soluble, 3	3	LGALS3-Hs00173587_m1	
YN	v-yes-1 Yamaguchi sarcoma viral related onco- gene homolog	2	LYN-Hs00176719_m1	
MAPK13	mitogen-activated protein kinase 13	2	MAPK13-Hs00559623_m	
/ISR1	macrophage scavenger receptor 1	2	MSR1-Hs00234007_m1	
ATT CO	mucin 2, oligomeric mucus/gel-forming	1	MUC2-Hs03005103_g1	
MUC2	macin 2, origonierie macas/ger-jorning	•		

Table 1 (continued)

Gene symbol	Gene name	Selection criteria	Detector MUC5B-Hs00861595_m1	
мис5В	mucin 5B, oligomeric mucus/gel-forming	1		
NCF2	neutrophil cytosolic factor 2	1	NCF2-Hs01084940_m1	
NFATC1	nuclear factor of activated T-cells, cytoplasmic,	2	NFATC1-Hs00542678_m1	
	calcineurin-dependent 1			
NFKBIZ	nuclear factor of kappa light polypeptide gene enhancer in B-cells inhibitor, zeta	1	NFKBIZ-Hs00230071_m1	
NLRP3	NLR family, pyrin domain containing 3	2	NLRP3-Hs00918082_m1	
NOS2A	nitric oxide synthase 2, inducible	1	NOS2-Hs01075529_m1	
ORMDL3	ORM1-like 3 (S. cerevisiae)	1	ORMDL3-Hs00918021_m1	
PHLDA1	pleckstrin homology-like domain, family A, member 1	1	PHLDA1-Hs00705810_s1	
PI3	peptidase inhibitor 3, skin-derived	1	PI3-Hs00160066_m1	
POSTN	periostin, osteoblast specific factor	1	POSTN-Hs01566734_m1	
PRKACA	protein kinase, cAMP-dependent, catalytic, alpha	2	PRKACA-Hs00427274_m1	
PRKACB	protein kinase, cAMP-dependent, catalytic, beta	2	PRKACB-Hs01086757_m1	
PTGER2	prostaglandin E receptor 2 (subtype EP2), 53 kDa	2	PTGER2-Hs04183523_m1	
PTPRC	protein tyrosine phosphatase, receptor type, C	3	PTPRC-Hs04189704_m1	
S100A9	S100 calcium binding protein A9	1	S100A9-Hs00610058_m1	
S1PR5	sphingosine-1-phosphate receptor 5	2	S1PR5-Hs00928195_s1	
SCD	stearoyl-CoA desaturase (delta-9-desaturase)	1	SCD-Hs01682761_m1	
SELL	selectin L	2	SELL-Hs00174151_m1	
SERPINB2	serpin peptidase inhibitor, clade B (ovalbumin), member 2	1, 2	SERPINB2-Hs01010736_m	
SERPINB4	serpin peptidase inhibitor, clade B (ovalbumin), member 4	1	SERPINB4-Hs01691258_g1	
SMURF1	SMAD specific E3 ubiquitin protein ligase 1	2	SMURF1-Hs00905759_m1	
SOS1	son of sevenless homolog 1 (Drosophila)	2	SOS1-Hs00362308_m1	
SPN	sialophorin	2	SPN-Hs01872322_s1	
SPP1	secreted phosphoprotein 1	3	SPP1-Hs00959010_m1	
STAT1	signal transducer and activator of transcription 1.91 kDa	3	STAT1-Hs01013996_m1	
SVIL	supervillin	1	SVIL-Hs00931028_m1	
TAGAP	T-cell activation RhoGTPase activating protein	2	TAGAP-Hs00299284_m1	
TCF21	transcription factor 21	1	TCF21-Hs00162646_m1	
TGFB1	transforming growth factor, beta 1	2	TGFB1-Hs00998133_m1	
TLR4	toll-like receptor 4	2	TLR4-Hs00152939_m1	
TNFA	tumor necrosis factor	3	TNF-Hs01113624_g1	
TNFAIP3	tumor necrosis factor, alpha-induced protein 3	1	TNFAIP3-Hs00234713_m1	
TRIM37	tripartite motif containing 37	2	TRIM37-Hs00248701_m1	
TSLP	thymic stromal lymphopoietin	1	TSLP-Hs00263639_m1	
VCAN	versican	2	VCAN-Hs00171642_m1	
ZAP70	zeta-chain (TCR) associated protein kinase 70 kDa	2	ZAP70-Hs00896347_m1	

Selection criteria were: 1. Relevant genes by differential expression or SNP studies in asthma/allergy, which were found in more than one independent work following a literature search; 2. Genes with differential expression found in results of previous studies from our laboratory; 3. Genes of interest because of their role in cellular plasticity, inflammation and/or regulation that could have been excluded by the other criteria. The detector refers to the specific primer of each gene used to carry out qRT-PCR.

