

## Blood Cell Indices in Ankylosing Spondylitis and Psoriatic Arthritis with Relation to the Disease Activity: A Cross-Sectional Study

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### ABSTRACT

**Background:** Red blood cell distribution width (RDW) and mean platelet volume (MPV) are influenced by autoimmune diseases as RA.

**Objective:** This study aimed to assess RDW, MPV levels and different hematological indices in ankylosing spondylitis (AS) and psoriatic arthritis (PsA) to investigate their relations with disease activity.

**Methods:** A total of 120 patients (60 AS and 60 PsA) were enrolled. Demographic data, disease activity scores, complete blood count (CBC), erythrocyte sedimentation rate (ESR) and C-reactive protein (CRP) levels were recorded. RDW, MPV and various ratios were measured.

**Results:** Patients mean age was  $44.750 \pm 12.187$  years were elevated in PsA and AS patients. RDW and MPV were significantly higher in patients with active PsA patients ( $14.495 \pm 1.9$  and  $8.903 \pm 1.166$ fl) than in inactive ones ( $11.975 \pm 0.5$ fl and  $8.220 \pm 0.5$  fl, p value  $<0.001$ ,  $0.014$  respectively). While in AS active patients RDW was elevated in patients with high bath ankylosing Spondylitis disease activity index (BASDAI) score. RDW% was positively correlated to disease activity among PsA and AS patients ( $r=0.614$  p  $<0.001$ ) and ( $r=0.567$  p  $<0.001$ ) respectively. In AS patients, RDW was at a criterion of 12.4, with sensitivity of 83.78% and specificity of 69.5% (p value  $< 0.001$ ), while in PsA patients' RDW showed good metrics [p value  $< 0.001$ ; Cutoff:13.2; sensitivity: 72.5%; specificity: 100%].

**Conclusion:** RDW and MPV were potentially elevated in active psoriatic patients and RDW were positively correlated with activity score of AS and PsA, which could be possible important index for disease activity.

**Key words:** Ankylosing spondylitis, Psoriatic arthritis, Mean platelet volume, Red cell distribution width.

### INTRODUCTION

Spondyloarthropathy (SPA) are immune-mediated disease characterized by systemic inflammation with functional affection of various organs including joints and others<sup>(1)</sup>. Chronic inflammatory arthritis is the predominate involvement of ankylosing spondylitis (AS) & psoriatic arthritis (PsA)<sup>(2)</sup>.

These patients usually alternate activity with remission state along the disease course. Of determinant of disease activity and inflammatory milieu are the Disease Activity Score and usual laboratory markers like erythrocyte sedimentation rate (ESR) and C-reactive protein (CRP)<sup>(3)</sup>.

A mass activation of the immune system leads to the autoantibodies overproduction, immune complexes, and inflammatory cytokines promotion, which interplay to produce eventual disease commencement that can affect different peripheral blood cells and their indices as activated platelets can trigger sequences of physiological and pathological reactions, liberating different platelet-derived proteins, chemokines and growth factors, which might result into immune-inflammatory disorders<sup>(4-6)</sup>. Through that, they react with neutrophils allowing the release of inflammatory cytokines.

Mounting evidence that mean platelet volume (MPV), red blood cell distribution width (RDW)<sup>(7)</sup>, and other parameters (e.g., neutrophil-to-lymphocyte ratio

[NLR]) are illustrated as reliable, accurate bio inflammatory markers in various autoimmune diseases<sup>(8)</sup>. Moreover, red blood cell distribution width-to-platelet ratio (RPR) has been presented as a surrogate and rapid index to predict morbidity and mortality in liver cirrhosis and acute pancreatitis.

The relations between blood indices and disease activity in spondyloarthropathy cases have not been widely investigated<sup>(9-10)</sup> yet the impact of those hemoparameters and ratios not extensively investigated among spondyloarthropathies that may give hand to structured model of different clinical and laboratory categories that may be used to assess disease activity and damage index in such diseases. Accordingly, this study aimed to assess the association between blood cell indices with the disease activity in AS and PsA, and to determine whether those indices can be used as a predictor of disease activity.

### MATERIALS AND METHODS

**Study design:** Observational study (cross-sectional) study. A convenient sample of patients were recruited from Outpatient Clinic of Internal Medicine Department and Rheumatology Department at Ain Shams and Kasr-Alainy hospitals.

**Study setting and sampling:** The study was conducted on 60 AS patients, 60 PsA patients. All AS patients were diagnosed according to the Assessment of Spondylarthritis International Society classification criteria for spondylarthritis <sup>(12)</sup> and PsA in line with Moll and Wright criteria for psoriatic arthritis <sup>(11)</sup>.

**Exclusion criteria:**

Anyone who had other inflammatory, autoimmune diseases, infectious diseases, malignancies, diabetes mellitus, or liver/kidney diseases.

**Calculation of sample size:**

Based on evidence from prior study done on autoimmune disease. Epi-calc 2000 was used to calculate the sample size of this cross-sectional study. Assuming 80% power, 0.05 level of significance, 60% proportion of exposed individuals, to detect odds ratio OR= 2.5 Sample size will be = 110 participants). Considering drop-outs rate of 10%, therefore the final sample size was 120 participants.

