

## Brain-Behavioral Systems in Patients with Comorbid Anxiety -Depression vs. Healthy Individuals

Saeedeh Azaraeen, M.A.<sup>1</sup>, Roshanak Khodabakhsh, Ph.D.<sup>2</sup>, Zohreh Khosravi, Ph.D.<sup>3</sup>

1- PhD Student of Psychology, Department of Psychology, AL Zahra University, Tehran, Iran (Corresponding author; saide.azaraeen@yahoo.com)

2- Associate Professor of Psychology, Department of Psychology, AL Zahra University, Tehran, Iran

3- Professor of Psychology, Department of Psychology, AL Zahra University, Tehran, Iran

Received: 22 July, 2017

Accepted: 21 November, 2018

### ARTICLE INFO

#### Article type:

Original Article

#### Keywords:

Brain -Behavioral Systems

Anxiety

Depression

Comorbidity

### Abstract

**Background:** According to the brain-behavioral systems theory, behavioral inhibition and behavioral activation systems contribute to the development of many psychopathological conditions. Given that anxiety and depression are the most common emotional disorders and the fact that they are highly overlapping, the aim of this study was to compare the brain-behavioral systems in the patients with comorbid anxiety-depression and healthy individuals.

**Method:** This study was cross-sectional. Sample includes Sixty-four patients with comorbid anxiety and depression attending to the community health centers of Jiroft city and 64 healthy individuals. They were selected using purposive sampling and matched for age and gender. After obtaining informed consent, they were asked to fill Jackson-5 scale, Beck Depression Inventory and Beck Anxiety Inventory. Data are analyzed by MANOVA.

**Results:** The results showed that the two groups had significant differences in behavior inhibition system, fight, flight and freeze. However, there was no difference in behavioral activation system.

**Conclusion:** Given the observed differences in the brain-behavioral systems between patients with comorbid anxiety-depression and healthy individuals, this theory may explain the comorbidity of anxiety and depression.

**Copyright:** 2018 The Author(s); Published by Kerman University of Medical Sciences. This is an open-access article distributed under the terms of the Creative Commons Attribution License (<http://creativecommons.org/licenses/by/4.0>), which permits unrestricted use, distribution, and reproduction in any medium, provided the original work is properly cited.

**Citation:** Azaraeen S, Khodabakhsh R, Khosravi Z. Brain-Behavioral Systems in Patients with Comorbid Anxiety -Depression vs. Healthy Individuals. Journal of Kerman University of Medical Sciences, 2018; 25 (4): 328-338.

### Introduction

A large number of studies documented depression and anxiety as highly comorbid disorders. It is reported that 75 percent of those afflicted with depression, also experience anxiety symptoms (1-3). Comorbidity of emotional disorders is linked with the most severe periods of the psychological disease and a weaker response to treatment (4). Several

models have been suggested to bring us to the better understanding of phenomenological comorbidity of depression and anxiety (5-9); Yet, Gray's reinforcement sensitivity theory (10) gained less attention in the comorbidity of depression and anxiety.

Inspired by Eysenck's theory, Gray (10) developed the Reinforcement Sensitivity Theory and based it on

three brain-behavioral systems: Behavioral Activation System (BAS), Behavioral Inhibition System (BIS), and Fight Flight System (FFS). He believed these brain-behavioral systems underlie the individual differences so that, their activations would arouse different emotional responses such as fear and anxiety.

BAS is the first system in the Gray model which responds to the conditioned stimuli of reward and non-punishment conditions. Activated and its sensitivity heightened, this system arouses positive emotions and active avoidance of punishment (11). BAS sensitivity is associated with the increase of positive emotions and the impulsive dimensions of the personality (12). The second system, BIS, is responsive to conditioned stimuli of punishment, non-reward, novel stimuli and also to innate fear stimuli. Activated, this system arouses anxiety, behavioral inhibition, passive avoidance and extinction and also increases attention and arousal. FFFS, which is the third system in Gray's model, is associated with activations of amygdale and hypothalamus and is sensitive to aversive stimuli. Studies on the role of these systems in clinical psychological disorders showed that the excessive activity of BAS and BIS makes people prone to mental disorders (13).

