Demodex Parazites in Schizophrenia

Mehmet Hanifi Kokaçya1, Berna Hamamcı2, Ümit Sertan Çöpoglu1, Özlem Aycan Kaya2
1Dept. of Psychiatry, Faculty of Medicine, 2Dept. of Parazitology, Faculty of Medicine, Mustafa Kemal University, Hatay, Turkey

Aim: Demodex parazites are commonly present all over the world, especially in facial region of humans. Demodex spp. are assumed to be more common in schizophrenia due to partial suppression of immune system and lack of good self-care. The present study aimed to investigate the prevalence of Demodex ectoparasites in schizophrenia patients. Material and Method: In the study, 31 patients with a diagnosis of schizophrenia and 30 subjects without any psychiatric disorder or skin disease were subjected to standard superficial skin biopsy technique to determine Demodex spp. Results: Demodex spp. were found positive in nine schizophrenia patients and it was found positive in two healthy controls. Considering the prevalence of Demodex spp., a significant relationship is found between schizophrenia patients and normal controls (p<0.05). Discussion: As a result, Demodex spp. are more common in schizophrenia due conditions of reduced self-care and immunosuppression, Demodex parasites should be considered in schizophrenia patients with skin lesions, especially on the face, and should to be treated if needed.

Keywords
Schizophrenia; Immune System; Ectoparasitic Infestations

Özet

Anahtar Kelimeler
Şizofreni; Imun Sistem; Ektoparasitik İnfestasyonlar

DOI: 10.4328/JCAM.3855 Received: 25.08.2015 Accepted: 15.09.2015 Printed: 01.02.2016 J Clin Anal Med 2016;1(suppl 1): 6-9 Corresponding Author: Mehmet Hanifi Kokacya, Department of Psychiatry, Faculty of Medicine, Mustafa Kemal University, 51100, Hatay, Turkey. GSM: +905052585954 F.: +90 3262294556 E-Mail: mhkokacya@mku.edu.tr
Introduction
Demodex species are commonly seen mites in the facial region of humans and widespread all over the world. Although there are numerous species of Demodex, two kinds of them can settle in the human body [1]. Demodex folliculorum and Demodex brevis live on the pilosebaceous glands and hair follicles and transmit from human to human by close contact interactions and result in infestation [2]. Demodex folliculorum is the most common ectoparasite found in humans [3]. For the diagnosis, cellophane-tape, skin scraping, punch biopsy and standard superficial skin biopsy methods are used [4]. Pathogenicity of Demodex spp. is still a debate and symptoms are very rare. It has been shown that Demodex spp. can cause severe infections only in immunosuppressed and middle and older-aged people [5]. Schizophrenia is a chronic and debilitating neuropsychiatric disorder that affects approximately 1 % of world population [6]. The main clinical symptoms of schizophrenia include: delusions and hallucinations, thought, speech and behaviour disorders, social withdrawal, loss of cognitive skills. Impairment in daily life skills, reduced self-care and weak immune system are the main symptoms reported for the disease [7]. The underlying neurobiological mechanisms of this disorder remains unclear however there are growing of evidence that oxidative stress and immune system abnormalities play an important role in the etiology. In one third of patients with schizophrenia shows immunological abnormalities such as an altered cytokine profile in serum and cerebrospinal fluid [6]. As immune system abnormalities, lack of hygiene and self care are seen in schizophrenia, it is hypothesized that Demodex spp. can be more prevalent in schizophrenia patients. To our knowledge, there have not been any reports focusing on the prevalence of Demodex spp. mites in patients with schizophrenia. This study aimed to identify the prevalence of Demodex spp. in patients diagnosed with schizophrenia.

Subjects and Method
Patients
Of 59 consecutive outpatients, aged 18 to 60 years, who had applied to the Mustafa Kemal University, School of Medicine, Research and Training Hospital Outpatient Psychiatry Clinic and had been diagnosed with schizophrenia according to Diagnostic and Statistical Manual of Mental Disorders IV-TR (DSM-IV-TR) between September 2013 and January 2014, 31 agreed to participate in the study. The control group included 30 healthy volunteers with no signs of skin disease. Ethics approval for this study was obtained from local ethic committee in accordance with the Helsinki Declaration. All patients provided written informed consent for participation in this research. Exclusion criteria were to be pregnant or breastfeeding, to have a chronic disease like hypertension, diabetes mellitus etc.., to use immunosuppressive agents and to have immunodeficiency syndromes like AIDS. The patients with any kind of comorbid dermatological or psychiatric disorder were excluded. All participants were subjected to a survey and samples were collected from participants by standard skin surface biopsy method using an adhesive band containing cyanoacrylate. Sample collection from study participants were achieved by taking samples from five different regions of their faces (nose, forehead, cheek, jaw). Before samples were taken, it was guaranteed that there was no cream or lotion on the faces of the patients and that the region where the samples would be taken was dry. Also, the region was cleaned with alcohol. To determine number of Demodex in cm² samples were examined using Hoyer solution and observed under the light microscope at x400 and x1000 magnifications. Five of more Demodex in cm² were accepted as positive for the Demodex.