**Methodology**

All patients were subjected to full medical history and clinical examination. Disease activity was assessed using Ankylosing Spondylitis Disease Activity Score (ASDAS) <sup>(13)</sup>, where patients with ASDAS scores < 1.3 were considered as inactive and those >1.3 were considered as active patients. Whilst, psoriatic disease activity was evaluated by Disease Activity in Psoriatic Arthritis (DAPSA) Score <sup>(14)</sup>. Patients with score 0-4 were considered inactive while those with score > 4 were considered active patients.

**Laboratory investigations:** Blood samples were drawn and chemically analyzed within 2 hrs from time of collection. Complete blood cell (CBC) analysis was done by the Coulter Counter mode including total white blood cell (WBC), with differential of neutrophil and lymphocyte count, RDW%, MCV, MCH and platelet counts, MPV and PDW.

**Statistical analysis**

-Data were coded and entered using the statistical package SPSS version 24. For quantitative variables comparisons, nonparametric Kruskal-Wallis and Mann-Whitney tests were used. While, chi-square ( $\chi^2$ ) test was performed for categorical data.

**Ethical consideration:**

The institutional ethical committee (Ain Shams University Hospitals) approved the study. An informed consents were taken from the patients with explaining the objectives and the methods of this study. Confidentiality was guaranteed on handling the data conformed to the revised biomedical ethics of Helsinki declaration.

**RESULTS**

In 60 PsA patients, average age in years was 44.750 ± 12.187, 38 were females (63.33%), 22 were males (36.67%), 20(33.33%) patients had inactive disease while 40 (66.67%) were active (Table 1). Among 60 AS patients the mean age was 39.600 ± 12.048 years, 41 (68.33%) patients were males while 19 (31.67%) were females. Patients with active disease were 37 (61.67%) while 23 patients were inactive (38.33%) as shown in table (2).

**Table (1):** Descriptive data of PSA patients

PSA			
Age (Years)	Median	46	
	IQR	38	53
DD (Years)	Median	10	
	IQR	6	15
		N	%
Sex	Male	22	36.67
	Female	38	63.33
Peripheral affection	Yes	48	80
	No	12	20
Axial affection	Yes	43	71.67
	No	17	28.33
No Tender j	Median	18.5	
	IQR	11	34
No Swollen j	Median	3	
	IQR	2	5
DAPSA score	Median	21.8	
	IQR	4	32
Disease Activity	Remission	20	33.33
	Mild	0	0
	Moderate	18	30
	Severe	22	36.67
Disease Activity	Inactive	20	33.33
	Active	40	66.67
WBCS (10 <sup>9</sup> /L)	Mean ±SD	6.558 ±	1.131
Neutrophils (10 <sup>9</sup> /L)	Mean ±SD	4.148 ±	1.58
Lymphocytes (10 <sup>9</sup> /L)	Mean ±SD	1.993 ±	0.3
HCT (%)	Mean ±SD	37.088 ±	1.567
RBCs (10 <sup>12</sup> /L)	Mean ±SD	4.793 ±	0.532
Hb (g/L)	Mean ±SD	12.6 ±	1.102
MCV (fL)	Mean ±SD	80.235 ±	7.156
MCH (pg)	Mean ±SD	24.933 ±	3.973
RDW%	Mean ±SD	13.655 ±	2.015
PLT(10 <sup>9</sup> /L)	Mean ±SD	247.833 ±	7.786
MPV/fl	Mean ±SD	8.675 ±	1.027
PDW%	Mean ±SD	11.945 ±	2.513
Hb/PLT ratio	Mean ±SD	0.055 ±	0.017
RDW%/PLT ratio	Mean ±SD	0.059 ±	0.018
RBCs/PLT ratio	Mean ±SD	0.021 ±	0.006
Hb/RDW% ratio	Mean ±SD	0.941 ±	0.153
NLR ratio	Mean ±SD	2.091 ±	0.491
PLR ratio	Mean ±SD	133.302 ±	8.824
ESR (mm/h)	Mean ±SD	60.475 ±	7.728
CRP (mg/L)	Mean ±SD	14.667 ±	7.097
HLA B27	Positive	43	71.67
	Negative	17	28.33

SD: Standard Deviation, No: number, %: percentage, mm/h: millimeter per hour, mg/L: milligram per Liter, IU/ml: International unit per milliliter, U/L: Unit per liter, mg/dL: milligram per Deci Liter WBCs: White blood cells, HCT: Hematocrit, RBCs: Red blood cells, Hb: Hemoglobin, MCV: Mean corpuscular volume, MCH: Mean corpuscular Hemoglobin, RDW: Red cell distribution width, PLT: Platelets, MPV: Mean platelet volume, PDW: Platelet distribution width, Hb/PLT ratio: Hemoglobin/Platelet ratio, RDW/PLT ratio: Red cell distribution width/ Platelet ratio, RBCs/PLT ratio: Red blood cells/Platelet ratio, Hb/RDW ratio: Hemoglobin/ Red cell distribution width ratio, NLR: Neutrophil/Lymphocyte ratio, PLR: Platelet/Lymphocyte ratio, /mm<sup>3</sup>: Per cubic millimeter, %: percentage, g/dL: Gram per deci Liter, fl: femtoliter, pg: picogram, t: independent sample t test, P: probability value, \*: p<0.05 is statistically significant, \*\*: p ≤ 0.001 is statistically highly significant.

