

RESEARCH ARTICLE

Clinical relevance of eosinophils, basophils, serum total IgE level, allergen-specific IgE, and clinical features in atopic dermatitis

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Abstract

Background: Atopic dermatitis (AD) is an inflammatory disease with diverse clinical features. Although AD is diagnosed mainly by clinical features, the laboratory abnormalities can be found in most patients and may be of diagnostic value. However, few studies have been performed on the clinical significance of laboratory abnormalities in adult and adolescent AD.

Methods: Adult and adolescent patients with AD were included in this study. The questionnaire and dermatological examination were completed by investigators. Laboratory tests included complete blood count, serum total IgE, and allergen-specific IgE.

Results: A total of 473 patients were recruited and 396 of them were diagnosed as AD. Increased serum total IgE level, peripheral eosinophils, and basophils were seen more frequently in AD patients than in non-AD patients ($P < .05$). Positive aeroallergens were seen more in AD patients than in non-AD patients ($P < .05$). Both total serum IgE level ($R = .286$, $P < .001$) and peripheral eosinophils ($R = .444$, $P < .001$) significantly correlated with EASI score. Serum total IgE level and extrinsic type AD decreased with age. Patients with elevated serum total IgE are more likely to have a personal history of atopic diseases ($P = .014$). AD-associated symptoms (such as flexural dermatitis, white dermographism, and anterior neck folds) are more frequently observed in AD patients with high serum IgE or eosinophilia ($P < .05$).

Conclusion: The serum total IgE level, allergen-specific IgE, peripheral eosinophils, and basophils are important for the diagnosis of AD. And they are associated with the severity, age groups, and clinical manifestations.

KEYWORDS

atopic dermatitis, basophils, clinical features, eosinophils, IgE

1 | INTRODUCTION

Atopic dermatitis (AD) is an inflammatory skin disease characterized by chronic recurrent dermatitis with profound pruritus. Patients with AD often have personal or family history of atopic diseases such as asthma, allergic rhinitis, and allergic conjunctivitis.¹

AD used to be diagnosed by explicit criteria that require information obtained from patient's history and physical examination according to different criteria.²⁻⁴ However, Liu et al⁵ reported that peripheral eosinophil and serum total IgE were also important in the diagnosis of AD. Elevation total serum IgE level, positive allergen-specific IgE, eosinophilia, and basophilia were common in AD. Studies have suggested a significant association between the severity of AD

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and systemic measures of allergic inflammation including serum IgE and blood eosinophil in infant and children patients. However, few studies were finished in adolescent or adult patients.

The clinical manifestations of AD are heterogeneous. Three stages are proposed including infantile AD, childhood AD, and adolescent/adult AD.⁶ Typical clinical presentations are similar in all the stages such as head and neck dermatitis and xerosis. However, some clinical features of AD are more specific to adults, including lichenification and prurigo. Nummular eczema, xerotic eczema, eczema of the hands, and feet are considered as special forms of AD.⁷ Few studies focused on the relationship between clinical characteristics and laboratory findings.

2 | MATERIALS AND METHODS

2.1 | Patients

A total of 396 adult/adolescent patients with AD (201 female and 195 male) and 77 non-AD patients were recruited. No significant difference in the median age and gender ratio between the groups was found. The informed consents were obtained from each patient. The diagnosis of AD was based on the Chinese criteria of AD.⁵ All patients were at an active stage of the disease, before the administration of systemic treatment. Atopic disorders or other inflammatory skin diseases were excluded in all healthy controls. The severity of AD was assessed by the eczema area and severity index (EASI) score at the first visit. Mild severity was defined as an EASI score of <7, moderate severity as ≥ 7 and <21, and severe AD as ≥ 21 .

2.2 | Questionnaire and dermatological examination

The patients were required to complete the questionnaire included demographic characteristics, atopic history (allergic rhinitis, allergic conjunctivitis, and allergic asthma), and family history. The patients also received a dermatological examination.

2.3 | Laboratory tests

The complete blood count, serum total IgE, and allergen-specific IgE were measured in each patient. Specific IgE antibody to aeroallergens and food allergens was measured in 226 AD patients and 65 non-AD patients. Food allergen-specific IgE antibody detection kit was used for allergen detection, including *D. pteronyssinus*, artemisia, mixture of epithelia, mixed molds, pellitory, willow, poplar, egg, milk, shrimp, mutton, beef, fish, crab, and staple foods (including wheat and corn).

