INTRODUCTION

Sleep is a vital process controlled at and affecting every level of a biological structure. Healthy maintenance of thermoregulation, endocrine system, metabolic functioning, and homeostatic system, including immune system, can be impaired because of sleep disorders (1).

Sleep deprivation is a method, which has been used in order to comprehend the functions of sleep both in healthy individuals and for the patients of depression with in treatment, for a long time. The objective of our present study is to examine the relation between hormonal values, which are known for being related to the effects of these said changes determined in the mood, dissociation and thought suppression in healthy individuals after one night of sleep deprivation implementation.

Methods: One night sleep deprivation was performed on a total of thirty-two healthy volunteers (16 males and 16 females) who were included in the study. Blood samples were taken from the individuals before and after sleep deprivation implementation in order to determine cortisol, dehydroepiandrosterone-sulfate (DHEA-S) and Thyroid Functions' Levels tests. In order to evaluate the effects of the sleep deprivation on moods, “White Bear Suppression Inventory (WBSI)” has been conducted, with an aim of evaluating thought suppression, “Profile of Mood States (POMS)”, “Dissociative Experiences Scale (DES)” with a purpose of realizing any dissociation tendency.

Results: On the individuals who have been implemented for sleep deprivation, a decrease on depression and vigor-activity sub-scales values was detected, and an increase was determined on fatigue sub-scales values of “POMS”. While the values of DES were found to have been statistically increased after sleep deprivation, also a significant decrease was determined on WBSI values. Even if there hasn't been any significant statistical change determined on cortisol levels after sleep deprivation, yet there had been some significant changes detected on Thyroid Stimulating Hormone (TSH), fT3, fT4, and DHEA-S levels. Decrease in “POMS” depression sub-scale values and increase on fatigue sub-scale values were determined on the individuals whose sT4 levels were found to be increased significantly in statistic manner after the sleep deprivation.

Conclusion: According to the results of our study, sleep deprivation for one night was determined to cause decrease on depressive mood, increase on dissociative symptoms and to lower the tendency of suppressing the unwanted thoughts, consciously. The fact of being obtained lower depression values, on the individuals with the increased DHEA-S levels after the sleep deprivation meets with the information claiming that the high DHEA-S levels may be deemed as protectors against the negative effects of the stress.

Keywords: Sleep deprivation, dissociation, mood, biochemical changes

The Effects of Sleep Deprivation on Dissociation and Profiles of Mood, and Its Association with Biochemical Changes

Yavuz SELVİ1, Sultan KILIÇ2, Adem AYDIN3, Pınar GÜZEL ÖZDEMİR4

1Department of Psychiatry, Division of Neuro Science, Selçuk University Faculty of Medicine, Konya, Turkey
2Clinic of Psychiatry, Afşin State Hospital, Kahramanmaras, Turkey
3Department of Psychiatry, Necmettin Erbakan University Faculty of Medicine, Konya, Turkey
4Department of Psychiatry, Yüzüncü Yıl University Faculty of Medicine, Van, Turkey

ABSTRACT

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Conclusion: According to the results of our study, sleep deprivation for one night was determined to cause decrease on depressive mood, increase on dissociative symptoms and to lower the tendency of suppressing the unwanted thoughts, consciously. The fact of being obtained lower depression values, on the individuals with the increased DHEA-S levels after the sleep deprivation meets with the information claiming that the high DHEA-S levels may be deemed as protectors against the negative effects of the stress.

Keywords: Sleep deprivation, dissociation, mood, biochemical changes
mer elements when new elements are presented. In addition, reasoning and concentration abilities also decrease because of mood changes (7,9).

