Thyroid Dysfunction in Libyan Vitiligo Patients

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Abstract
Background: Vitiligo is an acquired depigmenting disorder due to destruction of melanocytes, many hypotheses have been suggested for its pathogenesis. One of these hypotheses suggest that there are autoimmune and endocrine dysfunction involvement. The involvement of vitiligo with thyroid autoimmune diseases, with the increased prevalence of auto antibodies including thyroid auto antibodies in vitiligo support the hypothesis.

Patients and Methods: 50 Libyan patients under same age and gender with vitiligo, and 50 controls. Patients were excluded if they had a history of thyroid, or other autoimmune diseases. Data on age, onset of illness, duration and disease activity were determined. Serum T3, T4, TSH, and antibodies to TPO and TG were measured in both vitiligo patients & controls. All patients and control subjects underwent thyroid ultrasonography.

Results: Fifty patients with vitiligo and their 50 matched controls were studied. More than half of the patients (52%) were females and 48% were males, their mean of age was 40 ± 11 years, and the duration of vitiligo was 11± 9 years. Vitiligo vulgaris type was the most common form seen in 68% of the patients, and 42% reported at least one family member affected with vitiligo. Family history of thyroid disorder was seen in 20% of the patients. Thyroid functional abnormalities were significantly seen more in patients than control subjects. The frequency of TG and TPO thyroid autoantibodies was significantly higher in vitiligo patients than in healthy controls (P < 0.01). Abnormal thyroid ultrasound study was seen in 18 (36%) of the patients compared to 6 (12%) of the control subjects (P < 0.05).

Conclusion: Our findings pointed to a significant association between vitiligo and thyroid autoimmunity and showed that testing the level of thyroid autoantibodies is relevant in vitiligo patients.

Keywords: Thyroid; Autoimmune, Vitiligo, Melanocyte.

INTRODUCTION

Vitiligo is the most common, probably heritable, progressive depigmenting skin disorder caused by destruction of melanocytes. It is characterized by well-demarcated white patches of skin. Depending on the extent of the lesions, vitiligo can be classified into two main categories: generalized and localized. It affects 0.5-2% of the general population with no age, sex, or racial predilection [1]. Many different etiologic hypotheses have been presented for vitiligo. The most recent of these hypotheses supports the combination of environmental and genetic factors interacting to contribute to autoimmune melanocytes destruction [1, 2]. Several studies reported that there is an epidemiological association between generalized vitiligo and other autoimmune diseases; such as autoimmune thyroid diseases, pernicious anemia, alopecia areata and Addison's disease [3-5]. The aim of the study was to evaluate the association between vitiligo and thyroid autoimmune diseases.

Aim of the study: was to determine whether vitiligo is associated with thyroid autoimmunity.

PATIENTS AND METHODS

The aims of the study are to evaluate the association of vitiligo with thyroid autoimmune diseases patients. A random 50 healthy controlled trial was conducted on 50 patients with vitiligo and a
written consent was obtained after explaining the nature and possible consequences of the study to all patients attending at the Dermatology department of El-Jumhuriya hospital in Benghazi city. Patients and control subjects were excluded if they had a history of thyroid, or other autoimmune diseases. The diagnosis of vitiligo was mainly clinical.

A detailed history including the age of onset, duration of the disease, localization of the lesions, disease activity and presence of Koebner phenomenon, associated diseases and family history were obtained from each patient.

Laboratory investigations for fasting blood samples were collected between 08:30 and 10:00 h, after an overnight fasting of more than 12 hours. The blood samples were analyzed for total triiodothyronine (T3), total thyroxine (T4), thyroid stimulating hormone (TSH), thyroid peroxidase (TPO) antibody and thyroglobulin (TG) antibody. Thyroid gland sonography was performed for all patients and control subjects by the same radiologist.

STATISTICAL ANALYSIS

Data were fed to computer using Statistical Package for Social Sciences (SPSS) version 11.5 and reviewed. Descriptive statistics in the form of percentages mean and standard deviation of different parameters was used. Chi square test was used to compare qualitative parameters, P < 0.05.

RESULTS

Fifty patients with vitiligo and their 50 matched controls were studied. Of the 50 vitiligo patients, 26 (52%) are females and 24 (48%) are males, with a mean age of 40 years and a mean age of onset of the disease ± SD was 21 ± 7 years. The mean duration of vitiligo ± SD was 19 ± 10 years (range: 1 - 45 years).

The distribution pattern of the lesions, denoting the clinical type of vitiligo, is vitiligo vulgaris was the most common type is 34 (68%), followed by universal 7 (14%), acrofacial 6 (12%) and focal vitiligo 3(6%).

50 cases, 21 (42%) had a family history of vitiligo, first degree relatives (parents/siblings) were affected in 18 (36%) of the patients and second degree relatives (grandparents, uncles or aunts) in 3 (6%) of the patients. Ten (20%) of the patients had family history of thyroid diseases. Six (12%) of the patients had family history of hypothyroidism, 2 (4%) had hyperthyroidism, and 2 (4%) had family history of goiter.

