VITAMIN D STATUS IN EGYPTIAN PATIENTS WITH PSORIATIC ARTHRITIS AND ITS RELATIONSHIP WITH THE DISEASE ACTIVITY AND SEVERITY.

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ABSTRACT:

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Background: Many studies proved high percentage of vitamin D deficiency in psoriatic arthritis patients. Different records about the possible link between vitamin D deficiency and state of inflammatory & disease activity in psoriatic arthritis patients)

Aim of the work: To evaluate the vitamin D level in Egyptian patients with psoriatic arthritis and its association with activity of the disease as well as disease severity

Patients & Method: 45 Psoriatic Arthritis Patients fulfilled the Classification Criteria for the diagnosis of Psoriatic Arthritis (CASPAR) and 50 healthy controls were included. Medical history was taken, general and musculoskeletal examination were done. Disease activity was assessed using the Disease activity in psoriatic arthritis "DAPSA" score. VAS, HAQ, ESR & CRP were done. Serum level of 25(OH) vitamin D was measured by ELISA for patients and controls.

Results: Serum vitamin D level in psoriatic arthritis patients ranged from 10-48 ng/mL, with mean \pm SD 32.844 \pm 12.395. While in healthy controls it ranged from15-50 ng/mL with mean \pm SD33.940 \pm 8.998, without statistical significant difference. Vitamin D deficiency was detected in 12 patients (26.6%) and 5 (10%) healthy participants with statistically significant higher frequency of vitamin D deficiency in patients than controls (P-value <0.034). patients with vitamin D deficiency showed statistically significant longer disease duration, higher VAS, HAQ score, ESR, CRP titer and DAPSA score (P-value <0.001). Also, they had statistically significant higher frequency of axial affection (P-value <0.001). Vitamin D level was statistically negatively correlated with disease duration, VAS, HAQ, ESR, CRP and DAPSA score (P-value <0.001).

Conclusions: We concluded that psoriatic arthritis patients had higher frequency of vitamin D deficiency than healthy individuals. And that patients with vitamin D deficiency had more active and more severe disease compared to those without deficiency.

Keywords: Vitamin D, psoriatic arthritis, disease activity

INTRODUCTION:

Psoriatic arthritis (PsA) is a chronic autoimmune disease manifested by arthritis, enthesitis, dactylitis, skin lesions and axial affection 1,2,3 .

Lower vitamin D level was related to autoimmune diseases⁴. Vitamin D appears to affect the immune system response, it prevents the production of some cytokines (interleukin 1, interleukin 6, interleukin 12, Tumor Necrosis Factor $-\alpha$), and suppress the T helper 1 cells production of interleukin $2^{5\&6}$. In fact, vitamin D had immunomodulating, anti-inflammatory, antifibrotic and anti-oxidant effects ⁷.

Many previous studies proved high incidence of vitamin D deficiency in psoriatic arthritis patients^{8&9}. Different records about the probable relationship between vitamin D deficiency and inflammatory status & disease activity in psoriatic arthritis¹⁰.

Several studies proved higher frequency of vitamin D deficiency in psoriatic arthritis patients and its relation with the disease activity and severity, while other many studies didn't found that. So, we found that was an interesting research subject to investigate.

AIM OF THE WORK:

To evaluate the vitamin D level in Egyptian patients with psoriatic arthritis and its association with activity of the disease as well as disease severity

PATIENTS & METHODS:

Ninety five individuals participated in this cross sectional study. Forty five of them were diagnosed as psoriatic arthritis and fulfilled the Classification criteria for the diagnosis of Psoriatic Arthritis (CASPAR)¹¹. Patients were treated in internal medicine and Rheumatology clinics, Ain Shams University hospitals. Patients on vitamin D supplements were excluded. Another fifty healthy age and sex matched individuals participated as control group.

All participants gave an informed consent after explaining the purposes and the methodology of the study.

Ethical consideration:

Ethical approval was obtained from Ethical Committee of Scientific Research, Faculty of Medicine, Ain Shams University (FWA000017585 FMASU R 114/2023).

