

Could mean platelet volume be used as a marker for activity and severity index of alopecia areata?

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ABSTRACT

Background. We aimed to determine whether MPV can be used as a marker for the activity and the severity index of alopecia areata (AA).

Method. The charts of 71 children who received a diagnosis of AA and 70 age and gender-matched healthy children were retrospectively evaluated. The severity of hair loss was classified as S1 (<25%), S2 (25-49%), S3 (50-74%), S4 (75-95%), S4b (96-99%) (according to the percent of the area involved), alopecia totalis (AT), and alopecia universalis (AU). In the laboratory tests, the results of the complete blood count, anti-nuclear antibody (ANA), thyroid function tests (TSH, free/total T4, free/total T3), and autoimmune thyroid antibodies [anti-thyroid peroxidase antibody (anti-TPO) and anti-thyroglobulin antibody (AT)] were recorded.

Results. A total of 141 cases including 61 (43.3%) males and 80 (56.7%) females were included. There was no statistically significant difference between the groups according to the mean age ($p>0.05$).

The MPV measurements were statistically significantly higher in the AA group ($p<0.01$). There was no statistically significant difference between the types of AA according to the mean age, gender distribution, the presence of nail involvement, the presence of family history, and the presence of autoimmune disease ($p>0.05$). There was no statistically significant difference between the severity of AA according to the mean age, gender distribution, the presence of nail involvement, the presence of family history, and the presence of autoimmune disease ($p>0.05$).

Conclusion. MPV is helpful in assessing clinical activity in patients with AA. However, prospective studies involving more patients are needed to support our findings.

Key words: alopecia areata, mean platelet volume, inflammation.

Alopecia areata (AA) is a chronic autoinflammatory disease that is characterized by episodic and non-scarring loss of the scalp, eyebrows, eyelashes and other body hairs and by sharply demarcated patches.¹ Although its etiopathogenesis remains unclear, the autoimmune reactions, which target the hair matrix, are thought to be responsible for genetically susceptible individuals.² The lifetime risk of AA is 1.7-2%, regardless of age and sex.^{3,4} AA is seen especially in young

patients and approximately 60% of patients have their first attack before the age of 20.⁵ AA can also be accompanied by other autoimmune diseases such as autoimmune thyroid diseases, vitiligo, and lupus erythematosus.⁶ Mean platelet volume (MPV) is used as part of the complete blood count in full blood count analyzers and is a common marker used to demonstrate platelet function and activation.⁷ It has been shown in recent years that MPV is high in chronic inflammatory diseases and may be used as an inflammatory marker.⁸ In this study, we examined the relationship between MPV level and disease severity, nail involvement, family history, disease duration and other accompanying autoimmune diseases in pediatric patients with AA.

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Material and Methods

After receiving the ethical board approval (no:10, in 19.10.2015) of Istanbul Anatolia–North Region Public Hospitals Trust, 71 children under the age of 18 who received a diagnosis of AA at the Dermatology outpatient clinic of Beykoz State Hospital between 2015 and 2017 were retrospectively evaluated in terms of age, sex, disease severity, lesion distribution, nail findings, family history, and accompanying autoimmune diseases. The control group included 70 age and gender-matched healthy children who had no dermatologic or systemic disease. If the duration of AA was 3 months or less, it was considered acute. If the duration of AA was over 3 months, it was considered chronic. The severity of hair loss was classified as S1 (<25%), S2 (25-49%), S3 (50-74%), S4 (75-95%), S4b (96-99%) (according to the percent of the area involved), alopecia totalis (AT), and alopecia universalis (AU). In the laboratory tests, the results of the complete blood count, anti-nuclear antibody (ANA), thyroid function tests (TSH, free/total T4, free/total T3), and autoimmune thyroid antibodies [anti-thyroid peroxidase antibody (anti-TPO) and anti-thyroglobulin antibody (AT)] were recorded.

Statistical Analysis

NCSS-2007 software program (Number Cruncher Statistical System) (Kaysville, Utah, USA) was used for statistical analysis. Mean, standard deviation, median, frequency, ratio, minimum, and maximum were used for comparing the descriptive data. Moreover, the Mann Whitney U test was used for comparing the parameters without normal distribution

between two groups. The Kruskal Wallis test was used for comparing three or more groups without normal distribution. The Fisher-Freeman-Halton test, Fisher's Exact test, and Perason chi-square test were used for comparing the qualitative data. The significance level was considered as $p < 0.05$.

Results

A total of 141 cases including 43.3% (n=61) males and 56.7% (n=80) females who were admitted to the Dermatology outpatient clinic of Beykoz State Hospital between 2015 and 2017 were enrolled in the study. Their ages ranged from 1 to 18 years and the mean age was 12.57 ± 4.38 years.

In the AA group, their ages ranged from 1 to 18 years and the mean age was 12.24 ± 4.69 years. In the control group, their ages ranged from 2 to 18 years and the mean age was 12.91 ± 4.05 years. There was no statistically significant difference between the groups according to the mean age ($p > 0.05$) (Table I).