Statistical Analysis
The SPSS 18.0 (Statistical Package for Social Sciences Chicago,IL, USA) package program was used for the statistical analysis. Descriptive and analytic statistics were performed. Chi-square test was used to compare categorical variables. Continuous variables were presented as mean ± standard deviation p <0.05 was considered as statistically significant.

Results
Demodex spp. positivity was evaluated in terms of age and gender in patients diagnosed with schizophrenia. 14 female and 17 male schizophrenia patients aged between 18 and 60 years (mean: 36.64 ± 5.11 years ) and as a control group 12 females and 18 males aged between 19 and 52 years (mean: 32.44 ± 8.58 years ) were enrolled. The gender distribution of schizophrenia patients was similar to the control group (Table 1). Demodex spp. was detected on the faces of 29.03% (n=9) of patients diagnosed with schizophrenia and 9.5% (n=2) of the control group. (Table 2)

Table 1. Gender distributions of the schizophrenia and control groups.

<table>
<thead>
<tr>
<th>Gender</th>
<th>Schizophrenia</th>
<th>Control</th>
<th>p*</th>
</tr>
</thead>
<tbody>
<tr>
<td>Male</td>
<td>17(54.8)</td>
<td>18(60)</td>
<td>0.797</td>
</tr>
<tr>
<td>Female</td>
<td>14(45.2)</td>
<td>12(40)</td>
<td></td>
</tr>
<tr>
<td>Total</td>
<td>31(100)</td>
<td>30(100)</td>
<td></td>
</tr>
</tbody>
</table>

* chi-square test.

Table 2. Incidence of Demodex spp. in patients with schizophrenia and control groups according to gender.

<table>
<thead>
<tr>
<th>DIAGNOSIS</th>
<th>Demodex spp. Positive</th>
<th>Demodex spp. Negative</th>
<th>Total</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>N</td>
<td>%</td>
<td>N</td>
</tr>
<tr>
<td>Schizophrenia</td>
<td>Male: 17</td>
<td>6</td>
<td>35.3</td>
</tr>
<tr>
<td>Control</td>
<td>Male: 18</td>
<td>1</td>
<td>5.55</td>
</tr>
<tr>
<td></td>
<td>Female: 12</td>
<td>1</td>
<td>8.33</td>
</tr>
</tbody>
</table>

The existence of Demodex spp on the faces of patients diagnosed with schizophrenia was found to be significantly higher compared to the control group (p=0.043) (Table 3). Incidence of Demodex spp. was found to be 5.7 times higher in schizophrenia as compared to control group (Table 3). Positive results were observed mostly in nose area and the second most common region was determined as forehead. In one patient, all of the regions were positive. Additionally, forehead area was determined as the region that has the highest parasite concentration (Table 4).
Discussion

Demodex folliculorum and Demodex brevis are obligate parasites that only originates in human hair follicle and pilosebaceous unit. Demodex spp. normally present in intact skin, hair follicles, and sebaceous glands without any pathogenic effect. However, in conditions of poor hygiene and suppressed immune system, Demodex spp. can cause inflammatory dermatitis, keratitis and epitelio, acne and acne rosacea [8]. The density of Demodex spp. increases with age [9]. Aycan et al. investigated the incidence of Demodex spp. in various types of disease states and age groups [4]. They used standard biopsy technique and observed 92 (63.5) positive results among 172 patients that were 21 years old age and older. In our study, we found nine (63.5) positive among 31 patients whose mean age was 36.64. In control group two subjects were positive (6.7) among 30 controls whose mean age was 32.44. There was no significant relationship between the incidence of Demodex spp. and age.