2.4 | Statistical analysis

All the data were input using Epidata 3.1 software and analyzed by SPSS 22.0. Chi-square, Mann-Whitney *U* tests, and Kruskal-Wallis

TABLE 1 Comparison of laboratory findings in AD patients and non-AD patients

Laboratory tests	AD (n = 396)	Non-AD (n = 77)	P value
Serum total IgE (IU/mL) ^a	MD: 187.7	MD: 47.4	<.001
	25th: 56.0	25th: 19.3	
	75th: 722.0	75th: 132.5	
EOS (%) ^a	MD: 3.6	MD: 2.9	.022
	25th: 1.9	25th: 1.6	
	75th: 6.7	75th: 5.1	
EOS count ($\times 10^9/L$) ^a	MD: 0.24	MD: 0.18	.011
	25th: 0.12	25th: 0.10	
	75th: 0.45	75th: 0.37	
BAS (%) ^a	MD: 0.572	MD: 0.581	.383
	25th: 0.400	25th: 0.400	
	75th: 0.800	75th: 0.700	
Serum total >100 IU/mL, n (%) ^b	248 (62.6%)	15 (19.5%)	<.001
EOS% >5%, n (%) ^b	146 (36.9%)	12 (15.6%)	<.001
EOS count > $0.5 \times 10^9/L$, n (%) ^b	82 (20.7%)	6 (7.8%)	.006
BAS% >1%, n (%) ^b	55 (13.9%)	4 (5.2%)	.037

Abbreviations: MD, median values; 25th, 25th percentiles; 75th, 75th percentiles.

^aMann-Whitney *U* tests.

^bPearson chi-square test.

test were used for non-normally distributed variables. Simple linear regression analysis and Spearman's correlation index were applied to describe the correlation between measurement data. $P < .05$ was considered significant.

3 | RESULTS

3.1 | Laboratory findings in AD and non-AD patients

The laboratory findings in AD and non-AD patients are shown in Tables 1 and 2. Serum total IgE (62.6% vs 19.5%, $P < .001$), eosinophilia (36.9% vs 15.6%, $P < .001$), and basophilia (13.9% vs 5.2%, $P = .037$) in AD patients were significantly higher than in non-AD patients. The prevalence of positive IgE to aeroallergens was higher in AD patients than in non-AD patients (39.8% vs 24.6%, $P = .028$). A higher prevalence of positive IgE for *D. pteronyssinus* was found in AD patients than in non-AD patients (25.3% vs 10.8%, $P = .011$). No significant difference was found for food allergens.

3.2 | Laboratory findings and AD severity

The relationship between serum total IgE, blood eosinophil and basophil count, and EASI score is shown in Table 3. The severity was stratified according to EASI scores as follows: mild <7 ($n = 176$),

TABLE 2 Allergen-specific IgE in AD patients and non-AD patients

Allergens	AD (n = 226)	Non-AD (n = 65)	P value	χ^2
Any positive allergens	98 (43.4%)	19 (29.2%)	.045	4.194
Positive food allergens	36 (15.9%)	7 (10.8%)	.427	1.067
Positive aeroallergens	90 (39.8%)	16 (24.6%)	.028	5.043
D. pteronyssinus	57 (25.3%)	7 (10.8%)	.011	6.220
Artemisia	23 (10.2%)	5 (7.7%)	.640	0.370
Mixture of epithelia (cats and dogs)	19 (8.4%)	3 (4.6%)	.428	1.055
Mixed molds	21 (9.3%)	6 (9.2%)	1.000	0.001
Pellitory	6 (2.7%)	1 (1.5%)	1.000	0.273
Willow	9 (4.0%)	2 (3.1%)	1.000	0.118

Note: Pearson chi-square test was used.

moderate ≥ 7 and < 21 (n = 131), and severe ≥ 21 (n = 89). We found that serum total IgE (P = .025), blood eosinophil count (P < .001), EOS% (P < .001), and BAS% (P = .036) elevated significantly with severity. Linear regression analysis showed that the increased serum total IgE level and blood eosinophil count significantly associated with the EASI score (R = .286, R = .444, respectively, Figure 1).