In the AA group, the MPV measurements ranged from 6.4 to 11.4 and the mean MPV level was 8.33 ± 1.15 . In the control group, the MPV measurements ranged from 5.4 to 7.7 and the mean MPV level was 6.33 ± 0.41 . The MPV measurements were statistically significantly higher in the AA group ($p = 0.001$; $p < 0.01$) (Table I), (Fig. 1).

The MPV measurements showed no statistically significant differences according to the type of AA ($p = 0.344$; $p > 0.05$) (Table II). While the MPV measurements showed no statistically

Table I. Age and MPV evaluations according to groups.

		Group		p ^a
		Alopeci areata (n=71)	Control (n=70)	
Age	Min-Max (Median)	1-18 (13)	2-18 (14)	0.551
	Mean±SD	12.24±4.69	12.91±4.05	
MPV	Min-Max (Median)	6.4-11.4 (7.9)	5.4-7.7 (6.4)	0.001**
	Mean±SD	8.33±1.15	6.33±0.41	

^aMann Whitney U Test, ** $p < 0,01$
MPV: Mean platelet volume

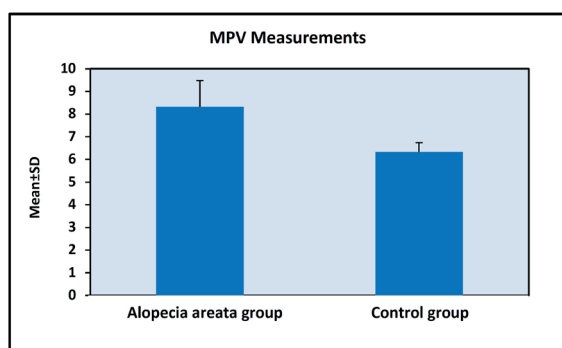


Fig. 1. Distribution of MPV measurements according to groups.

significant difference according to the severity of AA ($p>0.05$) (Table II), it was remarkable that the MPV measurements were high in the patients with AA severity S2.

The MPV measurements showed no statistically significant difference according to the presence of nail involvement, the presence of family history, the presence of autoimmune disease, and disease duration ($p>0.05$) (Table II).

There was no statistically significant difference between the types of AA according to the mean age and gender distribution ($p>0.05$) (Table III).

There was no statistically significant difference between the types of AA according to the presence of nail involvement, the presence of family history, and the presence of autoimmune disease ($p>0.05$) (Table III). It was remarkable that the patients who had a reticular type of AA had a higher presence of nail involvement than patients who had a siasphio or a ophiasis type of AA (Table III).

There was no statistically significant difference between the severity of AA according to the mean age and gender distribution ($p>0.05$) (Table IV).

There was no statistically significant difference between the severity of AA according to the presence of nail involvement, the presence of family history, and the presence of autoimmune disease ($p>0.05$) (Table IV).

Discussion

AA is a chronic inflammatory disease that is characterized by episodic and non-scarring loss of the scalp and/or other hairy areas of the body.⁹ Although genetic predisposition, non-specific immune reactions, organ-specific autoimmune reactions, environmental factors, and atopy

Table II. Evaluations of descriptive characteristics according to MPV measurements.

		MPV		P
		Min-Max (Median)	Mean±SD	
Type of alopecia areata	Reticular	6.4-11.4 (8.4)	8.57±1.25	^b 0.344
	Ophiasis	6.8-10.2 (7.7)	8.26±1.16	
	Siasphio	6.8-10.5 (7.7)	8.07±0.99	
Severity of alopecia areata	S1	6.8-10.5 (7.7)	8.05±1.10	^a 0.085
	S2	6.4-11.4 (8.5)	8.52±1.16	
Nail involvement	No	6.4-11.4 (7.9)	8.39±1.19	^a 0.522
	Yes	6.8-9.9 (8.2)	8.12±1.03	
Family history	No	6.4-11.4 (7.9)	8.36±1.19	^a 0.642
	Yes	7.2-9.4 (8)	8.06±0.76	
Autoimmune disease	No	6.4-11.4 (8)	8.30±1.13	^a 0.655
	Yes	6.8-10.9 (7.9)	8.49±1.29	
Disease duration	Acute hair loss	6.8-10.5 (8.5)	8.47±1.07	^a 0.166
	Chronic hair loss	6.4-11.4 (7.7)	8.16±1.25	

^aMann Whitney U Test, ^bKruskall Wallis Test
MPV: Mean platelet volume

Table III. Evaluation of descriptive characteristics according to type of alopecia areata.

		Type of Alopecia areata			P
		Reticular (n=31)	Ophiasis (n=17)	Siasphio (n=23)	
Age	Min-Max (Median)	4-18 (13)	1-18 (14)	4-17 (13)	^b 0.716
	Mean±SD	12.48±4.90	11.41±5.46	12.52±3.87	
Gender; n (%)	Male	17 (54.8)	5 (29.4)	10 (45.3)	^d 0.253
	Female	14 (45.2)	12 (70.6)	13 (56.5)	
Nail involvement; n (%)	No	28 (90.3)	11 (64.7)	16 (69.6)	^c 0.060
	Yes	3 (9.7)	6 (35.3)	7 (30.4)	
Family history; n (%)	No	27 (87.1)	17 (100)	21 (91.3)	^c 0.479
	Yes	4 (12.9)	0 (0)	2 (8.7)	
Autoimmune disease; n (%)	No	23 (74.2)	16 (94.1)	19 (82.6)	^c 0.260
	Yes	8 (25.8)	1 (5.9)	4 (17.4)	

^bKruskall Wallis Test, ^cFisher Freeman Halton Test, ^dPearson Chi-Square Test

Table IV. Evaluation of descriptive characteristics according to severity of alopecia areata.