Median (IQR) for non-parametric data , Mean±SD for parametric data

**Table (2):** Descriptive data of AS patients

AS			
Age (Years)	Median	41	
	IQR	33	46.5
DD (Years)	Median	6.5	
	IQR	3	10
		N	%
Sex	Male	41	68.33
	Female	19	31.67
Peripheral	Yes	38	63.33
	No	22	36.67
Axial	Yes	60	100
	No	0	0
Tender j	Median	5	
	IQR	2.5	7
Swollen j	Median	2	
	IQR	1	3
ASDAS score	Median	2.2	
	IQR	1.2	3.5
Disease Activity	Remission	23	38.33
	Mild	0	0
	Moderate	7	11.67
	Severe	13	21.67
	Very severe	17	28.33
Disease Activity	Inactive	23	38.33
	Active	37	61.67
WBCS (10 <sup>9</sup> /L)	Mean ±SD	6.853	1.737
Neutrophils (10 <sup>9</sup> /L)	Mean ±SD	3.988	1.311
Lymphocytes (10 <sup>9</sup> /L)	Mean ±SD	2.443	0.792
HCT (%)	Mean ±SD	38.14	3.213
RBCs (10 <sup>12</sup> /L)	Mean ±SD	4.915	0.644
Hb (g/L)	Mean ±SD	12.978	1.639
MCV (fL)	Mean ±SD	78.268	7.214
MCH (pg)	Mean ±SD	25.397	3.914
RDW%	Mean ±SD	13.232	1.6
PLT (10 <sup>9</sup> /L)	Mean ±SD	309.65	21.259
MPV/fl	Mean ±SD	9.695	2.322
PDW%	Mean ±SD	11.457	2.739
Hb/PLT ratio	Mean ±SD	0.048	0.018
RDW%/PLT ratio	Mean ±SD	0.048	0.016
RBCs/PLT ratio	Mean ±SD	0.018	0.007
Hb/RDW% ratio	Mean ±SD	0.997	0.179
NLR ratio	Mean ±SD	1.786	0.4
PLR ratio	Mean ±SD	139.872	6.929
ESR (mm/h)	Mean ±SD	38.383	4.654
CRP (mg/L)	Mean ±SD	9.95	2.819
HLA B27	Positive	37	61.67
	Negative	23	38.33

SD: Standard Deviation, No: number, %: percentage, mm/h: millimeter per hour, mg/L: milligram per Liter, IU/ml: International unit per milliliter, U/L: Unit per liter, mg/dL: milligram per Deci Liter WBCs: White blood cells, HCT: Hematocrit, RBCs: Red blood cells, Hb: Hemoglobin, MCV: Mean corpuscular volume, MCH: Mean corpuscular Hemoglobin, RDW: Red cell distribution width, PLT: Platelets, MPV: Mean platelet volume, PDW: Platelet distribution width, Hb/PLT ratio: Hemoglobin/Platelet ratio, RDW/PLT ratio: Red cell distribution width/ Platelet ratio, RBCs/PLT ratio: Red blood cells/Platelet ratio, Hb/RDW ratio: Hemoglobin/ Red cell distribution width ratio, NLR: Neutrophil/Lymphocyte ratio, PLR: Platelet/Lymphocyte ratio, /mm<sup>3</sup>: Per cubic millimeter, %: percentage, g/dL: Gram per deci Liter, fl: femtoliter, pg: picogram, t: independent sample t test, P: probability value, \*: p<0.05 is statistically significant, \*\*: p≤0.001 is statistically highly significant. Median (IQR) for non-parametric data , Mean±SD for parametric data

Patients then were divided into active group and inactive group, comparison between both groups was done regarding the different blood cell indices. On comparing the different blood cell indices in active and non-active psoriatic patients, we demonstrated significantly higher RDW% ( $p<0.001$ ) and MPV ( $p<0.014$ ) and significantly lower HB/RDW% ratio ( $p<0.001$ ) in patients who are active as opposed to inactive one (Table 3). However, active AS patients had significantly higher RDW% ( $p<0.001$ ) and lower Hb/RDW% ratio ( $p<0.003$ ) compared to inactive patients (Table 4).

