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ORIGINAL RESEARCH

Relationship Between Depression and Quality of Life Among Vitiligo Patients: A Self-assessment Questionnaire-based Study

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Background: Vitiligo is a common chronic autoimmune disease that is characterized by progressive loss of skin color due to melanocyte destruction. In addition to the physical effects of vitiligo, this condition exerts adverse psychological effects and causes social stigmatization. Earlier studies reported that individuals with vitiligo suffer from poor quality of life (QoL), but data about the QoL of Thai vitiligo patients is scarce.

Objective: This study aimed to investigate the QoL and prevalence of depression in Thai vitiligo patients, association between QoL and depression, and factors associated with QoL and depression among Thai people with vitiligo.

Methods: This cross-sectional self-assessment questionnaire-based study was conducted at the phototherapy and vitiligo clinics of the Department of Dermatology, Faculty of Medicine Siriraj Hospital, Mahidol University (Bangkok, Thailand). QoL was assessed using the Dermatology Life Quality Index (DLQI) questionnaire, and depression was evaluated via the Patient Health Questionnaire (PHQ-9).

Results: Among the 104 vitiligo patients that were recruited, the mean DLQI score was 7.46, and the prevalence of depression (PHQ-9 \geq 9) was 13.5%. Factors significantly associated with a higher mean DLQI score were skin phototype IV compared to phototype III, active disease, new lesions within the last three months, lesions involving upper extremities, and PHQ-9 \geq 9 (p<0.05). Moderate correlation was found between DLQI score and PHQ-9 score (Pearson's correlation coefficient: 0.524, p<0.001). Factors significantly associated with depression were working status and developing new lesions within the last three months (p<0.05). Patients with new lesions were 4.12 times more likely to be depressed than those without new lesions (OR: 4.12, 95%CI: 1.20–14.16; p=0.025).

Conclusion: Developing new lesions, active disease, dark skin phototype (IV), and lesion on upper extremity had significant adverse effects on QoL. Vitiligo patients who were employed and who had new vitiligo lesions are significantly more likely to be depressed.

Keywords: relationship, depression, quality of life, QoL, vitiligo patients, self-assessment questionnaire-based study, Thailand

Introduction

Vitiligo is a common chronic autoimmune disease that is characterized by progressive loss of skin color due to melanocyte destruction. The worldwide prevalence of vitiligo in adults ranges from 1-2%.¹ The pathogenesis of vitiligo is multifactorial, and includes autoimmune, genetic, and environmental factors.² In addition to the physical effects of vitiligo, this condition exerts adverse psychological effects and causes social stigmatization.³ Moreover, the effects of vitiligo may be more pronounced among

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The psychological problems that can occur in vitiligo include depression, anxiety, obsessive compulsive disorder, and suicidal thoughts.9-11 A meta-analysis conducted by Wang et al, found that vitiligo patients were 4.96 times more likely to have depressive symptoms compared to controls.¹² Psychodermatology is a relatively new area of patient care that focuses on the interaction between the skin and the mind. Thus, in addition to treating the clinical symptoms of vitiligo, dermatologists can also play an important role in helping patients to psychologically adjust to, and accept, their condition. However, an understanding of each vitiligo patient's psychological status is needed so that therapies can be customized to meet the individual needs of individual patients. To date, little is known about depression and QoL among those affected by vitiligo in Thailand. Accordingly, the aim of the study was to investigate the QoL of and the prevalence of depression in Thai vitiligo patients, to evaluate for association between QoL and depression, and to identify factors significantly associated with QoL and depression among Thai people with vitiligo.

Materials and Methods

Study Design and Sample Size Calculation

This cross-sectional self-assessment questionnaire-based study was conducted at the phototherapy and vitiligo clinics of the Department of Dermatology, Faculty of Medicine Siriraj Hospital, Mahidol University, Bangkok, Thailand during May 2019 to October 2019. Siriraj Hospital is Thailand's largest super-tertiary referral center. This study was conducted in accordance with the Declaration of Helsinki and was approved by the Siriraj Institutional Review Board (SIRB) (COA no. Si259/2019). The sample size was calculated using data from a study by Wong and Baba.¹³ Using an estimated standard deviation of 5.17, an allowable error of 1, an alpha of 0.05, and a beta of 0.20, a minimum sample size of 104 patients was calculated.