Detailed medical history was taken from all patients. General and musculoskeletal examination were done. Assessment of the disease activity was done using the Disease activity in psoriatic arthritis DAPSA score¹². Presence of axial affection (proved by MRI) was recorded. Assessment of pain intensity using the 0–10 visual analogue scale (VAS) as o = no pain; 10 = intense pain, and functional ability using health assessment questionnaire (HAQ) score was done as score less than 0.5 considered normal while score more than 1.5 considered severe disability 13 . Inflammatory markers were done included erythrocyte sedimentation rate (ESR), Creactive protein (CRP) titer.

Serum level of 25(OH) vitamin D was measured by ELISA for patients and controls. Results were interpreted as follows: (0-20 ng/mL);insufficiency deficiency (21-30 ng/mL); sufficiency (31-80 ng/mL) ¹⁴. Comparison between patients and controls regarding vitamin D level was done. Patients were divided according to the serum level of vitamin D into, group I: patients with vitamin D deficiency and group II: patients without vitamin D deficiency. Comparison between both groups was done regarding demographic data, clinical and laboratory markers. Then vitamin D level was correlated with parameters of disease activity and severity.

Data were coded and entered utilizing the statistical package SPSS version 24. Comparisons between quantitative variables will be done using the nonparametric Kruskal-Wallis and Mann-Whitney tests. For comparing categorical data, chi-square (χ 2) test will be performed.

RESULTS:

Table (1): Descriptive data of studied psoriatic arthritis patients.

Patier	Ν	%		
Sex	Male	14	31.11	
	Female	31	68.89	
Age (years)	Range	18	-	45
	Mean ±SD	31.933	+	7.650
Smoking	Yes	6	13.33	
	No	39		86.67
Disease duration (years)	Range	2	-	14
	Mean ±SD	5.578	+1	3.368
Axial affection	Yes 8			17.78
	No	37		82.22
VAS	Range	0	-	10
	Mean ±SD	4.289	+1	2.744
HAQ	Range	0.2	-	3
	Mean ±SD	0.667	±	0.517
ESR (mm/h)	Range	12	-	70
	Mean ±SD	34.933	±	15.049
CRP (mg/L)	Range	4	-	48
	Mean ±SD	17.000	+1	13.509
DAPSA score	Range	2	-	34
	Mean ±SD	11.178	±	9.803

There were 31(68.89%) females and 14 (31.11%) males, their ages ranged from 18 to 45 years with Mean ±SD 31.933 ± 7.650 . (Tab 1)

Table (2): Comparison between healthy controls and psoriatic arthritis patients regarding vitamin D level.

		Group					T-Test		
		Patient		Control			t	P-value	
VIT D	Range	10	-	48	15	-	50	-0.496	0.621
(ng/mL)	Mean ±SD	32.844	±	12.395	33.940	±	8.998		
Chi-	Square	N		%	N		%	X^2	P-value
VIT D	Deficient	12		26.67	5		10.00	4.478	0.034
	Non deficient	33		73.33	45		90.00		

Serum vitamin D level in psoriatic arthritis patients ranged from 10-48 ng/mL, with mean \pm SD 32.844 \pm 12.395. While in healthy controls it ranged from15-50 ng/mL with mean \pm SD33.940 \pm 8.998, without statistical significant difference. (Tab 2)

Vitamin D deficiency was more frequent in patients than controls [12 patients (26.6%) Vs 5(10%)]. This higher frequency had statistical significance (P-value <0.034).(Tab 2)