The results of some studies demonstrated the association between these three systems and a number of mental disorders; FFFS is proved to be linked with phobia and panic disorder (14). BAS is associated with addictive behaviors, social-emotional and psychological adaptation (15), bipolar disorder (16, 17), and attention-deficit and hyperactivity disorder (18, 19). BIS is linked with generalized anxiety disorder (20), obsessive

compulsive disorder (14), and also major depressive disorder [MDD] (17, 21).

MDD may occur due to the extremely insensitive BAS that fails to arouse positive emotions or to respond to encouraging environmental stimuli (7). Depressive individuals with lower levels of BAS are more likely to fail to respond to positive incidents and stimuli in their environment. They are less likely to search for positive stimuli and are less engaged in pleasure-giving activities (22). Indeed, low sensitive BAS seems to be common with patients who are suffering from unipolar depression (17). Moreover, BIS that has been confirmed to be associated with anxiety, has revealed strong link with unipolar depression (23). Though BAS low sensitivity is associated with depression, BIS high sensitivity is reported to be linked with a wide range of emotional problems (13).

Johnson, et al. (24) found out that high BIS score is a predictive factor for lifelong anxiety and depression. Examining the relation between brain-behavioral systems and anxiety, Ly and Gomez (25) found out that anxiety is positively related to BIS and punishment sensitivity, but it is negatively related to BAS. In the study of Vervoort et al. (26), the anxious group scores in BIS were higher than nonanxious group. In Kimberl, et al. study (27), higher BIS scores predicted anxiety and depression.

Findings that show relation of BIS high sensitivity to anxiety and depression (21, 28, 29) may explain depression-anxiety comorbidity. Yet, there is relatively little literature supporting this view. High BIS and low BAS may also provide a potential ground for the comorbid relation between anxiety and depression. Besides these, BIS is likely to be a common risk factor

for both disorders (30). In general, the study of relationship between personality theories and mental disorders can increase our understanding of the etiology and comorbidity of these disorders (31, 32) and suggest appropriate ways for preventing and treating them (33).

In addition to the corroborated findings, Gray's reinforcement sensitivity theory brings us to the understanding that irregularity in brain-behavioral systems is one of the factors leading to comorbidity of anxiety and depression. There has been relatively little research conducted on the activity of brain-behavioral systems in those patients suffering from both the anxiety and depression in other countries and there is no study about this topic in Iran. Therefore, the current study aimed to discern whether the normal cases and the patients with comorbid anxiety-depression differ in their brain-behavioral systems.

## Method

In this cross-sectional study, the brain-behavioral systems of normal individuals and patients afflicted with anxiety-depression comorbidity were investigated. The studied population consisted of patients with comorbid anxiety-depression referring to governmental and non-governmental health care centers in Jiroft city. They were selected through purposive sampling method and scored average to high according to Beck Depression Inventory-II [BDI-II] and Beck Anxiety Inventory [BAI]. The diagnostic results were also confirmed in consultation with a psychiatrist and a clinical psychologist. In addition, the normal cases consisted of people referring to health care centers to receive due services. Selected purposively due to their low score in BDI-II and BAI, they were matched to the

patient participants based on the age and gender variables. The sample size was estimated by G\*Power (34). Split into two groups, the sample yielded an effect size of  $f^2=.25$ , a relatively medium effect size, a power of .80 and 64 subjects per group. The results were then analyzed using SPSS version 21 by MANOVA.

## Study tools

Beck Depression Inventory-II: This inventory (35) is a revised version of BDI-II that was designed and validated to measure the severity of depression in Iranian samples (36). It is designed for individuals aged 13 and over and its scoring is based on Likert scale. The cut-off points are 0–13 that indicates minimal depression or no depression at all, 14–19 that indicates mild depression, 20–28 that indicates moderate depression, and 29–63 that indicates severe depression (37, 38). Studies on the second version of this inventory have reported that it has desirable validity, reliability and factor structure and also it is a proper substitute for the first version (36). An Iranian study, reported a strong alpha coefficient of 0.91, a 0.89 correlation coefficient between the two halves, a test-retest (1-week interval) coefficient of 0.94 and 0.93 correlations with BDI-I in an Iranian sample (36). In the current study, Cronbach's alpha was 0.92.