		Severity of Alopecia areata		P
		S1 (n=29)	S2 (n=42)	
Age	Min-Max (Median)	1-18 (13)	4-18 (15)	^a 0.167
	Mean±SD	11.28±5.01	12.90±4.40	
Gender; n (%)	Male	12 (41.4)	20 (47.6)	^d 0.603
	Female	17 (58.6)	22 (52.4)	
Nail involvement; n (%)	No	22 (75.9)	33 (78.6)	^d 0.788
	Yes	7 (24.1)	9 (21.4)	
Family history; n (%)	No	28 (96.6)	37 (88.1)	^c 0.390
	Yes	1 (3.4)	5 (11.9)	
Autoimmune disease; n (%)	No	25 (86.2)	33 (78.6)	^d 0.414
	Yes	4 (13.8)	9 (21.4)	

^aMann Whitney U Test, ^cPearson Chi-Square Test, ^dFisher's Exact Test

have been emphasized, the etiopathogenesis of AA has not been fully elucidated.⁹ AA is seen especially in young patients and also approximately 60% of patients have had their first attack before the age of 20.² Although AA can be diagnosed easily by clinical examination, diffuse AA may be diagnostic challenging. In this situation, the dermoscopic findings (broken hair, black dot, yellow dot, exclamation hair) facilitate the differential diagnosis; however, histopathological diagnosis may be needed in a small number of cases. The characteristic histopathologic finding of AA is an intense peribulbar and intrabulbar lymphocytic infiltrate of lymphocytes around the anagen

follicles in the appearance of a honey bee colony.⁹

In addition to homeostatic functions, thrombocytes interact with endothelial cells, leukocytes (monocytes, neutrophils, dendritic cells, T-cells), and progenitor cells and allow inflammatory cells to migrate to lesion sites and release abundant quantities of inflammatory cytokine and thus provide an inflammatory environment in the lesion area. It has been reported in recent years that MPV also reflects platelet function and activation and may be a marker of inflammation in different chronic diseases.⁸

In our study, the mean MPV levels of the patients with AA were significantly higher than those of the control group. The current hypotheses on the development of AA focuses on the deterioration of autoantigen presentation in the hair follicle as a result of immune privilege in the hair follicles and activation of autoreactive T lymphocytes.¹⁰ The autoreactive T-cell response, which is thought to be involved in the pathogenesis of AA, can be genetically determined.¹⁰ It can also be increased by IL-6 and other cytokines.¹¹ Although IL-6 is a proinflammatory cytokine produced by various cells such as fibroblasts, macrophages, B and T cells, it plays an important role in the hematopoiesis, immune cell activation, regulation of inflammation, and pathogenesis of autoimmune diseases.^{12,13} Inflammation is known to be an important stimulus for platelets. However, it has been suggested that increased levels of IL-6 may stimulate platelet production and release large platelets from bone marrow.¹³ Therefore, in our study, the mean MPV levels, which were significantly higher in the patients with AA than in the control group, may be thought to be associated with an increase in young platelets in circulation. When the literature is examined, it is seen that MPV levels vary depending on the severity of systemic inflammation in high- and low-grade inflammatory diseases, but the results are contradictory.¹⁴

When the relationship between the mean MPV level and the age of disease onset, disease duration, disease severity, family history, nail involvement, and accompanying autoimmune disease was evaluated, there was no difference between the AA and control groups. While the MPV measurements showed no statistically significant difference according to the severity of AA, it was remarkable that the MPV measurements were high in the patients with AA severity S2. When the mean MPV level was examined according to the type of AA, it was seen that the mean MPV level was a little bit higher in the reticular pattern, which is more common in the population.

In this study, although there was no significant relationship between the mean MPV level and the clinical activity markers of the patient group (early onset, family history, ophiasis pattern, nail involvement), we think that MPV can be used as a useful marker in the evaluation of AA.

Consequently, AA is a severe and chronic disease in the pediatric age group. Families often refer to a doctor for medical support and treatment because the disease has social and psychological effects. MPV is a simple marker that does not require advanced or expensive technology and also MPV level is determined in each complete blood count. MPV is helpful in assessing clinical activity at a glance in patients with AA. However, prospective studies involving more patients are needed to support our findings.

Ethical approval

Ethical board approval was received from the Istanbul Anatolia-North Region Public Hospitals Trust (no:10, in 19.10.2015).

Author contribution

The authors confirm contribution to the paper as follows: study conception and design: NDA; data collection: NDA; analysis and interpretation of results: NDA; draft manuscript preparation: NDA.

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Conflict of interest

The authors declare that there is no conflict of interest.

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