The hypothalamic-pituitary-adrenal axis (HPA) acts as a regulator of a behavioral and autonomic response to stress. Recent studies show that sleep deprivation acts as a stressor and leads to a moderate activation in the HPA axis and an increase in plasma glucocorticoid levels (10). Dehydroepiandrosterone-sulfate (DHEA-S), which is another hormone regulated by circadian rhythm, has significant anti-glucocorticoid effects through the inhibition of enzyme activities stimulated by glucocorticoids, and it is asserted that it can prevent destructive effects of hypercortisolemia in case of acute stress (11). Dehydroepiandrosterone (DHEA) and DHEA-S have neuroprotective effects; they increase neuronal plasticity and excitability. There is an interrelation between sleep, energy metabolism and thyroid functions. Thyroid stimulating hormone (TSH) release has a significant pulsatile profile. Sleep causes a fracture in the HPA axis, and sleep deprivation leads to a powerful increase in TSH and related thyroid hormones, with an increase in thyroid releasing hormones (TRH) (12). On the other hand, the interrelation between sleep deprivation, mood states and hormonal changes are not sufficiently clear. In our modern world today, healthy individuals have a limited sleep cycle, which differs from their natural biological rhythms, because of social activities. Besides, the shift system also causes sleep deprivation and it is described as chronic sleep deprivation (13).

This study is intended to discuss the relation between changes in mood states, dissociation and thinking processes that may appear in healthy individuals following one night sleeplessness and hormonal changes that have a circadian rhythm and are strongly.

METHODS

Participants
Thirty-two voluntary healthy university students consisting of 16 males and 16 females have been included in this study. The inclusion criteria for the study have been defined as follows: age >18 years, having no physical or psychiatric disorder; receiving no psychiatric diagnosis of Axis I or Axis II before the study period, using no drugs, alcohol, or substances that are known to affect sleep within the last 15 days and during the study period, and not working in a shift system.

Blood samples have been taken from the individuals meeting the study participation criteria for cortisol, DHEA-S, and thyroid function tests at 08:00 in the morning before and after sleep deprivation. The Dissociative Experiences Scale (DES), the Profile of Mood States (POMS), and the White Bear Suppression Inventory (WBSI), all of which are self-report measures, have been applied.

Measures

Dissociative experiences scale (DES): It is a self-report measure consisting of 28 questions used to screen dissociative experiences and measure their severity. Each item is scored between 0 and 100. The average score is obtained by dividing the sum of these scores by 28. Individuals whose overall score is equal to or above 30 are likely to have a dissociative disorder; the validity and reliability studies have been conducted by Yargıcı et al. (14).

White bear suppression inventory (WBSI): It is a self-report measure developed by Wegner and Zanakos and intended to evaluate the tendency to consciously suppress unwanted thoughts. It is also used in other anxiety disorders, particularly in obsessive compulsive disorder (OCD), and psychiatric disorders such as depression. It totally consists of 15 items. Each item is scored between 1 and 5 depending on the answer. The total score of answers to all the items gives the total measure score. The total score can vary from 15 to 75. High scores indicate that there is a stronger tendency or potential to consciously suppress unwanted thoughts. The validity and reliability studies in Turkey have been conducted by Ağırgün et al. (15).

Profile of mood states (POMS): It has been developed to quickly and reliably define and evaluate situational and short-term changes in mood states (16). It is used to evaluate the effects of psychotherapies and medications, measure the efficiency of sleep deprivation and other similar applications, and survey mood state changes in experimental and clinical studies. As the result of random ordering of the questions forming the measure, certain questions are categorized under 6 individual mood states. These are as follows: “tension-anxiety,” “depression-dejection,” “anger-hostility,” “vigor-activity,” “fatigue-inertia,” and “confusion-bewilderment.” The total POMS score is obtained by deducting the subscale score of “vigor-activity” from the sum of the other five subscale scores. If these five subscale and total scale scores are high, it indicates a higher disorder in mood states. The validity and reliability studies for the scale have been published by Selvi et al. (17) in Turkey.