Demographic and Clinical Characteristics of Vitiligo

Patients aged ≥ 18 years with dermatologist-diagnosed vitiligo were recruited. Written informed consent was obtained from all patients willing to join the study. Data, including gender, age, disease duration, skin phototype, occupation, education status, marital status, and thyroid disease, were collected and recorded. Type of vitiligo, current vitiligo activity, extent of lesions, and previous treatment were also documented. Active disease was defined as having one or more of these four clinical signs: confetti-like depigmentation, trichrome lesion, Köebner phenomenon, and/or inflammatory vitiligo.

Extent of vitiligo was assessed using Vitiligo Extent Score (VES). VES is a validated instrument that allows physicians to accurately and easily monitor the affected body surface area in a standardized way. Moreover, VES is considered a user-friendly and time-saving assessment tool.¹⁴ Vitiligo extent at each affected location is summed to calculate the total percentage of the body surface area that is affected by vitiligo.

Quality of Life and Depression Status

We used the Dermatology Life Quality Index (DLQI) questionnaire, which was designed by Finlay and Khan in 1994, to assess patient QoL.¹⁵ The DLQI questionnaire was translated into Thai language and validated by Kulthanan et ali.¹⁶ The DLQI questionnaire consists of 10 questions with six domains, including symptoms and feeling, daily activities, leisure, work and school, personal relationships, and treatment difficulties. DLQI score is stratified into the five following levels: 0–1=no effect, 2–5=small effect, 6–10=moderate effect, 11–20=very large effect, and 21–30=extremely large effect.¹⁷

Presence of depression was assessed using the Patient Health Questionnaire-9 (PHQ-9), which was translated into Thai language and validated by Lotrakul et al,¹⁸ and this assessment was reported to be a brief and user-friendly tool to improve the recognition of depression in primary care. The PHQ-9 is the major depressive disorder module of the full PHQ-9, and all nine components are scored 0 (not at all) to 3 (nearly every day).¹⁹ Lotrakul et al reported a PHQ-9 score \geq 9 to be the optimal cut-off score for diagnosing major depression in the Thai population, with a sensitivity of 0.84, specificity of 0.77, positive predictive value of 0.21, negative predictive value of 0.99, and positive likelihood ratio of 3.71. All patients were asked to complete both questionnaires by themselves.

Statistical Analysis

The data were analyzed using SPSS Statistics version 18.0 (SPSS, Inc., Chicago, IL, USA). Quantitative data were described using mean \pm SD, or median and range or IQR. Pearson's correlation coefficient was used to test association between two quantitative variables. Independent *t*-test, chi-squared test, and ANOVA were used to compare continuous and categorical data. Logistic regression analysis was used to calculate the OR to determine which factors may contribute to depression. All *p*-values <0.05 were regarded as being statistically significant.

Results

Demographic and Clinical Characteristics of the Study Population

A total of 104 individuals with vitiligo were included in this study. Table 1 summarizes the demographic and clinical characteristics of the study population. The mean age of patients was 41.6 ± 15.9 years, and most (70.2%) of them were female. The mean duration of disease was 8 ± 10.05 years. The most common type of vitiligo was nonsegmental vitiligo (76.9%), and most patients (56.7%) had Fitzpatrick skin phototype IV. Almost half of patients (49%) were suffering from active disease, and 42.3% of the patients had emergence of new lesions within the last three months. The mean VES was 3.3 ± 8.47 .

Dermatology Life Quality Index (DLQI)

The mean DLQI score was 7.46 ± 6.06 (range: 0–26). The three highest mean DLQI scores were from the questions relating to embarrassment (1.4), social activity (1.11), and clothing (1.01), consecutively (Figure 1). Table 2 shows the results of the analysis to identify factors significantly associated with QoL in vitiligo patients. That analysis

Table I Demographic Data and Clinical Characteristics of Patients with Vitiligo (n=104) $\,$