**Table (3):** Comparison between active & inactive PsA patients as regard WBCs, RBCs and platelet indices

PSA	Disease Activity						T-Test	
	Inactive			Active			t	P-value
	Mean	±	SD	Mean	±	SD		
NEUT( $10^9/L$ )	4.120	±	1.630	4.163	±	1.575	-0.097	0.923
HCT (%)	37.060	±	1.338	37.103	±	1.686	-0.098	0.922
RBCs( $10^{12}/L$ )	4.772	±	0.506	4.804	±	0.550	-0.220	0.827
Hb(g/L)	12.550	±	0.805	12.625	±	1.233	-0.246	0.806
MCV (fL)	80.575	±	4.835	80.065	±	8.124	0.258	0.797
MCH (pg)	24.805	±	3.067	24.998	±	4.392	-0.175	0.861
RDW%	11.975	±	0.587	14.495	±	1.951	-5.629	<0.001*
MPV/fl	8.220	±	0.562	8.903	±	1.132	-2.536	0.014*
PDW%	11.480	±	2.081	12.178	±	2.698	-1.014	0.315
NEUT/Lympho ratio	2.132	±	0.563	2.070	±	0.457	0.455	0.651
	Inactive			Active			Mann-Whitney test	
	Median	IQR		Median	IQR		Z	P-value
WBCS	6.000	4.150	- 8.250	6.300	5.250	- 7.400	0.660	0.509
Lympho	2.100	1.450	- 2.450	1.900	1.500	- 2.100	0.024	0.981
PLT	216.000	208.000	- 240.000	235.000	215.000	- 276.500	1.359	0.174
Hb/PLT ratio	0.060	0.050	- 0.060	0.050	0.040	- 0.060	1.074	0.283
RDW%/PLT ratio	0.055	0.050	- 0.060	0.060	0.050	- 0.070	1.407	0.159
RBCs/PLT ratio	0.020	0.020	- 0.030	0.020	0.020	- 0.020	0.659	0.510
Hb/RDW% ratio	1.045	0.975	- 1.100	0.845	0.750	- 0.995	3.831	<0.001*
PLT/Lympho ratio	139.855	90.950	- 154.315	132.110	91.355	- 160.000	0.031	0.975

**Table (4):** Comparison between active and inactive AS patients as regard WBCs, RBCs and platelet indices

AS	Disease Activity						T-Test	
	Inactive			Active			t	P-value
	Mean	±	SD	Mean	±	SD		
HCT (%)	38.896	±	3.064	37.670	±	3.255	1.450	0.153
RBCs( $10^{12}/L$ )	5.048	±	0.733	4.833	±	0.578	1.262	0.212
Hb(g/L)	13.091	±	1.759	12.908	±	1.581	0.418	0.678
MCV (fL)	78.235	±	9.023	78.289	±	5.961	-0.028	0.978
MCH (pg)	25.291	±	3.731	25.462	±	4.074	-0.163	0.871
RDW%	12.270	±	1.365	13.830	±	1.449	-4.144	<0.001*
MPV/fl	9.683	±	2.605	9.703	±	2.166	-0.032	0.974
	Inactive			Active			Mann-Whitney test	
	Median	IQR		Median	IQR		Z	P-value
WBCS	6.000	4.800	- 8.400	7.300	6.400	- 8.300	1.217	0.223
NEUT	3.200	2.400	- 4.800	3.600	3.500	- 4.700	1.204	0.229
Lympho	2.000	1.500	- 2.950	2.700	2.000	- 3.100	1.767	0.077
PLT	288.000	201.000	- 387.500	255.000	240.000	- 365.000	0.152	0.879
PWD	10.100	9.750	- 11.650	11.000	9.900	- 11.700	0.822	0.411
Hb/PLT ratio	0.050	0.030	- 0.060	0.050	0.030	- 0.060	0.139	0.889
RDW%/PLT ratio	0.040	0.030	- 0.060	0.050	0.040	- 0.060	1.307	0.191
RBCs/PLT ratio	0.020	0.010	- 0.020	0.020	0.010	- 0.020	0.312	0.755
Hb/RDW% ratio	1.100	0.960	- 1.255	0.980	0.800	- 1.070	2.792	0.005*
NEUT/Lympho ratio	1.470	1.380	- 2.105	1.440	1.240	- 1.970	0.548	0.584
PLT/Lympho ratio	166.000	84.780	- 210.670	124.140	70.000	- 185.240	1.886	0.059

No: number, %: percentage, mm/h: millimeter per hour, mg/L: milligram per Liter, IU/ml: International unit per milliliter, U/L: Unit per liter, mg/dL: milligram per Deci Liter WBCs: White blood cells, HCT: Hematocrit, RBCs: Red blood cells, Hb: Hemoglobin, MCV: Mean corpuscular volume, MCH: Mean corpuscular Hemoglobin, RDW: Red cell distribution width, PLT: Platelets, MPV:

Mean platelet volume, PDW: Platelet distribution width, Hb/PLT ratio: Hemoglobin/Platelet ratio, RDW/PLT ratio: Red cell distribution width/ Platelet ratio, RBCs/PLT ratio: Red blood cells/Platelet ratio, Hb/RDW ratio: Hemoglobin/ Red cell distribution width ratio, NLR: Neutrophil/Lymphocyte ratio, PLR: Platelet/Lymphocyte ratio, /mm<sup>3</sup>: Per cubic millimeter, %: percentage, g/dL: Gram per deci Liter, fl: femtoliter, pg: picogram, t: independent sample t test, P: probability value, \*: p<0.05 is statistically significant, \*\*: p<0.001 is statistically highly significant

Correlation of different blood cell indices with disease activity clinical and laboratory parameters showed that among psoriatic patients, RDW% was positively correlated with the number of tender and swollen joint, disease activity score, ESR and the C-RP titer. Hb/RDW% ratio was negatively correlated with all same parameters. While, the MCV was negatively correlated with the number of tender joints, swollen joint and ESR (Table 5). Regarding patients, results yielded that RDW% was positively correlated to disease activity score and tender joints. While, the Hb/RDW% ratio was inversely correlated to same parameters (Table 6).