Procedure
Psychiatric disorder in the individuals whose approvals have been taken and who have met the study participation criteria have been ruled out through the Structured Clinical Interview for DSM Disorders (SCID) evaluation. It was planned to put 32 individuals consisting of 16 males and 16 females through sleep deprivation. The individuals have been accepted to a clinic in 3-people groups each week for the application of total sleep deprivation. Individuals put through total sleep deprivation have been deprived of sleep from 22:00 at night to 07:00 in the morning and were continuously observed by the doctor in charge and the allied health personnel to prevent them from sleeping.

Before the sleep deprivation application, blood samples have been taken from all the subjects at 08:00 in the morning for cortisol, DHEA-S, and Thyroid Function Tests (TFT); POMS, DES, and WBSI scales have been applied. Blood sampling at 08:00 in the morning and scale applications have been repeated following sleep deprivation. Blood samples have been centrifuged at 5000 rpm for 10 min in the biochemistry laboratory to separate serum. Immunoassay kits compatible with fT3, fT4, and TSH Architect 2000 and Immulite 2000 for cortisol and DHEA-S and proper immunoassay kits have been used.

Statistical Analysis
Because all the individuals have fully met the study participation criteria, all of them have been put through statistical evaluation. Age, sex, marital status, all POMS values and sum of POMS subscale scores (six subscales), total POMS value, DES, WBSI, and cortisol, DHEA-S, fT3, fT4, and TSH values before and after sleep deprivation have been entered into the statistical package in SPSS 16.0. Pearson correlation coefficient has been used to analyze the correlation of POMS, DES, WBSI, and cortisol, DHEA-S, fT3, fT4, and TSH values before and after sleep deprivation. T-test has been applied to dependent groups for group comparison purposes. p<0.05 has been considered statistically significant.

RESULTS
The average age of the individuals was 24 years for males and 27 years for females. When POMS subscales were evaluated before and after sleep deprivation, it has been observed that depression scores show a significant
decrease after sleep deprivation, whereas fatigue scores increase significantly and vigor-activity scores decrease significantly (p<0.05). It has been determined that changes in tension-anxiety, anger-hostility, and confusion subscales and overall POMS scores after sleep deprivation are not statistically significant. A significant decrease has been observed in DES and WBSI scores after sleep deprivation (p<0.05) (Table 1).

The evaluation of biochemical parameters following sleep deprivation showed a significant increase in TSH, fT4, fT3, and DHEA-S levels following sleep deprivation (p<0.05). No statistically significant relation has been found between POMS subscales of “tension-anxiety,” “anger-hostility,” and “confusion-bewilderment,” overall POMS scores, cortisol levels, and DHEA-S/cortisol rates before and after sleep deprivation (Table 1).

Correlations of POMS subscales, overall POMS scores, and DES and WBSI scales have been evaluated with biochemical parameters. TSH indicated a positive correlation with “tension-anxiety,” “anger-hostility,” “fatigue-inertia,” and “confusion-bewilderment” subscales and overall POMS score and a negative correlation with “depression-dejection” and “vigor-activity” scores; however, these results were not statistically significant.

Free T4 (T4) indicated a positive correlation with “anger-hostility,” “confusion-bewilderment,” “fatigue-inertia,” and “vigor-activity” subscales and DES scores and a negative correlation with “depression-dejection,” “tension-anxiety,” and WBSI. Of these correlations, the correlations with “depression-dejection” and “fatigue-inertia” subscales were statistically significant.

Free T3 (T3) indicated a positive correlation with “depression-dejection,” “tension-anxiety,” “anger-hostility,” “fatigue-inertia,” and “confusion-bewilderment” subscales and overall POMS score and a negative correlation with “vigor-activity,” DES, and WBSI, although it was not statistically significant (Table 2).

Cortisol indicated a positive correlation with the “confusion-bewilderment” subscale only. This correlation was not statistically significant (Table 2).

DHEA-S had a positive correlation with “confusion-bewilderment,” “anger-hostility,” “fatigue-inertia,” DES, and WBSI and a negative correlation with “depression-dejection,” “tension-anxiety,” and “vigor-activity” subscale scores and overall POMS score. Of these, the correlations with “depression-dejection,” “fatigue-inertia,” and DES were significantly meaningful (p<0.001) (Table 2).