3.3 | Laboratory abnormalities in different age groups

The patients were divided into three different age groups: adolescence (12-18 years old), adults (19-60 years old), and seniors (>60 years old). No significant difference in severity was found

TABLE 3 Relationship between laboratory findings and severity of AD

Laboratory tests	Mild (n = 176)	Moderate (n = 131)	Severe (n = 89)
Serum total IgE (IU/mL) ^a	MD: 139.4***,# 25th: 51.7 75th: 429.8	MD: 302.0 25th: 54.3 75th: 831.3	MD: 356.0 25th: 81.1 75th: 1576.5
EOS (%) ^a	MD: 2.8***,### 25th: 1.7 75th: 4.6	MD: 4.5* 25th: 2.3 75th: 7.1	MD: 5.4 25th: 2.7 75th: 12.7
EOS count ($\times 10^9/L$) ^a	MD: 0.19***,### 25th: 0.08 75th: 0.31	MD: 0.27* 25th: 0.16 75th: 0.46	MD: 0.36 25th: 0.20 75th: 0.88
BAS (%) ^a	MD: 0.5* 25th: 0.3 75th: 0.7	MD: 0.6 25th: 0.4 75th: 0.8	MD: 0.6 25th: 0.5 75th: 0.8
Serum total >100 IU/mL, n (%) ^b	98 (55.7%)*	86 (65.6%)	64 (71.9%)
EOS% >5%, n (%) ^b	34 (19.3%)*,###	60 (45.8%)	52 (58.4%)
EOS count $> 0.5 \times 10^9/L$, n (%) ^b	15 (8.5%)*,##	27 (20.6%)*	40 (44.9%)
BAS% >1%, n (%) ^b	24 (13.6%)	18 (13.7%)	13 (14.6%)

Abbreviations: MD, median values; 25th, 25th percentiles; 75th, 75th percentiles.

^aKruskal-Wallis test.

^bPearson chi-square test.

*P < .05 vs severe group.

**P < .01 vs severe group.

***P < .001 vs severe group.

#P < .05 vs moderate group.

##P < .01 vs moderate group.

###P < .001 vs moderate group.