Beck Anxiety Inventory (BAI): Given the importance of the main dimensions of anxiety- that are the cognitive and physiological symptoms- Beck, et al. (39) designed a 21-questions self-report inventory with Likert scale. BAI scores the frequency of the subjects' anxiety symptoms on a scale value of 0 to 3 during the past week. The suggested cutoff points are 0–7 for minimal anxiety, 8–15 for mild anxiety, 16–25 for moderate anxiety, and 26–63 for severe anxiety. In

studying the internal consistency of the questionnaire in Iranian society, alpha coefficient was 0.92, there was a 0.91 reliability coefficient between the two halves, a test-retest (1-week interval) coefficient of 0.81 and a correlation of 0.62 with BDI-II (36). Cronbach's alpha for this study was 0.92.

Jackson Five-Scale Inventory: Jackson (40) developed a 30-question inventory to properly measure revised-Reinforcement Sensitivity Theory. The inventory consists of five subscales: BAS, BIS, and Fight, Flight and Freeze system (FFFS). Six questions have been designed for each subscale. Using confirmatory and exploratory factor analysis, Jackson developed and assessed new scales (Jackson Five-Scale). The results showed a satisfactory internal reliability and a desirable validity of the construct. The participant answered a 5-item Likert scale with 1= strongly agreed (Always) and 5= strongly disagreed (Never). Using double-translation method, Hasani, et al. (41) employed the Farsi version of Jackson Five-Scale Inventory for 308 participants (174 males & 134 females). The reliability of the inventory was examined by internal consistency, item-total correlations, and test-retest methods. Moreover, the scale's validity was examined by exploratory factor analysis, subscales' inter-correlations, and criterion validity. Cronbach's alpha value range (0.72 to 0.88), test-retest coefficients (0.64 to 0.78), and, item-total correlations (0.28 to 0.68) showed that the Farsi version of Jackson Five-factor Inventory is of a desirable validity. Confirmatory and exploratory factor analysis also supported the five main factors of the inventory. Subscales' internal consistency (ranged from 0.11 to 0.53) was reported as desirable, too. Finally, the scale's validity was reported

satisfactory since a specific pattern of correlation coefficient was recognized between inventory subscales from one hand and negative emotion, positive emotion, BAS and BIS, Eysenck's personality dimensions, and Bart's impulsivity dimensions on the other hand. Cronbach's alpha calculated for the subscales were: BAS: 0.52, BIS: 0.62, Fight: 0.75, Flight: 0.68 and Freeze: 0.63.

### Procedure

The study's inclusion criteria consisted of literacy for answering the inventory, being in the age range of 18-55 years and willingness to participate in the study. Exclusion criteria included lack of literacy, psychiatric disorders history except for anxiety and depression, bipolar mood disorder, mental retardation, physical diseases or any other conditions attributable to depression. These disorders assessed in the two groups through clinical interviews before the questionnaires were completed. Qualified participants were thus selected and asked to complete the questionnaires. Participants scored high ( $\leq 20$ ) and low (0-12) in BDI-II and high ( $\leq 16$ ) and low (0-7) in BAI were allocated to the patients and normal groups, respectively and individually completed the questionnaires. Participants were then thanked.

### Results

The demographic information of participants has been summarized in table 1.

Independent samples t-tests showed no significant age differences in the two groups,  $t(126) = 1.02$ ,  $p = 0.30$ . In addition, comparing two groups using chi-square test showed no significant differences in terms of gender ( $p =$

0.34), marital status ( $p= 0.84$ ), education ( $p= 0.06$ ), and addiction history ( $p= 0.09$ ). However, significant group differences are observed in terms of anxiety and depression history ( $p=0.001$ ) and psychiatric drugs use ( $p=.001$ ). The groups differed significantly only in psychiatric drug use and anxiety and on depression

history whereby those in the patient group reported more history of anxiety, depression disorder and psychiatric drug use.

Means, SDs, skewness, S.E. of skewness, kurtosis and S.E of kurtosis of the variables have been shown in Table 2.