Jiménez Díaz Hospital and Doce de Octubre Hospital from Madrid, Doctor Negrín Hospital from Las Palmas de Gran Canaria, Clinic Hospital and Sant Pau Hospital both from Barcelona). These patients fulfilled the following criteria: severe, mild, or moderate asthma diagnosis assigned according to the GEMA [1]; no treatment was given before or during the collection of the samples. Pulmonary function test was determined by percentage of forced vital capacity (FVC) and forced vital volume in one second (FVE₁). Patients with allergic asthma showed a positive skin prick test result for some of the airborne allergens from a battery of common allergens.

HCs were healthy subjects with no history of respiratory diseases. HCs and patients with allergy (rhinitis) without asthma were recruited and diagnosed at the Allergy Service of two hospitals in

 Table 2

 Differential genes among clinical phenotypes.

Comparison	Number of genes with sig- nificant differential expression*	Number of genes upregulated	Number of genes downregulated	Genes statistically significant with a RQ $>$ 10
NAvs AA	74	74 in NA	-	CCL5, CHI3L1, CTSG, GMH, IL1-R2
NAvs AR	66	64 in NA	2 in NA	CCL5, CRTAP, GPX3, HLA-DQB1, IL-10, IL2RB, MSR1, NLRP3, PHLDA1, SERPINB2, PI3
AAvs AR	14	4 in AA	10 in AA	CHI3L1, CPA3, CTSG, PI3

NA: Nonallergic asthma group; AA: allergic asthma group; AR: nonasthmatic allergy (rhinitic) group; RQ: relative quantification.

Andalusia (Spain), *Virgen del Rocío* University Hospital from Seville, and *San Cecilio* University Hospital from Granada, Spain. AR patients fulfilled the following criteria: seasonal rhinitis without asthma, positive skin prick test for some of the airborne allergens from a battery of common allergens, and no previous immunotherapy.

HC and AR biological samples that were not used in this work were stored in the FJD Biobank, IIS-Fundación Jiménez Díaz Madrid.

Informed consent was obtained from each subject. Ethical approval for the study was obtained from the Ethical and Research Committee of the participating hospitals.

2.2. Gene selection criteria

Ninety-four genes (Table 1) were chosen following three main criteria for a gene expression analysis [2] through quantitative real time PCR with RNA of the study population described before:

- a. Relevant genes associated with asthma and allergic diseases in more than one independent work, selected after a Pubmed literature search of analyses of differential gene expression, or polymorphic variants (SNPs) related to the disease.
- b. Relevant genes previously described by our group [3].
- c. Genes excluded by the other two criteria but that could be interesting due to their implication in cellular plasticity, inflammation, and/or regulation of the disease.

2.3. Gene expression analysis

Gene expression analysis between the 3 clinical groups is summarized in Table 2. The statistical analysis for testing differential gene expression was performed by using the StatMiner program (http://www.integromics.com/StatMiner). This program follows a simple, step-by-step analysis workflow guide that includes parametric, non-parametric, and paired tests for relative quantification of gene expression, as well as 2-way ANOVA for two-factor differential expression analysis. Significance was defined by RQ (relative quantification) < -2 or > 2 and corrected P value (< 0.05) adjusting the P value with the Benjamini–Hochberg FDR method.

2.4. MSR1 protein analysis by Western Blot

The protein expression of MSR1 was analyzed [2]. Specific protein was extracted from PBMCs (10⁶ cells) using the TRIzol method (Invitrogen, Carlsbad, CA, USA) and quantified by the BCA method (Thermo Scientific, Rockford, IL USA). Western blot used was the Invitrogen WesternBreeze Chemiluminescent Western Blot Immunodetection Kit (Life Technologies) following the

^{*} Significance established at an adjusted p < 0.05 and a RQ < -2 or > 2. All genes mentioned in the last column are overexpressed except the ones marked in bold which are underexpressed.

manufacturer's instructions with minor modifications. Briefly, 40 μg of proteins from each subject were running in a 12% SDS-PAGE Novex BoltTM Mini gels (Life Technologies) and transferred using the Invitrogen BlotTM Dry Blotting System to nitrocellulose membranes. After 30 min of incubation with blocking solution, were incubated overnight at 4 °C with rabbit anti-human polyclonal CD204/Macrophage Scavenger Receptor I antibody (dilution 1:2500) (Thermo Scientific) as specific antibody and, with a rabbit anti-human monoclonal β-Actin antibody (dilution 1:1000) (Cell Signalling Technology, Danvers, MA, USA) as control. The result was visualized by chemiluminiscence using a luminescent image analyzer: ImageQuant LAS 4000 (GE Healthcare Life Science, Little Chalfont, Buckinghamshire, UK). Data of MSR1 results were relativize to β-Actin expression.

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Transparency document. Supporting information

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References

- [1] GEMA (Guía española del manejo del asma), Arch. Bronconeumol. 45 (2009) 2-35.
- [2] S.Baos, D.Calzada, L.Cremades, J.Sastre, J.Quiralte, F.Florido, et al.Biomarkers associated with disease severity in allergic and nonallergic asthma. Molecular Immunol. In press.
- [3] M. Aguerri, D. Calzada, D. Montaner, M. Mata, F. Florido, J. Quiralte, et al., Differential gene-expression analysis defines a molecular pattern related to olive pollen allergy, J. Biol. Regul. Homeost. Agents 27 (2013) 337–350.