Patient		VIT D						T-Test	
		With deficiency		Without deficiency			t	P-value	
Age(years)	Range	23	-	44	18	-	45	0.915	0.365
	Mean ±SD	33.667	ŧ	7.572	31.303	±	7.695		
Disease duration	Range	3	-	14	2	-	12	3.776	< 0.001
(years)	Mean ±SD	8.333	±	3.257	4.576	±	2.840		
VAS	Range	6	-	10	0	-	6	8.939	< 0.001
	Mean ±SD	7.917	±	1.311	2.970	±	1.741		
HAQ	Range	0.9	-	1.6	0.2	-	3	4.669	< 0.001
	Mean ±SD	1.158	±	0.202	0.488	±	0.479		
ESR (mm/h)	Range	39	-	70	12	-	51	6.978	< 0.001
	Mean ±SD	52.917	ŧ	10.808	28.394	±	10.289		
CRP (mg/L)	Range	28	-	48	4	-	20	21.242	< 0.001
	Mean ±SD	38.167	ŧ	5.875	9.303	±	3.157		
DAPSA score	Range	16	-	34	2	-	14	12.514	< 0.001
	Mean ±SD	25.417	ŧ	5.915	6.000	±	4.054		
Chi-Square		N		%	N		%	X ²	P-value
Sex	Male	2		16.67	12		36.36	1.593	0.207
	Female	10		83.33	21		63.64		
Smoking	Yes	1		8.33	5		15.15	0.354	0.552
	No	11		91.67	28		84.85		
Axial affection	Yes	8		66.67	0		0.00	26.757	< 0.001
	No	4		33.33	33		100.00		

Table (3): Comparison between patients with vitamin D deficiency and patients without vitamin D deficiency.

Comparison between patients with vitamin D deficiency and those without vitamin D deficiency showed statistically significant longer disease duration, higher VAS, HAQ score, ESR, CRP titer and DAPSA score (P-value <0.001) in patients with vitamin D deficiency. Also they had statistically significant higher frequency of axial affection (P-value <0.001). (Tab 3)

Table (4): Correlation between vitamin D level and parameters of disease severity and activity in psoriatic arthritis patients.

	VIT D				
	r	P-value			
Disease duration	-0.571	< 0.001			
VAS	-0.920	< 0.001			
HAQ	-0.524	< 0.001			
ESR (mm/h)	-0.829	< 0.001			
CRP (mg/L)	-0.934	< 0.001			
DAPSA score	-0.935	< 0.001			

Vitamin D level was statistically negatively correlated with disease duration, VAS, HAQ, ESR, CRP and DAPSA score (P-value <0.001). (Tab 4).

DISCUSSION:

Frequency of vitamin D deficiency in PSA and its relationship with the inflammatory status and activity is an interesting topic to investigate, so current study was designed to assess level of vitamin D in Egyptian patients with PSA and its association with activity of the disease as well as disease severity

In current study there was female predominance in our psoriatic arthritis patients as there were 31 females (68.89%) and 14 males (31.11%). These results are consistent with **El Helaly et al.**¹⁰ who found that there was female sex predilection in

psoriatic arthritis. While disagree with **Vekić Mužević et al.**¹⁵ who didn't observe that.

In our psoriatic arthritis patients, serum vitamin D level ranged from 10-48 ng/mL, with mean \pm SD 32.844 \pm 12.395. Twelve patients (26.6%) were deficient. These findings are partially agree with those of **Urruticoechea-Arana et al.**¹⁶ who observed that vitamin D level in psoriatic arthritis patients ranged from 14.0 – 28.8 ng/ml with median 20.0 ng/ml, but they found higher frequency of vitamin D deficiency than our findings as they detected 40.9% of their patients were deficient.

Our research didn't reveal statistically significant difference between patients and controls regarding vitamin D level that disagree with **Radić et al.**¹⁷ **Mohammed et al.**¹⁸, **Ibrahim et al.**¹⁹, **Petho et al.**²⁰, and **Touma et al.**²¹ who detected lower Vitamin D levels in psoriatic arthritis patients rather than healthy participants. This disagreement could be explained by our small sample size.

Despite the fact that there was no statistically significant difference in vitamin D levels between patients and controls, there was no statistically significant difference between patients and controls regarding vitamin D level, but, there was statistically significant higher frequency of vitamin D deficiency in patients than controls. The same was recorded by **El Tawab et al.**²² who found that vitamin D level didn't show significant difference in psoriatic arthritis patients and healthy participants had lower frequency of vitamin D deficiency than patients with psoriatic arthritis.