**Table (5):** Correlation between the WBCs, RBCs, Platelets indices and parameters of disease activity in PsA

PSA	Tender j		Swollen j		Activity score		ESR		CRP	
	r	P-value	r	P-value	r	P-value	r	P-value	r	P-value
WBCS (10 <sup>9</sup> /L)	-0.161	0.219	-0.153	0.266	-0.042	0.750	-0.125	0.342	0.110	0.402
NEUT (10 <sup>9</sup> /L)	-0.186	0.154	-0.197	0.150	-0.102	0.436	-0.191	0.144	0.052	0.694
Lympho(10 <sup>9</sup> /L)	-0.247	0.057	-0.239	0.078	-0.119	0.366	-0.196	0.134	0.083	0.526
HCT (%)	-0.025	0.847	-0.022	0.873	0.021	0.873	0.057	0.667	-0.093	0.480
RBCs(10 <sup>12</sup> /L)	0.054	0.680	0.067	0.627	0.108	0.413	0.154	0.241	-0.024	0.858
Hb(g/L)	0.084	0.523	0.050	0.716	0.058	0.662	-0.041	0.755	-0.043	0.742
MCV (fL)	-0.250	0.054*	-0.347	0.009*	-0.213	0.102	-0.266	0.040*	-0.186	0.155
MCH (pg)	-0.205	0.117	-0.205	0.133	-0.130	0.321	-0.165	0.209	-0.001	0.995
RDW%	0.529	<0.001*	0.380	0.004*	0.614	<0.001*	0.433	0.001*	0.388	0.002*
PLT (10 <sup>9</sup> /L)	0.039	0.770	0.004	0.978	0.050	0.707	-0.040	0.764	0.039	0.768
MPV/fl	0.263	0.042	0.172	0.210	0.196	0.134	0.187	0.153	0.172	0.188
PDW%	0.099	0.452	-0.054	0.694	0.037	0.779	-0.070	0.596	-0.040	0.760
Hb/PLT ratio	0.044	0.738	0.100	0.466	-0.007	0.959	0.095	0.472	-0.053	0.685
RDW%/PLT ratio	0.233	0.074	0.204	0.135	0.241	0.063	0.219	0.092	0.107	0.415
RBCs/PLT ratio	-0.024	0.855	0.061	0.660	-0.057	0.668	0.093	0.480	-0.050	0.705
Hb/RDW% ratio	-0.426	0.001*	-0.316	0.019*	-0.525	<0.001*	-0.410	0.001*	-0.366	0.004*
NLR ratio	-0.002	0.987	-0.061	0.661	-0.063	0.633	-0.107	0.415	-0.067	0.612
PLR ratio	0.148	0.258	0.127	0.355	0.053	0.685	0.043	0.743	-0.097	0.460

**Table (6):** Correlation between the WBCs, RBCs, Platelets indices and parameters of disease activity in AS

AS	Tender j		Swollen j		Activity score		ESR		CRP	
	r	P-value	r	P-value	r	P-value	r	P-value	r	P-value
WBCS(10 <sup>9</sup> /L)	0.246	0.071	0.005	0.978	0.204	0.118	0.100	0.447	0.368	0.004*
NEUT(10 <sup>9</sup> /L)	0.114	0.408	-0.019	0.906	0.148	0.260	0.364	0.004*	0.355	0.005*
Lympho(10 <sup>9</sup> /L)	0.285	0.035*	-0.004	0.978	0.150	0.253	-0.085	0.521	0.193	0.139
HCT (%)	-0.180	0.188	-0.164	0.313	-0.099	0.451	-0.017	0.899	0.084	0.522
RBCs(10 <sup>12</sup> /L)	-0.110	0.425	-0.331	0.037*	-0.060	0.647	0.092	0.483	0.235	0.070
Hb(g/L)	-0.237	0.082	-0.150	0.354	-0.056	0.671	-0.224	0.085	-0.073	0.578
MCV (fL)	-0.178	0.193	0.492	0.001*	0.069	0.603	0.015	0.907	-0.157	0.232
MCH (pg)	-0.188	0.170	0.205	0.203	-0.019	0.884	-0.182	0.164	-0.357	0.005*
RDW%	0.551	<0.001*	0.101	0.533	0.460	<0.001*	0.090	0.493	0.140	0.285
PLT(10 <sup>9</sup> /L)	-0.060	0.666	0.182	0.261	-0.056	0.673	0.266	0.040*	0.223	0.087
MPV/fl	-0.020	0.886	-0.188	0.244	-0.128	0.330	-0.313	0.015*	-0.244	0.061
PDW%	-0.004	0.979	-0.278	0.082	-0.036	0.782	-0.278	0.031*	0.055	0.675
Hb/PLT ratio	-0.027	0.844	-0.314	0.048*	-0.066	0.614	-0.319	0.013*	-0.161	0.220
RDW%/PLT ratio	0.219	0.108	-0.243	0.131	0.103	0.434	-0.272	0.036*	-0.210	0.107
RBCs/PLT ratio	0.020	0.885	-0.553	<0.001*	-0.101	0.443	-0.269	0.038*	-0.037	0.777
Hb/RDW% ratio	-0.543	<0.001*	-0.166	0.306	-0.364	0.004*	-0.189	0.148	-0.137	0.297
NLR ratio	-0.172	0.208	0.081	0.619	0.033	0.803	0.300	0.020*	0.087	0.509
PLR ratio	-0.305	0.023*	0.175	0.279	-0.127	0.332	0.208	0.111	-0.031	0.812