Patients with Vitiligo (n=104)	
Demographic Data	Total
Gender, n (%)	
Male	31 (29.8)
Female	73 (70.2)
Age, mean ±SD, years	41.55±15.92
Duration of disease, median (range), years	5.00 (0.16-60)
Marital status, n (%)	
Married	50 (48.1)
Single	49 (47.1)
Divorced	5 (4.8)
Education, n (%)	
Primary	15 (14.4)
Secondary	15 (14.4)
Undergraduate	57 (54.8)
Postgraduate	16 (15.4)
Doctoral	1 (1.0)
Occupation, n (%)	
Housewife	12 (11.5)
Public servant	17 (16.3)
Business owner	14 (13.5)
Office worker	19 (18.3)
Unskilled labor	14 (13.5)
Unemployed/retired	9 (8.7)
Student	19 (18.3)
Skin phototype, n (%)	
	29 (27.9)
IV	59 (56.7)
V	16 (15.4)
Thyroid diseases, n (%)	
Yes	9 (8.7)
No	95 (91.3)
Type, n (%)	
Segmental	24 (23.1)
Nonsegmental	80 (76.9)
Disease activity, n (%)	
Stable	53 (51.0)
Active	51 (49.0)
Poliosis, n (%)	
Yes	12 (11.5)
No	92 (88.5)
VES, median (range)	0.69 (0.01–58.37)
Previous treatment, n (%)	
Topical medication	100 (96.2)
Systemic medication	31 (29.8)
Phototherapy	85 (81.7)
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(Continued)

Table I (Continued).

Demographic Data	Total
Duration of treatment, median (range), month	19 (0–720)
DLQI, mean ±SD	7.46±6.06
DLQI, median (range)	6 (0–26)
PHQ-9, mean ±SD	4.09±4.49
PHQ-9, median (range)	3 (0–25)

Abbreviations: DLQI, Dermatology Life Quality Index; VES, Vitiligo Extent Score; PHQ-9, Patient Health Questionnaire-9.

revealed skin phototype IV compared to phototype III, active disease, new lesions appearing within the last three months, and lesions involving the upper extremities (excluding bony prominences) to be significantly associated with QoL. Regarding skin phototype, we found no significant difference between skin phototype V and other skin phototypes. Importantly, a high DLQI score was strongly significant associated with depressive status reflected in PHQ-9 \geq 9. We found no significant difference in higher DLQI score between women and men, between the working age group (age 30–59 years) compared to other age groups, employed patients compared to unemployed patients, location other than upper extremities, longer treatment duration, or VES extent.

Depression

The mean PHQ-9 score was 4.09 ± 4.49 (range: 0–25). Fourteen patients (13.5%) had PHQ-9 \geq 9, 65 cases had a PHQ-9 score of 0–4, and 25 cases had a score of 5–8 (mild) (Figure 2). When we compared the DLQI score between the depressive group (PHQ-9 \geq 9) and the PHQ-9 \leq 9 group, the three highest scores in the depressive group were the scores for social activities, embarrassment, clothing, shopping and homecare (Figure 1). This finding helps to explain the moderate correlation observed between the DLQI score and the PHQ-9 score (Pearson's correlation coefficient: 0.524, *p*<0.001) (Figure 3).

Table 3 summarizes the factors significantly associated with depression in vitiligo patients. We found a significantly higher prevalence of depression in employed individuals than students, unemployed, or retired people (p=0.046). Depression was considerably more frequent in patients with active vitiligo than those with stable disease. In addition, individuals with new lesions appearing within the last three months were significantly more likely to be depressed than those with no new lesions (p=0.018), and logistic regression analysis showed patients with new lesions to be 4.12 times more likely to show depression compared to those without new lesions (OR: 4.12, 95%CI: 1.20–14.16; p=0.025).

There were no statistically significant differences in PHQ-9 score for gender, age group, education level, skin phototype, or type of vitiligo. Nonetheless, depression was not associated with involved locations, treatment duration or VES.

Discussion

Current pathogenesis of vitiligo is complex, mainly involved heat shock protein 70, interferon-gamma, JAK-STAT, CXCL10, effector and memory CD8⁺ T cell lymphocyte.²⁰ Treatment of vitiligo included topical corticosteroid, calcineurin inhibitors, systemic immunosuppressants, UVB