Similarly, **Gamonal et al.**²³ found that vitamin D deficiency was more frequently seen in patients with PsA (82.2%).

Vitamin D deficiency in psoriatic arthritis patients could be clarified by many causes such as less exposure to sun, taking drugs that interfere with Vitamin D metabolism like, steroids or immunosuppressive drugs, and decreased Vitamin D intake **Filoni et al.**²⁴. Also, chronic diseases lead to decreased synthesis or increased catabolism of vitamin D **Urruticoechea-Arana et al.**¹⁶

While investigating the relationship of vitamin D with the disease severity, our results revealed that vitamin D deficiency was linked to more severe disease indicated by statistically significant longer disease duration, higher VAS and HAQ score in deficient patients and by presence of statistical negative correlation of vitamin D level with these parameters. Also, presence of higher frequency of axial affection in patients with vitamin D deficiency.

In line, Results of **Mohammed et al.**¹⁸ showed that Vitamin D deficiency was linked to longer disease duration. And **Sag et al.**²⁵ found that the duration of psoriatic arthritis was inversely correlated with serum Vitamin D level.

In harmony, **Rotondo et al.**²⁶ noted a high frequency of axial affection in the psoriatic patients with vitamin D deficiency which go hand by hand with **Fernandes et al.**²⁷ who considered presence of axial affection as indicator of disease severity.

Kincse et al.²⁸ noticed that vitamin D level was significantly linked to the functional ability in psoriatic arthritis that supports our findings. While **Montolio-Chiva et al.**²⁹ didn't observe any relationship between vitamin D and HAQ score.

The link between VAS and vitamin D come in agreement with the theory of the relationship between vitamin D deficiency and musculoskeletal pain **Plotnikoff et al.**³⁰. Also it may be due to the higher disease activity in the deficient patients.

Vitamin D level may be correlated with disease activity and inflammatory markers **Deng et al.**³¹, **Zhao et al.**³² and **Cai et al.**³³. In current study, vitamin D deficiency was linked to more active disease as we found that inflammatory markers and DAPSA score

were significantly higher in patients with vitamin D deficiency (P-value <0.001), also same parameters were negatively correlated with vitamin D level (P-value <0.001).

Our results agree with **Kincse et al.**²⁸ who confirmed negative correlation between serum vitamin D and DAPSA score in psoriatic arthritis. And **El Helaly et al.**¹⁰ who found that patients with high CRP showed lower levels of vitamin D.

Montolio-Chiva et al.²⁹ observed inverse correlation between vitamin D level and number of tender joint and swollen joint and that patients with vitamin D deficiency had higher DAPSA score. On the other hand **Braun-Moscovici et al.**³⁴ didn't found Vitamin D to be correlated with disease activity scoring in psoriatic arthritis.

Furthermore, our results agree with **Mohammed et al.**¹⁸ who found that Vitamin D level is lower in psoriatic arthritis with high disease activity score.

Our findings support those of **Pavlov et al.** ³⁵ who found that vitamin D was significantly correlated with CRP and reported that vitamin D deficiency may be used as indicator of more active disease.

Current results partially agree with those of **El Helaly et al.**¹⁰ and **El Tawab et al.**²² who observed that vitamin D deficiency was frequent in psoriatic arthritis patients. However, they did not correlate vitamin D deficiency with the activity or severity in their patients.

On the contrary, **Rotondo et al.**²⁶ showed that vitamin D level was not correlated with inflammatory markers, furthermore, **Ibrahim et al.**¹⁹ and **Sag et al.**²⁵ reported that vitamin D level was not correlated with CRP.

This study had some limitations such as being a single center study and small sample size and absence of follow up of deficient patients after giving vitamin D supplements. This study had some limitations, including the fact that it was conducted in a single center with mall sample size, and the lack of follow-up with patients who were vitamin D deficient after receiving supplements.

Conclusions:

Finally we concluded that psoriatic arthritis patients had higher frequency of vitamin D deficiency than healthy individuals. And that patients with vitamin D deficiency had more severe and more active disease compared to those without deficiency.