In addition, correlations of changes in POMS subscales following sleep deprivation have been identified. “Tension-anxiety,” “depression-dejection,” and “anger-hostility” subscales showed a positive correlation, whereas the “vigor-activity” subscale showed a negative correlation. The “Depression-dejection” subscale had a positive correlation with “tension-anxiety,” “anger-hostility,” and “fatigue-inertia” subscales and a negative correlation with “vigor-activity.” The “Anger-hostility” subscale showed a positive correlation with “tension-anxiety” and “depression-dejection” subscales in addition to the “fatigue-inertia” subscale and a negative correlation with the “vigor-activity” subscale. A negative correlation has been found between “fatigue-inertia” and “vigor-activity” subscales. Evaluation of the correlation of biochemical parameters with each other following sleep deprivation indicated a positive correlation between changes in TSH levels that appeared following sleep deprivation and changes in T3 levels and these discrepancies were statistically significant (p<0.001). The correlation between T3 values and DHEA-S values was positive and statistically significant (p<0.05). There was a positive correlation between cortisol and other blood parameters, which was not statistically significant.

**DISCUSSION**

In this study, 32 healthy individuals were put through total sleep deprivation to study the effect of sleep deprivation on mood states, dissociation, and thought suppression as well as changes in cortisol and DHEA-S levels and thyroid functions that are known to affect mood states and reveal circadian release.

The result of our study showed a significant decrease in the subscale predicting depressive complaints in healthy individuals following sleep deprivation. In addition, sleep restriction led to a decrease in vigor and activity levels while increasing fatigue levels. It caused an insignificant decrease in the “confusion-bewilderment” subscale, thereby indicating confusion.

The sleep deprivation application provides a controlled regulation in the sleep-wake rhythm. In addition to preclinical and in vivo studies, clinical studies also show that sleep deprivation increases serotonin, norepinephrine, and dopamine neurotransmission, increases thyroid hormone levels, and affects targets in treatment of mood state disorders such as glutamate (18,19,20). The results indicating an improvement in the depressive mood of healthy individuals following sleep deprivation showed that, individual psychobiological differences in endogenous characteristics, diurnal characteristics, and sleep-wake pattern can be the predictors for the response to the treatment (4).

<table>
<thead>
<tr>
<th>Table 1. Changes in mood, dissociation, thought suppression and biochemical parameters of healthy individuals before and after total sleep deprivation according to the POMS, DES and WBSI evaluations</th>
</tr>
</thead>
<tbody>
<tr>
<td><strong>Before sleep deprivation</strong></td>
</tr>
<tr>
<td><strong>Mean</strong></td>
</tr>
<tr>
<td>Depression</td>
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<tr>
<td>Anxiety</td>
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<tr>
<td>Anger</td>
</tr>
<tr>
<td>Confusion</td>
</tr>
<tr>
<td>Fatigue</td>
</tr>
<tr>
<td>Vigor</td>
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<tr>
<td>POMS total</td>
</tr>
<tr>
<td>DES</td>
</tr>
<tr>
<td>WBSI</td>
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<tr>
<td>TSH</td>
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<tr>
<td>T4</td>
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<tr>
<td>T3</td>
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<tr>
<td>DHEA-S</td>
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<tr>
<td>Cortisol</td>
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<tr>
<td>Cortisol/ DHEA-S rate</td>
</tr>
</tbody>
</table>

*p<0.05; **p<0.01
SD: standard deviation; POMS: profile of mood states; DES: dissociative experiences scale; WBSI: White Bear Suppression Inventory; TSH: thyroid stimulating hormone; DHEA-S: dehydroepiandrosterone-sulfate
The studies indicate a decrease in activity, vigor, and excitability in individuals subjected to a restricted sleep time during consecutive days and an increase in fatigue, sleepiness, anxiety, confusion, bewilderment, and irritability (21), whereas in our one night sleep deprivation study, it has been determined that there is no significant change in tension-anxiety, anger, and confusion. This indicates that cumulative effects of chronic partial sleep deprivation may be different from those of acute deprivation and is important to foresee mood state changes that may occur in depressive patients during treatment.