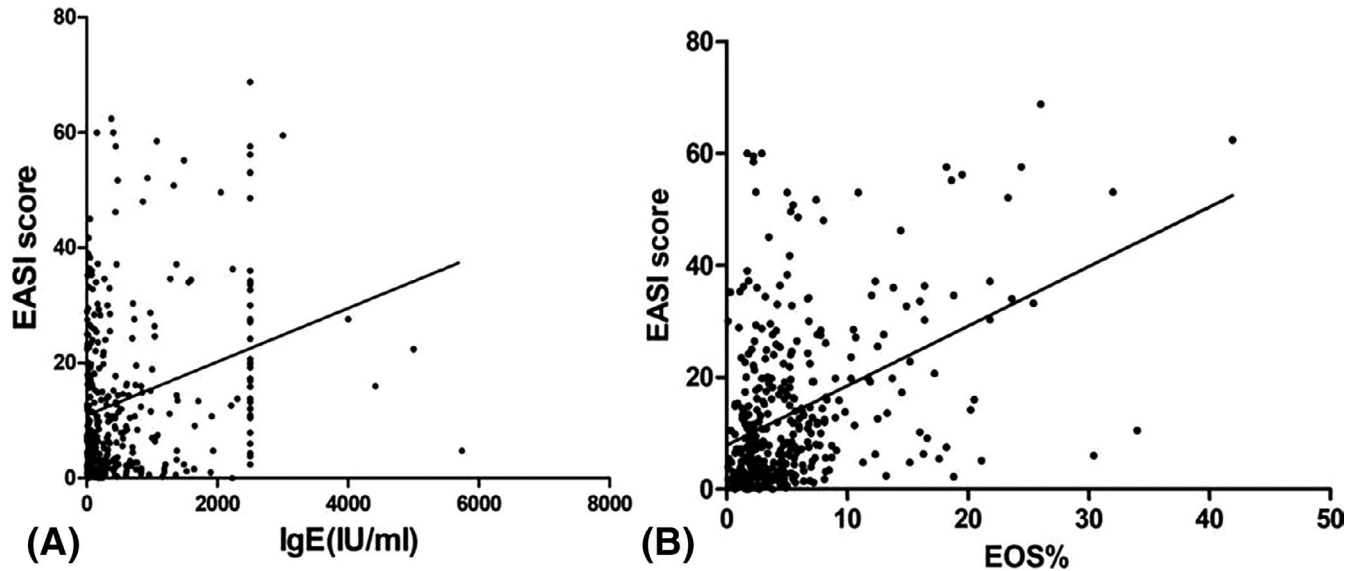


FIGURE 1 Simple linear regression analysis and Spearman's correlation analysis between IgE, EOS%, and EASI score. A, Association between total serum IgE and EASI score ($R = .286$, $P < .001$). B, Association between eosinophil count and EASI score ($R = .444$, $P < .001$). EOS%: percentage of peripheral eosinophils. EASI, eczema area and severity index

between age groups. The serum total IgE level decreased significantly with age ($P < .001$, Figure 2A). No significant difference was found in peripheral eosinophils and basophils between age groups. The intrinsic AD was defined as a total IgE of <150 IU/mL and with no sensitization to allergens, whereas the extrinsic type

was defined as a total IgE of >150 IU/mL or a positive sensitization to allergens. We found that the proportion of extrinsic AD decreased with age. The extrinsic AD was significantly higher in the adolescent group than in the senior group (90.2% and 73.9%, respectively, $P = .029$, Figure 2B).