 Table 2 Factors Associated with Quality of Life in Patients with

 Vitiligo

Factors	Mean DLQI ±SD	þ value
Gender		
Male (n=31)	6.45±6.10	0.270
Female (n=73)	7.89±6.04	
Age (years)		
<30 (n=28)	6.79±4.92	0.133
30-59 (n=63)	8.10±6.59	
≥60 (n=13)	5.85±5.57	
Education		
Below undergraduate (n=30)	6.73±5.07	0.438
Undergraduate and above (n=74)	7.76±6.43	
Occupation		
Unemployed/retired/students	6.15±5.26	0.081
(n=40)		
Employed (n=64)	8.28±6.42	
Skin phototype		
III (n=29)	5.38±4.31	0.032*
IV (n=59)	8.92±6.55	
V (n=16)	5.88±5.69	
Types of vitiligo		
Segmental (n=24)	8.38±7.13	0.403
Non-segmental (n=80)	7.19±5.73	
Disease activity		
Stable (n=53)	6.21±5.54	0.031*
Active (n=51)	8.76±6.36	
New lesion in three months		
Yes (n=44)	9.09±6.36	0.018*
No (n=60)	6.27±5.59	
Involved area		
Head and neck		
Yes (n=75)	7.59±6.11	0.737
No (n=29)	7.14±6.03	
Trunk		
Yes (n=60)	8.12±6.07	0.200
No (n=44)	6.57±6.01	
Acral		
Yes (n=68)	7.78±5.83	0.465
No (n=36)	6.86±6.52	
Upper extremities		
Yes (n=32)	9.53±6.53	0.020*
No (n=72)	6.54±5.65	
Lower extremities		
Yes (n=34)	7.74±6.06	0.750
No (n=70)	7.33±6.10	

(Continued)

Table	2	(Continued)
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Factors	Mean DLQI ±SD	p value
Bony prominence		
Yes (n=26)	8.15±5.64	0.504
No (n=78)	7.23±6.22	
Mucosal		
Yes (n=29)	6.72±5.34	0.443
No (n=75)	7.75±6.33	
Treatment duration		
Less than one year (n=30)	8.30±7.07	0.668
I–2 years (n=42)	7.19±5.99	
More than two years (n=32)	7.03±5.19	
VES		
VES <5 (n=89)	7.27±6.15	0.434
VES ≥5 (n=15)	8.60±5.55	
PHQ-9 (Depression; PHQ-9 ≥9)		
PHQ-9 <9 (n=90)	6.29±5.22	<0.001*
PHQ-9 ≥9 (n=14)	15.00±5.83	

Note: *Significant variables.

Abbreviations: DLQI, Dermatology Life Quality Index; VES, Vitiligo Extent Score; PHQ-9, Patient Health Questionnaire-9.

phototherapy and surgery.^{21–23} Novel targeted therapy focused on JAK inhibitor with some success.²¹ However, no current treatment can cure the disease. After treatment discontinuation, vitiligo lesions frequently return, supporting the hypothesis of autoimmune memory at vitiligo location.²⁴ Disease stabilization and stimulation of repigmentation are the main concepts of treatment and should be performed as early as first lesions appear. Prognosis is correlated with disease location and onset of treatment. Due to the unpredictable clinical course and long-standing treatment, vitiligo undoubtedly affects the psychosocial impact state and QoL.

Quality of Life

Earlier studies have reported mean DLQI scores of vitiligo ranging from 1.82 to 15.⁴ Our study showed mean DLQI score of 7.46 which represented a moderate effect on patients' QoL. This was similar to QoL in studies by Dolatshahi et al and Wang et al.^{25,26} The three main effects were embarrassment, social activities, and choice of clothing which were similar to previous studies.^{7,27} When compared to other dermatologic conditions in Thailand, vitiligo DLQI is higher than fungal



Figure 2 Severity of depression according to Patient Health Questionnaire (PHQ)-9 scoring among Thai vitiligo patients.

infection $(6.5)^{28}$ but lower than seborrheic dermatitis $(8.1)^{29}$ and acne $(10.6)^{.28}$

Amer and Gao demonstrated that individuals with vitiligo in Asian countries such as India and Saudi Arabia seemed to have a higher DLQI score compared to those in Caucasian countries due to darker skin phototypes and differences in culture.⁴ Our study found individuals with skin phototype IV had significantly higher DLQI than those with phototype III (p=0.032), but lack of significant differences between skin phototype V and type III and IV. Dolatshahi et al, found that patients with skin phototype V had significantly poorer QoL than in those other types (p<0.001) due to the contrast between vitiligo lesions and darker skin.²⁵ These can be explained from the small number of our patients with skin phototype V.