In this study, a significant decrease following sleep deprivation has been determined in total scores from the WBSI scale applied to evaluate the conscious suppression of thoughts or imaginations that individuals do not want to have and perceive as ego-dystonic. These low scores obtained provide the impression that the tendency or the potential to consciously suppress unwanted thoughts decrease following sleep deprivation. Even though such a decrease looks like a positive finding, it may also indicate the effect of a decreased cognitive function along with increased sleepiness.

A significant increase has been determined in dissociation levels following sleep deprivation in our study. This result conforms with the information obtained from the recent research, thus asserting that deviant sleep experiences are associated with dissociative symptoms.

The research has been focused on subjective experiences involving de-realization, absorption, and amnesic complaints, which are dissociative symptoms and abnormal experiences related to sleep (22,23). It has been observed that sleep loss increases dissociative symptoms in long-term shift workers and in groups with a regularly restricted sleep (24). Impairment in the sleep-wake cycle leads to cognitive errors and is also associated with the susceptibility to daydream about imaginary occupations. It also increases hypnotizability and suggestibility (25,26). In our study, both the increase in dissociation symptoms following sleep deprivation and the decrease in the ability to consciously suppress thoughts indicate that cognitive function disorders caused by changes in sleep-wake cycle can be caused through dissociation or deteriorations in both areas. This is supported by the fact that individuals with high dissociation scores have higher memory neglect errors; this cannot be explained with decreased working memory capacity and high emotional reactivity, and neglect errors are typical in individuals with high dissociation scores (24,26,27).

In our study, a significant increase in dissociation levels following sleep deprivation can be considered as the result of intrusion of electrophysiological characteristics of sleep into wake consciousness. In addition, because norepinephrine, which is a neurotransmitter regulating excitement and alertness, is associated with dissociative symptoms and people prone to dissociation have cholinergic system hypersensitivity, it can be the biological explanation to increased dissociation levels following sleep deprivation (28,29). Acetylcholine, which is released at the highest level in the cortex during wake and Rapid Eye Movement (REM) periods, and neurons localized at peri-brachial area of pons are responsible for cortical desynchronization, rapid eye movements, and muscle paralysis, which are REM components. Because cognitive and behavioral disorders occur due to on one hand, increased serotonergic and noradrenergic activity following sleep deprivation due to mutual interaction between cholinergic and amineergic neurons, resulting in an improvement in mood states, and on the other hand, cholinergic activity is the regulator of conditions associated with cognitive functions such as consciousness, memory, and attention and is related to motor behaviors during sleep, this can shed light on the underlying neurobiological mechanism. In particular, following sleep deprivation causing dysregulation of neurotransmitters such as acetylcholine, serotonin, and norepinephrine, duration of sleep stages that will also occur electrophysiologically and intrusion of failures in transition between such stages and sleep-wake into consciousness may have contributed to Table 2. Correlation between changes in mood, dissociation, thought suppression and hormonal changes occurring in healthy individuals following total sleep deprivation