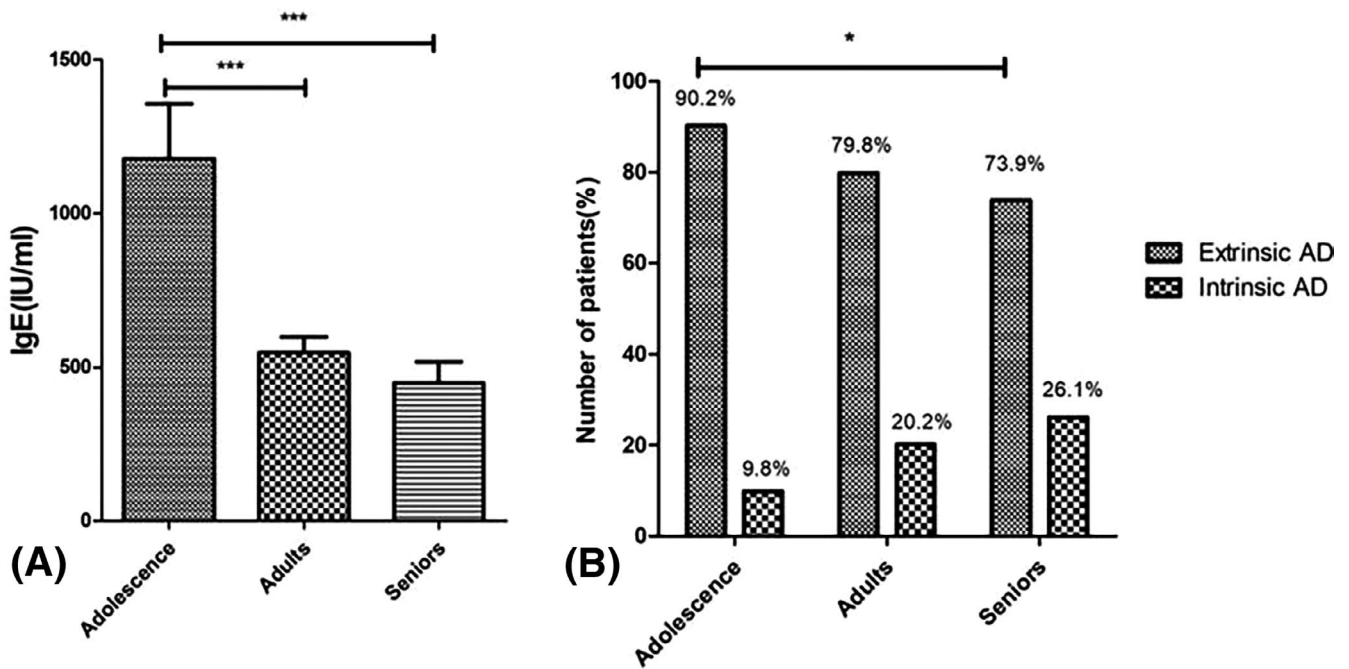


FIGURE 2 A, The mean value of total serum IgE in the adolescent group, adult group, and senior group was 1178.4, 546.8, and 449 IU/mL. Mann-Whitney U test was used. $***P < .001$. B, The proportion of extrinsic and intrinsic types according to different age groups. The intrinsic type was defined as a total IgE of <150 IU/mL and with no sensitization to allergens, whereas the extrinsic type was defined as a total IgE of >150 IU/mL or a positive sensitization to allergens. The age groups were defined as follows: adolescence >12 and ≤ 18 years old, adults >18 and ≤ 60 years old, and seniors >60 years old. Pearson chi-square test was used. $*P < .05$

TABLE 4 Clinical features in high-IgE patients and low-IgE patients

Variables	Serum total IgE		P value	χ^2
	≥ 100 IU/mL (n = 248)	<100 IU/mL (n = 148)		
Pruritus	242 (97.6%)	145 (98.0%)	.800	0.064
Flexural dermatitis	128 (51.6%)	53 (35.8%)	.002	9.327
Xerosis	204 (82.3%)	122 (82.4%)	1.000	0.002
Ichthyosis/palmar hyperlinearity/ keratosis pilaris	54 (21.8%)	24 (12.6%)	.178	1.810
Perifollicular accentuation	41 (16.5%)	24 (16.2%)	.935	0.007
immediate skin test reactivity, RAST test positive	79 (31.9%)	42 (28.4%)	.467	0.528
Tendency for cutaneous infections (especially <i>Staphylococcus aureus</i> and herpes simplex virus)	50 (20.2%)	27 (18.2%)	.641	0.218
Tendency to nonspecific hand/foot dermatitis	61 (24.6%)	44 (29.7%)	.263	1.253
Nipple eczema	20 (8.1%)	9 (6.1%)	.464	0.537
Cheilitis	48 (19.4%)	21 (14.2%)	.190	1.719
White dermographism	46 (18.5%)	14 (9.5%)	.023	7.505
Facial pallor/facial erythema	96 (38.7%)	50 (33.8%)	.326	0.966
Pityriasis alba	23 (9.3%)	6 (4.1%)	.054	3.721
Itch when sweating	143 (57.7%)	84 (56.8%)	.860	0.031
Intolerance to wool and lipid solvents	86 (34.7%)	50 (33.8%)	.856	0.033
Food hypersensitivity	91 (36.7%)	43 (29.1%)	.120	2.416
Recurrent conjunctivitis	36 (14.5%)	17 (11.5%)	.392	0.734
Dennie-Morgan infraorbital fold	27 (10.9%)	10 (6.8%)	.172	1.867
Orbital darkening	32 (12.9%)	7 (4.7%)	.008	6.974
Anterior neck folds	43 (17.3%)	10 (6.8%)	.003	8.953
Keratoconus	0 (0.0%)	0 (0.0%)	—	—
Anterior subcapsular cataracts	6 (2.4%)	11 (7.4%)	.017	5.669
Urticaria/angioedema	67 (27.0%)	32 (21.6%)	.230	1.439
Nummular eczema	65 (26.2%)	35 (23.6%)	.570	0.322
Periauricular eczema	78 (31.5%)	40 (27.0%)	.352	0.867
Eyelid eczema	58 (23.4%)	27 (18.2%)	.228	1.455
Scalp eczema/seborrheic dermatitis	90 (36.3%)	54 (36.5%)	.969	0.002
Perineum eczema	56 (22.6%)	33 (22.3%)	.948	0.004
Pompholyx of hand/foot	32 (12.9%)	25 (16.9%)	.274	1.197

Note: Pearson chi-square test was used.