There are several investigations about gender and Demodex spp. Kokacya et al have investigated Demodex spp. Prevalence in 63 depressive patient They found four positive results in 27 male (26.7%) and 11 (34.2%) positive results in 36 female patient. They found no significant difference between gender and Demodex spp. prevalence [2].

In the present study, we found six (35.3) positive among 17 males and 3 (21.4) positive among 14 females. These results were not statistically significant (p<0.05). Although, these results were insignificant, high prevalence of Demodex spp. in schizophrenia (35.3) in males can be associated with reduced self-care. Demodex infestation might progress heavily in elderly and immunsuppressed people.

It has been reported that Demodex spp. are widespread in patients with AIDS, leukemia, cancer, diabetes, rheumatoid arthritis, and in pregnant women and hemodialysis patients [10]. Parasite concentrations may increase due to suppressed immune system in these disease manifestations [10-13]. There are studies investigating the incidence of Demodex spp. in immunocompetent people. In a study, Ozcelik et al. identified 25.53 % positivity of D. folliculorum in patients with chronic renal failure and suppressed immune system [1]. In an another study, patients who were diagnosed with type 2 diabetes mellitus examined in terms of D. folliculorum 24.6 % of patients were positive. Also, parasite concentration were high in samples taken from cheek area compared to control group [14].

Schizophrenia is a, affecting approximately 1% population and starts before the age of 25. Schizophrenia, a complex disease with multifactorial etiology, is seen in all social classes, distorts the interpersonal and occupational capability and shows a chronic course [15]. Reduced self-care and weak immune functions are the early symptoms of schizophrenia [7,16]. Generally, schizophrenia is characterized with impartment of cell-mediated immune system, atypical lymphocytes in peripheral blood, decrease in the number and the function of lymphocytes, abnormal lymphoproliferative response to mitogens, abnormal cell-mediated and humoral response to neurons, changes in the number and the ratio of T cells [17,18]. Thus, there is a strong relationship with immune system functions and schizophrenia [19,20]. Müller et al showed that the cellular immune parameters were related to the course of the psychopathological symptoms in schizophrenic and, associated with therapeutic outcome of neuroleptic treatment [21].

Down-regulation of endogenous antioxidant and anti-inflammatory mechanisms has been identified in schizophrenia [22]. For instance prolidase activity which is shown correlated with oxidative stress is found to be higher in schizophrenia patients [23]. Furthermore there are evidences that increased serum levels of chemokines, pro-inflammatory cytokines and, monocytosis, raised inflammatory gene expression in monocytes and altered T-cell function in schizophrenic patients [24-27].

Environmental conditions, nutrition, education level, economic status, hygiene conditions and factors such as the person's immune status is directly related to the prevalence of parasitic infections. Demodex species which are the most common types of permanent parasites of humans, leads to the formation of various dermatological problems in lesions and in skin regions that lack well cleansing. In patients with schizophrenia, especially in cases in which the negative symptoms were observed, they may neglect daily care and cleaning because the social withdrawals are at the forefront. Therefore Demodex spp. can be more prevalent in schizophrenia patients due to lack of self-care.

In the present work, the difference was statistically significant for the positivity of Demodex spp. among patients with schizophrenia than those patients were diagnosed with schizophrenia as compared with the control group (p<0.043) and incidence of these parasites in schizophrenic patients were found to be significantly higher. Although in schizophrenia, self-care is almost always being affected, high prevalence of Demodex spp. in schizophrenia patients appears to be associated with loss of functionality and self care rather than disease’s direct effect. This study has some limitations. Firstly, the sample size is small (n=31). Secondly with data being retrieved from only one hospital in Hatay, results may not be generalized for all schizophrenia patients. Thirdly as severity of schizophrenia and functionality are not measured by scales, the correlation between demodex spp presence and severity of schizophrenia could not be evalu-
ated. Despite these limitations this is the first study that evaporates Demodex spp prevalence in schizophrenia patients. To generalize these results, multicentre studies with more schizophrenia patients are needed.

**Conclusion**

Last of all, the weakening of the immune system, frequently impaired social behavior, especially the lack of hygienic self-care can be risk factors for the Demodex spp. infestations in schizophrenia cases. In these patients, especially in skin lesions, Demodex spp. have to be taken into consideration by informing patients about the treat and explaining the protection-control methods and anti-parasitic treatment should be applied in parasite positive patients.

**Competing interests**

The authors declare that they have no competing interests.

**References**


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