<table>
<thead>
<tr>
<th></th>
<th>Depression</th>
<th>Anxiety</th>
<th>Rage</th>
<th>Confusion</th>
<th>Prostration</th>
<th>Vigor</th>
<th>Mdp Total</th>
<th>DES</th>
<th>Sea bear</th>
<th>TSH</th>
<th>sT4</th>
<th>sT3</th>
<th>DHEA-S</th>
<th>Cortisol</th>
<th>DHEA-S/Cortisol</th>
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</thead>
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<td>0.24</td>
<td>0.19</td>
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<td>-0.12</td>
<td>0.51***</td>
<td>-0.49**</td>
<td>-0.08</td>
<td>-0.04</td>
<td>-0.40</td>
<td>0.26</td>
<td>-0.61**</td>
<td>-0.18</td>
<td>-0.04</td>
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<td>0.11</td>
<td>0.24</td>
<td>-0.37</td>
<td>0.67**</td>
<td>0.07</td>
<td>0.20</td>
<td>0.13</td>
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<td>0.21</td>
<td>-0.12</td>
<td>-0.05</td>
<td>-0.04</td>
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<td>Rage</td>
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<td>0.13</td>
<td>0.66**</td>
<td>-0.55**</td>
<td>0.84**</td>
<td>0.21</td>
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<td>0.11</td>
<td>0.26</td>
<td>0.25</td>
<td>0.14</td>
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<td>0.84**</td>
<td>0.36**</td>
<td>0.64**</td>
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<tr>
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<td>0.02</td>
<td>-0.32</td>
<td>0.11</td>
<td>0.13</td>
<td>-0.09</td>
<td>0.41</td>
<td>0.11</td>
<td>-0.42</td>
<td>-0.01</td>
<td>-0.16</td>
<td>-0.42</td>
<td>0.35</td>
<td>1</td>
</tr>
</tbody>
</table>

DHEA-S: dehydroepiandrosterone-sulfate; DES: dissociative experiences scale; POMS: profile of mood states; WBSI: White Bear Suppression Inventory; TSH: thyroid stimulating hormone.
the occurrence of confusion, attention deficit, and dissociative symptoms. Therefore, this study which has a clinical aspect must be supported with neurobiological and electrophysiological studies.

In this study, it has been found that sleep deprivation does not lead to a significant change in cortisol levels of healthy individuals. Conflicting results obtained from former studies also show that a definite result cannot be provided (30). On the other hand, it has been shown that sleep deprivation is perceived as a stressor and thus results in a slight activation in the HPA axis and an increase in plasma concentrations of glucocorticoids (10). In addition, a significant increase has been observed in DHEA-S levels following sleep deprivation. It is known that DHEA-S levels increase due to acute stress in healthy individuals. Preclinical and clinical studies provide indirect evidences that DHEA-S levels and DHEA-S/cortisol ratio can have an important role in the regulation of the effect of stress (31). Clinical studies show that peritraumatic symptoms of dissociation is a risk factor for the development of post-traumatic stress disorder, and these symptoms are positively associated with glucocorticoid release induced by stress (32). It is thought that increased DHEA-S/cortisol ratio can be protective against the negative effects of stress (31). It has been determined that DHEA-S/cortisol ratios during stress are significantly higher in individuals with lesser dissociative symptoms and higher performance (33).

Our study has shown that DHEA-S levels have a positive correlation with DES scores and a negative correlation with depression scores. It has been concluded that lower depression scores in individuals with increased DHEA-S levels following sleep deprivation can be because of the high DHEA-S’s protective effect against the negative effect of stress and the improvement in cognition.

In this study, significant increases have been observed in TSH, fT3, and fT4 levels following sleep deprivation. Certain studies state that thyroid function can be predictive in response to sleep deprivation and show that there is an increase in TSH following sleep deprivation (34). Because it shows that thyroxin (T4) safely elongates clinical response occurring because of sleep deprivation, it strongly supports the relation of antidepressant effect and changes in thyroid functions (35). Similarly, our study is intended to define the biological indicators serving to improve depressive mood following sleep deprivation and the increase in T4, levels seems to provide this result. As stated before, the increase in T4 levels following sleep deprivation in our study has been found to be correlated with the decrease in subscales predicting depressive mood. If this result is interpreted correctly, it can guide to determinants studied for the treatment of sleep deprivation in the former studies.

Because individuals have not been evaluated for their morning and evening chronotypes, it is the main limitation of this study. Besides, it constitutes a preliminary study for research that can evaluate the effects of sleep deprivation in neurobiological terms and new research is required to be conducted in this respect.

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