3.4 | Association of serum total IgE and clinical features of AD

Table 4 shows the clinical features relevant to the diagnostic criteria and their association with serum total IgE levels. All the AD patients were divided into high serum IgE group (≥ 100 IU/mL) and low serum IgE group (<100 IU/mL). Flexural dermatitis (51.6% vs 35.8%, $P = .002$), anterior neck folds (17.3% vs 6.8%, $P = .003$), white dermographism (18.5% vs 9.5%, $P = .023$), and orbital darkening (12.9% vs 4.7%, $P = .008$) were significantly found more in high-IgE group

compared with low-IgE group. Furthermore, patients with high-IgE levels had higher personal atopic history than in those with normal IgE levels (64.5% vs 52.0%, $P = .014$, Table 5).

3.5 | Association of peripheral eosinophils and clinical features of AD

Table 6 shows the association of clinical features and peripheral eosinophils. A total of 146 patients had eosinophilia (>5%), while 250

Variables	Serum total IgE		P value	χ^2
	≥ 100 IU/mL (n = 248)	<100 IU/mL (n = 148)		
Eczema/AD before 2 years old	95 (38.3%)	44 (29.7%)	.084	2.993
Eczema/AD before 12 years old	101 (40.7%)	46 (31.1%)	3.694	0.055
Personal history of atopic diseases (Eczema, AD, asthma or allergic rhinitis)	160 (64.5%)	77 (52.0%)	.014	6.016
Family history of atopic diseases	161 (64.9%)	106 (71.6%)	.169	1.896

Note: Pearson chi-square test was used.

patients had normal peripheral eosinophils. We found that clinical features such as flexural dermatitis (53.4% vs 41.2%, $P = .021$), xerosis (89.0% vs 78.4%, $P = .009$), nipple eczema (11.0% vs 5.2%, $P = .045$), white dermographism (20.5% vs 12.0%, $P = .029$), facial pallor/facial erythema (44.5% vs 32.4%, $P = .018$), and anterior neck fold (19.2% vs 10.0%, $P = .014$) were seen more in patients with eosinophilia. No difference in personal and family atopic history was found between the two groups (57.5% vs 60.2% and 58.9% vs 72.4%, respectively, Table 7). In fact, patients with eosinophilia had a less frequent atopic personal and family history. History of periauricular eczema (38.4% vs 24.8%, $P = .006$), eyelid eczema (30.1% vs 16.4%, $P = .002$), and seborrheic dermatitis (45.2% vs 30.0%, $P < .001$) were found more frequently in patients with eosinophilia. No significant difference was found in clinical features and medical history in patients with or without basophilia (not shown in the table).

4 | DISCUSSION

Immunoglobulin E, eosinophils, and basophils are involved in the pathogenesis of atopic disease. IgE is a key molecule that can activate effector cells involved in allergic inflammation.⁸ In a case-control study of 2201 East German schoolchildren, elevated serum total IgE levels were found in 75% children with AD.⁹ In our study, 62.6% of patients with AD were found to have elevated serum IgE levels, whereas only 19.5% in non-AD group. Eosinophilia has been shown to be present in AD patients.¹⁰ Liu et al⁵ found that 31.8% of AD patients revealed eosinophilia (>5%). In our study, 36.9% of AD patients were found to have eosinophilia. However, the diagnostic importance of eosinophilia has been unclear since elevated blood eosinophils can be caused by other diseases such as parasite infection. And some AD patients may also have normal blood eosinophil levels. Basophils and mast cells are effector cells in an IgE-dependent type I hypersensitivity reaction in skin inflammation.¹¹ In our study, we also found basophilia was more prevalent in AD patients than in non-AD patients, which was rarely discussed in previous studies. The sensitization of aeroallergens was seen more in AD patients. However, no significant differences were found for food allergen sensitization between AD and non-AD patients for adults in the present study. Previous studies have produced similar results. It was indicated¹² that sensitization was predominate to foods initially, but then shifted toward inhalant allergens, as the AD patients became older. The occurrence of food allergy

TABLE 5 Medical history in high-IgE patients and low-IgE patients

such as cow milk, egg, and peanut in children was called "Class I food allergy." The prevalence of "Class I food allergy" decreased in adolescence and adult AD patients but the sensitization of aeroallergens remained unchanged.¹³ It was predicted that the prevalence of "Class I food allergy" decreased due to their "self-care behavior" as the increase with age. Adult patients might pay more attention to avoiding unconscious exposure to suspicious food allergens and thus avoiding the elevation of sIgE of allergens. However, it still needs to be further researched.