SD: Standard Deviation, No: number, %: percentage, mm/h: millimeter per hour, mg/L: milligram per Liter, IU/ml: International unit per milliliter, U/L: Unit per liter, mg/dL: milligram per Deci Liter WBCs: White blood cells, HCT: Hematocrit, RBCs: Red blood cells, Hb: Hemoglobin, MCV: Mean corpuscular volume, MCH: Mean corpuscular Hemoglobin, RDW: Red cell distribution width, PLT:

Platelets, MPV: Mean platelet volume, PDW: Platelet distribution width, Hb/PLT ratio: Hemoglobin/Platelet ratio, RDW/PLT ratio: Red cell distribution width/ Platelet ratio, RBCs/PLT ratio: Red blood cells/Platelet ratio, Hb/RDW ratio: Hemoglobin/ Red cell distribution width ratio, NLR: Neutrophil/Lymphocyte ratio, PLR: Platelet/Lymphocyte ratio, /mm<sup>3</sup>: Per cubic millimeter, %: percentage, g/dL: Gram per deci Liter, fl: femtoliter, pg: picogram, t: independent sample t test, P: probability value, \*: p<0.05 is statistically significant, \*\*: p≤0.001 is statistically highly significant.

The following ratios, regarding PSA patients, were statistically significant with + ve HLA B27 test such as Hb/PLT ratio, Hb/RDW% ratio and PLT/Lympho ratio with p< 0.025, <0.002 and <0.012 respectively as shown in table (7). While the AS patient all ratios were statistically insignificant as shown in table (8).

**Table (7): Relation of HLA B27 test in PSA patients**

PSA	HLA B27						T-Test	
	Positive			Negative			t	P-value
	Mean	±	SD	Mean	±	SD		
<b>WBCS</b>	6.207	±	1.513	7.447	±	1.094	-2.088	0.041*
<b>HCT</b>	37.072	±	1.659	37.129	±	1.351	-0.127	0.900
<b>RBCs</b>	4.713	±	0.533	4.994	±	0.487	-1.882	0.065
<b>Hb</b>	12.398	±	0.939	13.112	±	1.333	-2.346	0.022*
<b>MCV</b>	77.970	±	5.643	85.965	±	7.522	-4.488	<0.001*
<b>MCH</b>	24.019	±	4.069	27.247	±	2.610	-3.026	0.004*
<b>RDW%</b>	14.002	±	2.111	12.776	±	1.463	2.190	0.033*
<b>MPV/fl</b>	8.607	±	0.921	8.847	±	1.271	-0.814	0.419
<b>PWD</b>	11.858	±	2.153	12.165	±	2.324	-0.423	0.674
<b>NEUT/Lympho ratio</b>	2.073	±	0.12	2.135	±	0.445	-0.436	0.665
	Positive			Negative			Mann-Whitney Test	
	Median	IQR		Median	IQR		Z	P-value
<b>WBCS</b>	6.100	5.200	- 7.050	7.300	4.300	- 9.200	1.232	0.218
<b>NEUT</b>	4.000	2.850	- 4.800	5.100	3.200	- 6.100	1.380	0.168
<b>Lympho</b>	1.800	1.500	- 2.100	2.100	1.600	- 2.700	1.855	0.064
<b>PWD</b>	11.900	10.100	- 13.300	10.800	10.000	- 13.100	0.296	0.767
<b>Hb/PLT ratio</b>	0.050	0.040	- 0.060	0.060	0.050	- 0.070	2.626	0.009*
<b>RDW%/PLT ratio</b>	0.060	0.050	- 0.060	0.060	0.050	- 0.070	1.286	0.198
<b>RBCs/PLT ratio</b>	0.020	0.020	- 0.020	0.020	0.020	- 0.030	1.700	0.089
<b>Hb/RDW% ratio</b>	0.920	0.760	- 1.020	1.050	0.960	- 1.100	3.022	0.003*
<b>PLT/Lympho ratio</b>	145.330	118.155	- 155.770	80.000	77.620	- 134.380	2.840	0.005*

Parametric data represented by mean SD -Non parametric data represented by median (IQR)