**Table 1.** Demographic characteristics of the studied groups

Descriptive Variable		Normal group (n=64)		Patient group (n=64)	
		frequency	percent	frequency	percent
<b>Gender</b>	Female	41	64.1	46	71.9
	Male	23	35.9	18	28.1
<b>Marital Status</b>	Single	17	26.6	18	28.1
	Married	47	73.4	46	71.9
<b>Education</b>	Elementary	12	18.8	3	4.7
	Intermediate	10	15.6	11	17.2
	High school Diploma	23	35.9	34	53.1
	Bachelor	18	28.1	16	25
	Master	1	1.6	0	0
<b>Addiction History</b>	Yes	1	1.6	5	7.8
	No	63	98.4	59	92.2
<b>Anxiety and depression History</b>	Yes	5	7.8	41	64.1
	No	59	92.2	23	35.9
<b>Psychiatric Drugs Use</b>	Yes	2	3.1	23	35.9
	No	62	96.9	41	64.1

**Table 2.** Means, SDs and the other distribution statistics of the studied variables

Group	Variable	Age	Depression	Anxiety	BAS	BIS	Fight	Flight	Freeze
<b>Normal</b>	Mean	32.33	4.84	3.52	19.52	20.47	14.05	16.66	15.67
	SD	9.26	4.09	2.98	4.54	6.22	4.85	4.75	3.90
	Skewness	.431	.477	.953	-.645	-.535	.489	-.194	-.001
	S.E. Skewness	.299	.299	.299	.299	.299	.299	.299	.299
	Kurtosis	-.471	-.899	1.891	1.130	-.344	.184	-.354	-.222
	S.E Kurtosis	.590	.590	.590	.590	.590	.590	.590	.590
<b>Patients</b>	Mean	30.86	30.23	29.45	19.94	22.77	18.59	19.38	19.77
	SD	6.76	8.26	9.39	4.10	4.41	5.44	4.82	4.80
	skewness	.538	.728	.804	-.301	-.621	-.135	-.722	-.142
	S.E. skewness	.299	.299	.299	.299	.299	.299	.299	.299
	kurtosis	.356	.210	1.283	-.422	-.190	-.324	.111	-.484
	S.E kurtosis	.590	.590	.590	.590	.590	.590	.590	.590

MANOVA analyses confirmed that there was a significant multivariate effect: Pillai's Trace= .265,  $F_{(5,122)}= 8.81$ ,  $p=0.001$ , when compared with the normal group, participants in the patient group were significantly higher in the mean scores of BIS,  $F_{(1,126)}= 5.79$ ,  $p= 0.018$ , Fight,  $F_{(1,126)}=24.88$ ,  $p= 0.001$ , Flight,  $F_{(1,126)}=10.30$ ,  $p= 0.002$ , and Freeze,  $F_{(1,126)}= 27.97$ ,  $p= 0.001$ . There was no significant difference in BAS,  $F_{(1,126)}= .30$ ,  $p=.58$ .

## Discussion and Conclusion

The current study aimed to compare brain-behavioral systems in the normal individuals and patients afflicted with comorbid anxiety and depression. Congruent with the previous studies, the results validated the hypothesis that individuals with comorbid anxiety-depression differ in their brain-behavioral systems from healthy cases (23, 24, 42, 43). In the similar way, Gray (44) believed that anxiety and neurotic depression, i.e., comorbid depression-anxiety, is the result of BIS excessive activity. Yusuke, et al. (45) have investigated the personality characteristics and their probable relations to anxiety and depression symptoms. They have reported both anxiety and depression as positively and significantly linked with BIS.

In general, high BIS sensitivity leads to increased attention to threat, increased negative affect and increasing behavioral inhibition which ultimately leads to anxiety (46). In other words, the higher BIS sensitivity increases the response to negative events (47). Hence, this finding suggests that BIS may play a role both in depression and anxiety. In r-RST, BIS is responsible for resolving the conflict, i.e., situations involving both reward and threat (48). This applies to conflicts between competing goals of the FFFS and the BAS (avoiding pain and approaching reward) but can also be between FFFS-FFFS goals or BAS-BAS goals (e.g. making a choice between two potential rewards). According to r-RST, r-BIS operates in either one of two modes. When in “checking” mode, its role is to be a risk-assessor, meaning that it monitors the environment

and scans memory of previous aversive events in order to detect potential danger. When in “control” mode, r-BIS becomes activated and attention to the environment increases. In the case of a FFFS-BAS conflict, this is when it would assess the merits of avoiding versus approaching the stimulus in making a decision about the best response (49). As a decision-making system, the corresponding emotions are feelings of anxiety and worry in the face of unfamiliar stimuli or frustration when faced with the absence of reward. Therefore, people with comorbid depression and anxiety are likely to be more in the conflict situations.