For the relationship between laboratory tests and the severity of AD, Kiiski et al¹⁴ reported that the total serum IgE level and specific IgE were significantly associated with the severity of AD. Clinical features of 5000 patients with AD in South Korea were studied, and a significant correlation between increased EASI scores and the total serum IgE levels was found.⁷ Our studies have produced similar results. We also found that eosinophils count was associated with the severity of AD. We found that basophil count in peripheral blood of severe AD patients was significantly higher than in mild AD patients (0.698% vs 0.591%, $P < .05$). The relationship between peripheral basophils and severity of AD is rarely discussed. Eosinophils contain granules that contain chemical mediators including major basic protein-1 (MBP-1), MBP-2, eosinophil-derived neurotoxin (EDN), and eosinophil cationic protein (ECP).¹⁵ They can induce tissue damage of target tissue and stimulate the recruitment of basophils, mast cells, and neutrophils. Activated eosinophils can produce a large number of cytokines and chemokines, such as IL-16, IL-12, TGF- β 1, and IL-13, and play an important immunoregulatory role in the pathogenesis of AD.¹⁶

We also found that elevated serum total IgE level was less seen in elderly patients than in adolescent and adult patients ($P < .01$). Somani et al⁸ reported that 88% of the AD patients have elevated serum IgE levels with the highest IgE elevation seen in those between 10 and 20 years old.¹⁷ That is similar to our results. AD may be classified into two different types, extrinsic type and intrinsic type, which are also called allergic and non-allergic types. The intrinsic type, or non-allergic type, shows normal IgE levels in addition to the absence of any sensitization to allergens, whereas the extrinsic or allergic type has increased specific IgE levels with sensitization to specific allergens. The overall proportion of extrinsic type and intrinsic type were 79.8% and 20.2% in our patients, which was similar to that reported from Netherland¹⁸ (78.2% vs 21.8%) and Korea¹⁹ (80% vs 20%). Besides that, we also found that the number of AD patients with intrinsic type

increased with age, reaching 9.8% in adolescence and 26.1% in seniors. In the previous study, children with AD were reported to have a high prevalence (about 80%) of sensitization to foods and a medium prevalence (about 40%) to aeroallergens.¹¹ However, in our research sIgE prevalence in AD patients was 39.8% for aeroallergens and only 15.9% for food allergens, so food sensitization in adult patients was much lower than in children patients with AD. So it was predicted that sensitization to food allergens decreases with age, leading to the decrease in the proportion of the intrinsic types of AD.

We also found a significant difference in personal atopic history between high-IgE group and low-IgE group. Celakovska et al²⁰ found that AD patients with elevated serum total IgE and food allergy suffer significantly more often from allergic rhinitis, bronchial asthma, and persistent eczematous lesions and have more positive personal or family history of atopic diseases. It indicates that serum total IgE as a key effector involved in allergic sensitization, contributing to the aggregation of several atopic diseases, thus contributing to high prevalence of personal atopic history.