Disclosure and Conflict of Interest:

Competing interests: The authors declare that they have no competing interests concerning this article.

Ethical approval: All procedures performed in the study were in accordance with the ethical standards of the faculty of medicine, Ain Shams university research and ethical committee. We obtained approval from Research Ethics Committee (REC) No. FWA 000017585. FMASU R 114/2023, On 1/5/2023. Written informed consent was obtained from participants for participation in this study.

The FMASU REC is organized and operated according to guidelines of the International Council on Harmonization (ICH) and the Islamic Organization of Medical Sciences (IOMS), the United States Office for Human research Protections and the United States Code of Feral Regulations and operates under Federal Wide Assurance No. FWA 000017585. FMASU R 114/2023.

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Authors contributions:

All authors have participated in the concept, design, collect, analysis and interpretation of data, writing, drafting and revising the manuscript. FM: recruited patients, carried out clinical examination and assessment, and generated the result sheets. underwent data tabulation MAZ: and statistical analysis, and interpreted the patient's data and wrote the final results. MM: recruited patients, carried out clinical examination and assessment, and revised data interpretation and manuscript. NN: was the major contributor in writing and editing the manuscript, designed the protocol, carried out the Ethical approval, and data collection. All authors have agreed to conditions noted on the Authorship Agreement Form and have read and approved the final version submitted. The content of the manuscript has not been published, or submitted for publication elsewhere. All authors read and approved the final manuscript.

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حالة فيتامين د لدى مرضى التهاب المفاصل الصدفي المصريين وعلاقته بنشاط المرض وشدته فاطمه محمد عبود^{رو۲}, منه الله زكريا ابو الوفا^۳, منى محمد البابلى³, نرمين نصحى عزيز ⁽ 'باطنه عامه طب عين شمس, ^۲باطنه عامه طب القوات المسلحه, ^۳كلينيكال باثولوجى, طب عين شمس,³طب طبيعى وتاهيل طب عين شمس

الخلفية: وصفت الكثير من الأدلة العلمية ارتفاع معدل نقص فيتامين (د) في مرضى الصدفية والتهاب المفاصل الصدفى . تم الإبلاغ عن بيانات متناقضة حول الارتباط المحتمل بين المستويات المنخفضة من فيتامين (د) وحالة الالتهاب أو نشاط المرض.

الهدف من البحث: الهدف من هذه الدراسة هو تقييم حالة فيتامين (د) في المرضى المصريين المصاريين المصاريين المصاريين المصاريين بالتهاب المفاصل الصدفي والتعرف على تأثير نقص فيتامين (د) على شدة المرض ونشاطه.

الطريقة: الدراسة شملت ٤٥ مريض التهاب المفاصل الصدفي تم تشخيصهم طبقا لتوجيهات المنظمة الأمريكية للأمراض الروماتيزمية وتم تجميعهم من عيادة الروماتيزم وقسم الباطنة العامة. و٥٠ شخص اصحاء. تم فحص المرضى بعد اخذ التاريخ المرضى ثم عمل فحوصات شملت سرعه الترسيب وبروتين سى النشطز تم قياس معدل فيتامين (د) في الدم لكلا من المرضى و الاصحاء.

النتائج: وجدنا ان معدل حالات نقص فيتامين (د) في المرضى اكثر من الاصحاء. بعد تقسيم المرضى الى مجموعتين, وجدنا ان المجموعه التي تعانى من نقص فيتامين (د) لديها مده مرض اطول و احساس بالالم اعلى و نسبه الاعاقه الحركيه اعلى و اصابه العمود الفقرى اعلى و كذلك لديهم سرعه ترسيب وبروتين سي نشط اعلى من المجموعه التي لا تعانى من ن نقص فيتامين (د).

الاستنتاجات: نقص فيتامين (د) منتشر بين مرضى الاتهاب المفاصل الصدفى اكثر من الاصحاء. نقص فيتامن (د) مرتبط بزيلده مده المرض و اريتفاع معدل نشاط المرض و شدته.