**Table (8):** Relation of HLA B27 test in AS patients.

AS	HLA B27						T-Test	
	Positive			Negative			t	P-value
	Mean	±	SD	Mean	±	SD		
<b>HCT</b>	38.146	±	3.146	38.130	±	3.391	0.018	0.986
<b>RBCs</b>	4.959	±	0.749	4.846	±	0.434	0.657	0.514
<b>Hb</b>	12.973	±	1.582	12.987	±	1.763	-0.032	0.975
<b>MCV</b>	78.327	±	7.160	78.174	±	7.460	0.079	0.937
<b>MCH</b>	25.376	±	4.430	25.430	±	2.995	-0.052	0.959
<b>RDW%</b>	13.241	±	1.713	13.217	±	1.437	0.054	0.957
<b>PLT</b>	322.568	±	16.804	288.870	±	41.543	1.047	0.299
<b>Hb/RDW% ratio</b>	0.997	±	0.178	0.996	±	0.186	0.029	0.977
	Positive			Negative			Mann-Whitney Test	
	Median	IQR		Median	IQR		Z	P-value
<b>WBCS</b>	6.900	5.000 -	8.300	7.500	5.450 -	8.600	0.723	0.470
<b>NEUT</b>	3.600	2.800 -	4.700	4.400	3.200 -	4.850	0.541	0.588
<b>Lympho</b>	2.500	1.800 -	2.900	2.400	1.600 -	3.300	0.099	0.921
<b>PLT</b>	312.000	245.000 -	373.000	253.000	201.000 -	300.000	1.666	0.096
<b>MPV/fl</b>	9.200	8.600 -	10.200	9.300	8.700 -	10.900	0.891	0.373
<b>PWD</b>	10.100	9.800 -	11.400	11.000	9.900 -	12.600	1.485	0.138
<b>Hb/PLT ratio</b>	0.050	0.030 -	0.060	0.050	0.050 -	0.060	1.821	0.069
<b>RDW%/PLT ratio</b>	0.040	0.030 -	0.060	0.050	0.040 -	0.065	1.384	0.166
<b>RBCs/PLT ratio</b>	0.020	0.010 -	0.020	0.020	0.020 -	0.020	1.602	0.109
<b>NEUT/Lympho ratio</b>	1.440	1.290 -	1.750	1.440	1.250 -	2.105	0.221	0.825
<b>PLT/Lympho ratio</b>	133.330	92.590 -	201.720	141.480	76.065 -	190.955	0.844	0.399

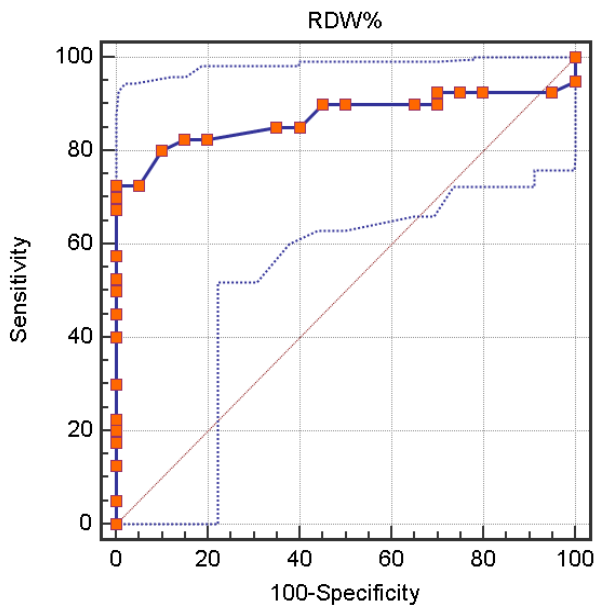
WBCs: White blood cells, HCT: Hematocrit, RBCs: Red blood cells, Hb: Hemoglobin, MCV: Mean corpuscular volume, MCH: Mean corpuscular Hemoglobin, RDW: Red cell distribution width, PLT: Platelets, MPV: Mean platelet volume, PDW: Platelet distribution width, Hb/PLT ratio: Hemoglobin/Platelet ratio, RDW/PLT ratio: Red cell distribution width/Platelet ratio, RBCs/PLT ratio: Red blood cells/Platelet ratio, Hb/RDW ratio: Hemoglobin/ Red cell distribution width ratio, NLR: Neutrophil/Lymphocyte ratio, PLR: Platelet/Lymphocyte ratio, /mm<sup>3</sup>: Per cubic millimeter, %: percentage, g/dL: Gram per deci Liter, fl: femtoliter, pg: picogram, t: independent sample t test, P: probability value, \*: p<0.05 is statistically significant, \*\*: p≤0.001 is statistically highly significant.



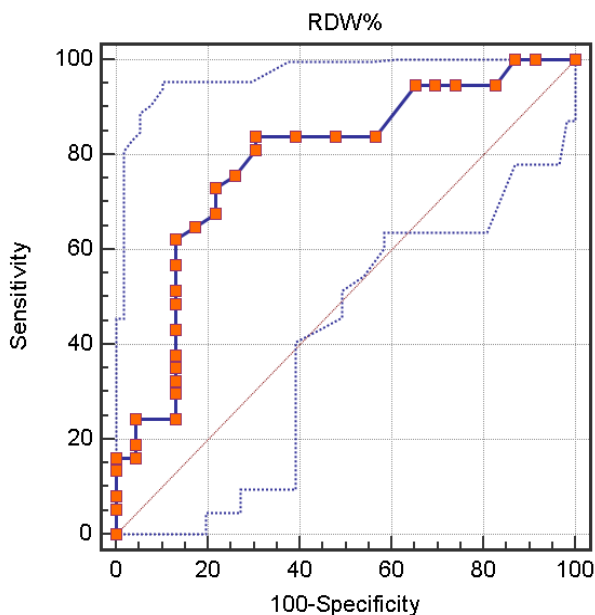
### Diagnostic role of RDW to predict disease activity

The ROC curve revealed that the best RDW% cut-off value for disease activity prediction in AS was >12.4. The sensitivity was 83.78% and the specificity was 69.57% with positive predictive value of 81.6%.