TABLE 6 Clinical features in patients with or without eosinophilia

Variables	EOS%		P value	χ^2
	≥5% (n = 146)	<5% (n = 250)		
Pruritus	145 (99.3%)	242 (98.6%)	.163	2.625
Flexural dermatitis	78 (53.4%)	103 (41.2%)	.021	5.551
Xerosis	130 (89.0%)	196 (78.4%)	.009	7.172
Ichthyosis/palmar hyperlinearity/keratosis pilaris	24 (16.4%)	54 (21.6%)	.240	1.553
Perifollicular accentuation	22 (15.1%)	43 (17.2%)	.674	0.305
immediate skin test reactivity, RAST test positive	47 (32.2%)	74 (29.6%)	.651	0.292
Tendency for cutaneous infections (especially <i>Staphylococcus aureus</i> and herpes simplex virus)	26 (17.8%)	51 (20.4%)	.599	0.395
Tendency to nonspecific hand/foot dermatitis	34 (23.3%)	71 (28.4%)	.290	1.236
Nipple eczema	16 (11.0%)	13 (5.2%)	.045	4.504
Cheilitis	26 (17.8%)	43 (17.2%)	.891	0.024
White dermographism	30 (20.5%)	30 (12.0%)	.029	5.239
Facial pallor/facial erythema	65 (44.5%)	81 (32.4%)	.018	5.818
Pityriasis alba	14 (9.6%)	15 (6.0%)	.230	1.749
Itch when sweating	87 (59.6%)	140 (56.0%)	.528	0.485
Intolerance to wool and lipid solvents	57 (30.0%)	79 (31.0%)	.154	2.263
Food hypersensitivity	53 (36.3%)	81 (32.4%)	.443	0.627
Recurrent conjunctivitis	22 (15.1%)	31 (12.4%)	.449	0.556
Dennie-Morgan infraorbital fold	18 (12.3%)	19 (17.6%)	.151	2.433
Orbital darkening	18 (12.3%)	21 (8.4%)	.223	1.602
Anterior neck folds	28 (19.2%)	25 (10.0%)	.014	6.698
Keratoconus	0 (0.0%)	0 (0.0%)	—	—
Anterior subcapsular cataracts	6 (4.1%)	11 (4.4%)	1.000	0.019
Urticaria/angioedema	28 (19.2%)	64 (25.6%)	.205	3.165
Nummular eczema	37 (25.3%)	58 (23.2%)	.877	0.263
Periauricular eczema	56 (38.4%)	62 (24.8%)	.006	8.097
Eyelid eczema	44 (30.1%)	41 (16.4%)	.002	10.318
Scalp eczema/seborrheic dermatitis	66 (45.2%)	75 (30.0%)	.000	15.437
Perineum eczema	39 (26.7%)	47 (18.8%)	.094	4.739
Pompholyx of hand/foot	20 (13.7%)	34 (13.6%)	.561	1.157

Note: Pearson chi-square test was used.

Variables	EOS count%		P value	χ^2
	≥5%(n = 146)	<5%(n = 250)		
Eczema/AD before 2 years old	49 (33.6%)	90 (36.0%)	.663	0.241
Eczema/AD before 12 years old	55 (37.7%)	92 (36.8%)	.914	0.030
Personal history of atopic diseases (Eczema, AD, asthma, or allergic rhinitis)	84 (57.5%)	153 (61.2%)	.524	0.515
Family history of atopic diseases	86 (58.9%)	181 (72.4%)	.008	7.643

Note: Pearson chi-square test was used.

In our study, the clinical differences in AD patients grouped by serum total IgE level and eosinophils count were also investigated. Flexural dermatitis, anterior neck folds, white dermographism, and orbital darkening were seen more in AD patients with elevated IgE. We also found that flexural dermatitis, xerosis, nipple eczema, white dermographism, facial pallor/facial erythema, anterior neck fold, periauricular eczema, eyelid eczema, and seborrheic dermatitis were seen more in patients with eosinophilia in our research. Direct evidence supporting the relationship between serum total IgE, eosinophil, and clinical features was scarce. However, EDN and ECP, released by activated eosinophils, were reported to play a role in the clinical phenotypes of AD. Kim et al²¹ found that EDN and ECD not only reflected the severity of AD, but also correlate with AD-associated minor clinical features such as asthma, cheilitis, NS-HFD (non-specific hand or foot dermatitis), and scalp scale. The dysfunction of protein makes the skin more permeable to environmental allergens, which contribute to relevant symptoms such as xerosis, facial erythema, and anterior neck fold. In conclusion, we found that serum total IgE level and peripheral eosinophils count were seen more in AD patients and they correlated with clinical features of AD.

In conclusion, we considered that the serum total IgE, allergen-specific IgE, blood eosinophil count, EOS%, and BAS% can be used in the diagnosis of AD. The serum total IgE, blood eosinophil count, and EOS% are related to the severity of disease. We also found a correlation in laboratory tests, medical history, and clinical features in AD patients. This analysis can lead to a better understanding of the clinical application of laboratory tests in AD.

CONFLICT OF INTEREST

The authors have no conflict of interest to declare.

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TABLE 7 Medical history in patients with or without eosinophilia

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