Active disease is clinically defined as rapid spreading of news lesions, confetti-like depigmentation, trichrome and Köebner phenomenon. Active vitiligo should receive prompt and aggressive treatment to halt and prevent further spreading of depigmentation. This mainly included immunosuppressants and phototherapy. Our study demonstrated individuals with active vitiligo and patients who developed new lesions within the last three months had significantly higher DLQI score denoting a higher impact on their QoL than those with stable vitiligo and no new lesions. These were also observed by Karelson et al, in which the DLQI score was significantly associated with active stage of the disease (p=0.006).³⁰ The rapid spreading of disease can turn to widespread lesions and prognosis is unpredictable. As a result, the activity of the disease greatly impaired QoL.

Based on the location, our study showed that individuals with vitiligo lesions on upper extremities, which are visible parts, were significantly associated with poorer QoL Previous studies supported the concept of lesions occurring on visible or exposed regions of the body having significantly higher DLQI scores than those without lesions.^{13,27} In addition, Silverberg and Silverberg's study showed that lesions on the arms were significantly associated with self-consciousness about vitiligo,⁷ and therefore visible lesions could affect patients' confidence and disturb QoL.

Our study showed that vitiligo patients with depression had significantly higher DLQI scores than those without depression (p<0.001). This relationship was also observed by Chan et al, where vitiligo patients with a DLQI score of six and above were 4.88 times more likely to develop depression compared to those with a score of 0–5.³¹ In addition, we also found that the DLQI score displayed moderate correlation with the PHQ-9 score. QoL impairment would probably affect patients' mood and eventually they would develop depressive symptoms.

We found a trend of QoL impairment in female than male patients, although lack of significant effect, similar to previous studies.^{13,25} Women have more cosmetic concerns than men, regardless of age groups, socioeconomic and living status, and therefore seek more treatment.³²

Our study also showed a trend of higher mean DLQI score among individuals who were working than those who were students, unemployed or had retired but we did not find statistical significance between the two groups. However, Wong and Baba's study revealed that patients who had jobs



Figure 3 Box plot showing Patient Health Questionnaire (PHQ)-9 scoring compared between the PHQ <9 group and the PHQ-9 ≥9 group relative to Dermatology Life Quality Index (DLQI) score.

Notes: The o symbol represents score in one case of patient that is not in the bar range.

had significantly higher DLQI scores compared to those who were retired. People who were employed or had to go out for work and meet other people might feel more embarrassed, therefore, resulting in reduction of QoL.¹³ Regarding age groups, age 30–59 years which was a working age group also seemed to have higher DLQI scores compared to other groups (Table 2).

Depression

The prevalence of depression among our vitiligo patients based on PHQ-9 \geq 9 was 13.5%, which was lower than earlier studies.^{33–35} Most of our patients had none or mild depressive symptoms similar to a study from Iran.¹¹ Prevalence of depression and depressive symptoms varies among the studies depending on the diagnostic criteria or instruments used.¹² Reported screening instruments included Hamilton Depression Rating scale (HDRS) and Beck Depression Inventory (BDI).¹² However, Dabas et al, by using PHQ-9, found that the prevalence of depression among their vitiligo population in India was 27.4% which was higher than that in our study.³⁶ The reason for this could be that vitiligo lesions might be more noticeable on darker skin color. It should be noted that the sensitivity of PHQ-9 is high but the positive predictive value of PHQ-9 is low. Therefore, clinical assessment by psychiatrists is recommended.

We observed that depression was significantly associated with working status of the patients (p=0.046) and significantly associated with developing new lesions with the last three months (p=0.018). Individuals with new lesions developing in three months were 4.12 times more likely to have depression compared to those without new lesions (OR: 4.12, 95%CI: 1.20–14.16; p=0.025). This could be explained by the rapid spreading and the patients might feel more concerned about the occurrence of new lesions since the disease has no cure.

According to the earlier meta-analysis, the prevalence of depression in women with vitiligo was significantly higher than that in men.¹² However, we found no statistically significant differences in depression between genders, age groups, skin phototypes, types of vitiligo and involved areas.