Thyroid functional abnormalities were seen in 20 (40%) of the patients compared to 8 (16%) of the control subjects and this difference was statistically significant (Table 2).

The T3 and T4 were within normal range in 42 (84%) and 44 (88%) of the patients, and 47 (94%) and 47 (94%) of control group, respectively, however, no significant difference was detected between the two groups (Table 3).

The frequency of high TSH was 26% in vitiligo patients and 8% in control subjects and this difference was statistically significant (P = 0.016) (Table 3). None of the patients or control subjects had low TSH level.

Anti-TG antibody in 11 (22%), and anti-TPO antibody in 10 (20%) of the patients were higher than the normal antibodies titer. Moreover, 5(10%) of the patients had both anti-TG and anti-TPO antibodies higher than the normal antibody titer. In the control group, one subject (2%) had positive anti-TG and none had positive anti-TPO (Table 4). This difference was statistically significant.

Abnormal thyroid ultrasound study was seen in 18 (36%) of the

| Type of vitiligo       | No. (%) |
|------------------------|---------|
| Vitiligo vulgaris      | 34 (68%)|
| Vitiligo universalis   | 7 (14%) |
| Acrofacial vitiligo    | 6 (12%) |
| Focal vitiligo         | 3 (6%)  |

| Normal TFT | Abnormal TFT |
|------------|--------------|
| Vitiligo patients | 30 (60%) | 20 (40%) |
| Control subjects  | 42 (84%) | 8 (16%) |
| P value           | 0.007**      |

| Vitiligo patients | Control subjects | P value |
|-------------------|------------------|---------|
| T3 Normal 43 (84%) | 47 (94%) | 0.1     |
| T4 Normal 44 (88%) | 47 (94%) | 0.4     |
| TSH High 13 (26%)  | 4 (8%)   | 0.01*   |
patients compared to 6 (12%) of the control subjects (P < 0.05) (Table 5).

**DISCUSSION**

Vitiligo is an acquired pigmentary disorder of unknown etiology characterized by depigmentation of the skin resulting from the loss of melanocytes. The prevalence of the disease ranges from 0.1% to 8% with female predominance [1]. The onset of the disease usually occurs before 20 years of age. Generalized vitiligo is the most common clinical presentation and often involves the face and extremities. The course of the disease is unpredictable, with a severe psychological impact on the patient quality of life and the response to treatment varies [1].

The pathogenesis of vitiligo is complex and involves the interplay of a series of variables [5, 6]. There is a multifactorial genetic component predisposing certain individuals to vitiligo and family history is a variable found in approximately one-third of the people with the disease [7]. There is also strong genetic evidence of a link between vitiligo and other autoimmune diseases [8].

The autoimmune hypothesis of vitiligo proposes that an immune system disorder results in the destruction of melanocytes. The epidemiological association with several autoimmune diseases is one of the various evidence which support an autoimmune origin of vitiligo [2, 3]. Among these diseases, lichen planus, pernicious anemia, Hashimoto’s thyroiditis, hypothyroidism, endemic goiter, diabetes mellitus, lupus erythematosus, and Graves’ disease. However, the chance of coexistence with autoimmune thyroid diseases is higher than the others [9-12].

In 1941, Robert suggested that vitiligo might be connected with an increased activity of the thyroid gland; and reported a distinct rise of the basal metabolism in 10 out 20 vitiligo patients tested [3]. Later, several studies reported a significantly increased prevalence of autoimmune thyroid disease in vitiligo patients; the rate of positivity of thyroid autoantibodies varied from 2.2% to 50% [2, 4].

In the present study Anti-TG antibody was found in 22%, and anti-TPO antibody was found in 20% of the patients. Our results are consistent with previous studies [11, 13, 14]. A recent study of 82 vitiligo patients demonstrated positive anti-TG titers in 23% of the patients and anti-TPO antibody were positive in 24.1% of the patients, and the results were found significantly higher when compared to healthy controls [14]. Moreover, study of Kasumagic-Halliovic et al., found high levels of anti-TPO in 11 (27%) out of 40 vitiligo patients and they demonstrated significantly elevated levels of anti-TPO compared to the control group [11].

Daneshpazhooh et al., measured the serum level of anti-TPO antibody and reported significantly high levels in vitiligo patients compared to healthy controls [10]. Moreover, a study was carried out in India revealed that the anti-TPO antibody was positive in 31.4% cases [15].

However, Gulec Saylam Kurtipek, et al., found high levels of anti-TPO in 16 (14.8%) and anti-TG in 9 (8.3%) out of 106 vitiligo patients, these rates are lower than our study [16]. In addition, previous studies reported a significantly increased prevalence of vitiligo in patients with autoimmune thyroid disease compared to patients with non-autoimmune thyroid disease [10, 16].