In PsA patients, the best RDW% cut-off value for disease activity prediction was >13.2 with specificity of 100.0% and sensitivity of 72.50% with 100.0% positive predictive value. While, the best MPV cut-off value for predicting disease activity was > 8.4 with specificity of 70.0% and sensitivity of 67.50% with positive predictive value is 81.8% (Figures 1 & 2).



**Fig (1)** ROC curve showed the validity of RDW for disease activity prediction in PsA patients



**Fig (2)** ROC curve showed the validity of RDW for disease activity prediction in AS patients

### DISCUSSION

The etiopathogenesis of spondyloarthropathy is the result of interplay between genetic, immunoregulatory, ethnic, factors, and several key points of these multifactorial connections are still unclear <sup>(15)</sup>, with summit of inflammatory status, which influence the disease severity.

MPV and RDW could reflect the extent of heterogeneity of circulating erythrocytes volume and platelets, which are influenced in inflammatory conditions <sup>(16)</sup>. Despite a great amount of evidence about the relations between MPV, RDW, platelet count and other hematological parameters in rheumatic diseases has been elucidated, but controversial results have been found <sup>(17)</sup>. However, disease activity relation in either PsA or AS and various blood cell indices has not been fully demonstrated yet <sup>(18)</sup>.

In current study, trying to assess the relation between the various blood cell indices and the clinical, laboratory parameters of disease activity. In psoriatic patients we demonstrated significantly lower HB/RDW% ratio ( $p < 0.001$ ) and significantly higher RDW% ( $p < 0.001$ ) among patients who are active vs those who are not. Also, the number of tender joints, swollen joints, disease activity score, ESR and the CRP titer were positively correlated with RDW%, while they were negatively correlated with Hb/RDW% ratio. The MCV was negatively correlated with the number of tender joints, swollen joints and ESR. That may reflect the significance of these indices as indicators of disease activity and inflammation. This is similar to the results of **Gisoni** <sup>(19)</sup> who revealed that RDW is elevated in psoriatic patients, but unsimilarly, RDW did not seem to be associated with disease severity. Moreover, **Ozisler and Sandikci** <sup>(20)</sup> concluded that increased RDW found in active disease and it may be indicator of the inflammation similar to the CRP and ESR. And that RDW could be a biomarker of disease activity in PsA.

Although MPV ( $p < 0.014$ ) was significantly higher in active patients but MPV was not correlated to activity score. This is matching with a previous study where the correlation was not significant between PASI values and MPV level <sup>(21, 22, 23)</sup>. This contradiction might be attributed to different ethnic population and sample size.

Regarding AS, active patients had significantly higher RDW% ( $p < 0.001$ ) and lower Hb/RDW% ratio ( $p < 0.003$ ) compared to inactive patients. Also, RDW% was positively correlated to the number of tender joints and disease activity score. While, the Hb/RDW% ratio was inversely correlated to same parameters. This agrees with **Melek and Coworkers** <sup>(24)</sup> who found RDW showed higher significance in AS patients than in the controls. They had detected that RDW differed significantly in active compared to non-active patients. In line with

current study, activity score of AS was positively related to RDW and not related to MPV, other studies found MPV was not correlated to BSDAI<sup>(24, 25)</sup>. In contrast to the above-mentioned results, according to certain research, AS with active AS had lower MPV compared to inactive patients<sup>(26, 27)</sup>. this may be attributed to super production of acute reactants and pro-inflammatory cytokines that reduce the size of platelets<sup>(28)</sup>.

RDW and MPV has been shown to be strongly associated with CRP and ESR in a various inflammatory disease as RA, IBD. Moreover, CRP has been observed to be linearly related to elevated RDW and MPV in prior studies<sup>(29-31)</sup> and these results partially disagree with ours as we didn't find correlation between MPV, CRP and RDW values although there was negative correlation of MPV to the ESR.

On evaluation of the performance validity of hemoparameters, the current study concluded that RDW% higher specificity with good sensitivity for prediction of psoriatic activity, while among AS, RDW showed good sensitivity with fair specificity inconsistent with another study that showed RDW and MPV were more specific than sensitive for disease activity<sup>(24)</sup>.

Chronic inflammation can adversely influence, in view of these results, thrombopoiesis erythropoiesis through variable routs such as cell apoptosis and increased oxidative stress. Thus, inflammation may lead to anisocytosis and change in platelet maturation and size in spondyloarthropathy patients. Once CBC is cheap and easy to be done, blood cell indices could be used as tool for detection and following the activity of the disease.

Regarding the HLA-B 27, PSA patients with + ve HLA B27 test had lower Hb/PLT ratio, with higher Hb/RDW% ratio and PLR, while there was no significant difference between the AS patients in both HLA B27 groups.

The current study has some shortcomings: It is cross-sectional in design, small sized sample and no accessible information about the nutritional status or nutritional elements as levels of vitamin B12, folic acid or iron, which may result in change in CBC parameters.

## CONCLUSION

The current study has suggested that blood cell indices were related to both clinical and laboratory parameter of the disease activity in both AS and PsA. Also, the RDW% was the most related index to the disease activity score and inflammatory markers and it may be used to distinguish between active from inactive disease with best cut-off value >13.2 and >12.4 in PsA and AS respectively.

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