We found a significant correlation between DLQI score and PHQ-9 in vitiligo. Patients whose QoL was affected had depressive symptoms and vice versa. These were similar to

Table 3	Factors	Associated	with	Depression	Among	Patients
with Vitil	igo					

Factors	Number of Cases PHQ-9 < 9	Number of Cases PHQ-9 ≥ 9	p value
	.,	~ /	
Gender			
Male	27	4	1.000
Female	63	10	
Age group			
Age <30	26	2	0.097
Age 30–59	51	12	
Age ≥60	13	0	
Education			
Below undergraduate	27	3	0.752
Undergraduate and	63	11	
above			
Occupation			
Unemployed/retired/	38	2	0.046*
students		-	0.0 10
Employed	52	12	
Skin phototype			
	27	2	0.234
IV	51	8	
V	12	4	
Types of vitiligo			
Segmental	20	4	0.733
Nonsegmental	70	10	
Disease progression			
Stable	49	4	0.072
Active	41	10	
Neurise is there are the			
New lesion in three months	34	10	0.018*
Yes No	56	4	0.018
	50	•	
Involved area			
Head and neck			
Yes	63	12	0.340
No	27	2	
Trunk			
Yes	49	11	0.089
No	41	3	
Acral	50		0.741
Yes	58	10	0.766
No	32	4	
Upper extremities Yes	28	4	1.000
No	62	4	1.000
Lower extremities	V2		
Yes	31	3	0.541
No	59	11	0.5 11
Bony prominence			
Yes	24	2	0.509
No	66	12	

(Continued)

Table 3 (Continued).

Factors	Number of Cases PHQ-9 < 9	Number of Cases PHQ-9 ≥ 9	p value
Mucosal			
Yes	24	5	0.527
No	66	9	
Treatment duration			
Less than one year	27	3	0.388
I–2 years	34	8	
More than two years	9	3	
VES			
VES <5	76	13	0.405
VES ≥5	14	1	

Note: *Significant variables.

Abbreviations: PHQ-9, Patient Health Questionnaire-9; VES, Vitiligo Extent Score.

previous studies.^{36,37} Therefore dermatologists should be concerned and keep in mind depressive symptoms when they detect and recognize the high DLQI score. Coping with depressive symptoms could improve QoL.

Nowadays, psychodermatology and liaison psychiatry clinics play an important role in treating and supporting patients with chronic intractable dermatologic conditions, including vitiligo. The multidisciplinary approach provides the patients with an individualized holistic care with the cooperation of dermatologist, psychiatrist and other specialized personnel. In addition to treating preexisting psychiatric diseases, psychiatrists also play a critical role in early detection of mild psychiatric conditions, especially depression. They can provide early interventions to prevent the progression of mild conditions to serious psychiatric disorders.

Few studies focused on treatment for depression in individuals with vitiligo. Papadopoulos et al found that cognitive behavioral and person-centered therapies in patients with vitiligo can significantly improve patients' ability to cope with the disease assessed by the General Health Questionnaire.³⁸ Thus, early psychological intervention might be helpful to prevent depression and enhance patients' coping mechanism to improve their QoL.

Limitations

This study was conducted in a single tertiary-care hospital which might not represent all Thai individuals with vitiligo or might not be generalizable to other countries. Although PHQ-9 is the standard measurement for screening of depression, its limitation is that it is a self-rated questionnaire.

Additional clinical information, eg, patient's function, effects of medication, illness, and psychiatric evaluation are recommended for definite diagnosis and comprehensive evaluation. Moreover, patients were informed that they were tested for depression and QoL before answering the questionnaire and thus their responses may be biased.

Conclusion

Vitiligo showed moderate QoL impairment in the Thai population and moderate prevalence of depression. These two factors are well correlated. Development of new lesions, active disease, dark skin phototype (IV) and lesions on upper extremities had significant effects on QoL. In addition, new vitiligo lesions and employed workers had risks of depressive symptoms. Vitiligo patients who were working or in active stage of the disease had a significant impairment in QoL and were associated with depression. These patients might need an appropriate health-care consultation, more aggressive interventions to halt disease progression and regular screening for depression. Liaison psychiatry clinics can play an important role in supporting patients with vitiligo and dermatologist should take part in this psychosocial issue to provide a holistic approach for this chronic, visible, and unpredictable condition.

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Disclosure

The authors report no conflicts of interest in this work.

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