In present study, thyroid dysfunction manifesting as hypothyroidism occurred in 26% of the patients and in 8% of controls, this difference being statistically significant. A high prevalence of hypothyroidism in vitiligo patients has been reported by Kumar et al., (40%) and Akay et al., (31%) [17, 18]. However, a lower occurrence was noted in other study [19]. Although, previous study have noticed an association of vitiligo with hyperthyroidism, these findings were not noticed in our study [20].

**CONCLUSION**

The study revealed a significant association between vitiligo and thyroid autoimmunity and screening vitiligo patients for thyroid autoantibody seems plausible.

**REFERENCES**

[1]. Alkhan A, Felsten LM, Daly M, Petronic-Rosic V (2011) Vitiligo: A comprehensive overview Part I. Introduction, epidemiology, quality of life, diagnosis, differential diagnosis, associations, histopathology, etiology, and work-up. J Am Acad Dermatol. 65(3): 473–91.

[2]. Al-Mutarini N, Sharma A (2006) Profile of vitiligo in Farwaniya region in Kuwait. Kuwait Med J. 38(2): 128–131.

[3]. Robert P (1941) Ueber die vitiligo. Dermatology. 84: 317–319.

[4]. Mandy RC, Ortiz LJ, Lugo-Somolinos A, Sánchez JL (1996) Organ-specific autoantibodies in vitiligo patients and their relatives. Int J Dermatol. 35(1): 18–21.

[5]. Schallreuter KU, Bahadoran P, Picardo M, Slominski A, Eliausi Y, et
El-Dibany SA, El-Sherif NA, Greiw A, Matmati NA, Belkhair N (2017) Thyroid Dysfunction in Libyan Vitiligo Patients. Int J Clin Dermatol Res, S2:002, 4-7.

[6]. Richmond JM, Frisoli ML, Harris JE (2013) Innate immune mechanisms in vitiligo: danger from within. Current Opinion in Immunology. 25(6): 675–682.

[7]. Faris AR, Mira MT, Tarl’e RG, Silva de Castro CC, Dellatorre G (2014) Vitiligo-part 2-classification, histopathology and treatment. An Bras de Dermatologia. 89(5): 784–790.

[8]. Spritz RA (2013) Modern vitiligo genetics sheds new light on an ancient disease. J Dermatol. 40(5): 310–318.

[9]. Niepomniszcze H, Amad RH (2001) Skin disorders and thyroid diseases. J Endocrinol Invest. 24(8): 628–638.

[10]. Daneshpazhooh M, Mostofizadeh GM, Beljati J, Akhyani M, Robati RM (2006) Anti-thyroid peroxidase antibody and vitiligo: a controlled study. BMC Dermatology. 6: 3.

[11]. Kasumagic-Halilovic E, Ovcina-Kurtovic N, Jukic T, Karamenih J, Begovic B, et al., (2013) Vitiligo and autoimmunity. Med Archiv. 67(2): 91–93.

[12]. Vrijman C, Knoone MW, Limpens J, Leeflang MM, Luiten MM, et al., (2012) The prevalence of thyroid disease in patients with vitiligo: a systematic review. Br J Dermatol. 167(6): 1224–1235.

[13]. Sedighie M, Gholamhossein G (2008) Thyroid dysfunction and thyroid antibodies in Iranian patients with vitiligo. Indian J Dermatol. 53(1): 9–11.

[14]. Yang Y, Huang G, Yan X, Qing Z (2014) Clinical analysis of thyroglobulin antibody and thyroid peroxidase antibody and their association with vitiligo. Indian J Dermatol. 59(4): 357–360.

[15]. Dave S, Dsouza M, Thapp DM, Reddy KS, Bobby Z (2003) High frequency of thyroid dysfunction in Indian patients with vitiligo. Indian J Dermatol. 48(2): 68–72.

[16]. SaylamKuruppek G, Cihan FG, EraymanDemirbas S, Ataseven A (2015) The Frequency of Autoimmune Thyroid Disease in Alopecia Areata and Vitiligo Patients. BioMed Res Int. ID : 435947.

[17]. Kumar KV, Priya S, Sharma R, Kapoor U, Saini M, et al., (2012) Autoimmune thyroid disease in patients with vitiligo: Prevalence study in India. Endocr Pract. 18(2): 194–9.

[18]. Akay BN, Berkir M, Anadolu Y, Gullu S (2010) Epidemiology of vitiligo, associated autoimmune diseases and audiological abnormalities. Ankara study of 80 patients in Turkey. J Eur Acad Dermatol Venereol. 24(10): 1144–50.

[19]. Narita T, Otso N, Fukai K, Kabashima K, Kawada A, et al., (2011) Generalized vitiligo and a associated autoimmune diseases in Japanese patients and their families. Allergol Int. 60(4): 505–8.

[20]. Iacovelli P, Sinagra JL, Vidolin AP, Marenda S, Capitanio B, et al., (2005) Relevance of thyroiditis and of other autoimmune diseases in children with vitiligo. Dermatology. 210(1): 26–30.

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