In the last decades, researchers paid attention to Reinforcement Sensitivity Theory (RST) in different disorders. Depression is turned out to be one of the disorders gaining considerable attention in those studies. According to RST, depression is characterized with motivational deficiencies or lack of positive reinforcement and as well by an increase in avoidance behaviors such as social withdrawal. In addition, the mentioned theory associates depression with low BAS and high BIS (50). Quilty and collaborators (17), reported a strong link between depression symptoms and higher sensitivity to punishment (BIS). On the other hand, anxiety disorders are characterized by an increase in the vigilance to threat stimuli and they are linked with higher BIS (23). Moreover, it is believed that higher levels of BIS would arouse more anxiety (45).

The neuroanatomical bases of BIS are located in serotonergic and noradrenergic pathways in orbitofrontal cortex, septo-hippocampal system and the Papez circuit (10, 51). Due to the BIS higher sensitivity

in those patients with comorbid anxiety-depression, and given the fact that these individuals respond well to serotonin reuptake inhibitors (52), the findings of the present study supports Gray's model which assumes that the serotonergic irregularities of septo-hippocampal system form the basis for depression and anxiety. Levita et al. (53) in their study on the BIS, anxiety and hippocampal volume in the nonclinical population found that greater levels of BIS were positively associated with right hippocampal volume.

Pourmohammad-Rezai-Tajrishi and Mirzamani-Bafghi (54) found a significant link between BIS and depression. Mansuri and Bakhshipour-Roudsari (55) showed that BIS is associated with pathological and non-pathological anxiety. Basharpour and Mozafari (56) studied the role of BAS and BIS in predicting students' state-trait anxiety. They reported a link between high BIS and state-trait anxiety. They also showed that the lower BAS is linked only with trait anxiety. Karsazi and Hashemi (57) examined the structural relations of brain-behavioral systems and the difficulty of emotion-regulation to depression and social anxiety symptoms. Their findings revealed that the brain-behavioral systems can act out as personality-neurobiological basis for depression and social anxiety disorder. These findings show that these neurobiological systems are vulnerable to depression and anxiety.

The present study demonstrated that normal individuals and those with comorbid anxiety-depression did not differ significantly in BAS. According to Gray (44), BAS activity differs in anhedonic depression and anxiety-depression comorbidity. Examining for the BAS

and BIS activity in these two subtypes of depression, Kimbrel, et al. (27) found out that low BAS only predicts anhedonic depression; and high BIS is associated with anxiety-depression comorbidity. Thus, anxiety-depression comorbidity might well explain the reason for the finding that the groups in the present study showed no difference in BAS. Likewise, Yusuke, and colleagues (45) found no significant correlation between depression and BAS. Yet, using hierarchical regression to control anxiety symptoms, they found a link between depression and BAS. In their study of brain-behavioral systems and depression/anxiety dimensions, Spielberg, et al. (58) reported anhedonic depression and anxiety dimensions (cognitive and physical arousal) to be linked with high BIS. They also reported that only anhedonic depression is linked with low BAS. Thus, it is likely that high BIS leads to anxiety-depression comorbidity. Yet, more studies needed to be done to uncover the link between BAS and depression subtypes and their due symptoms.

Small sample size and the employment of cross-sectional data gathering method were the limitations of the current study. Yet, the study was an unprecedented one in Iran that contrasted the brain-behavioral systems in normal individuals and those with depression-anxiety comorbidity. So, this theory may explain Iranian patients with comorbidity of anxiety and depression. As we know, human behavior is strongly influenced by cultural differences, and this theory may explain the psychological phenomenon occurred to Iranian patients instead of patients in other cultural contexts.

## References

1. Essau CA. Comorbidity of depressive disorders among adolescents in community and clinical settings. *Psychiatry Research* 2008; 158(1):35-42.
2. Essau CA, editor. *Epidemiology, comorbidity, and course of adolescent depression*. Oxford, UK: Oxford University Press; 2009.
3. Garber J, Weersing VR. Comorbidity of anxiety and depression in youth: implications for treatment and prevention. *Clin Psychol (New York)* 2010; 17(4):293-306.
4. Fichter MM, Quadflieg N, Fischer UC, Kohlboeck G. Twenty-five-year course and outcome in anxiety and depression in the upper Bavarian longitudinal community study. *Acta Psychiatrica Scandinavica* 2010; 122(1):75-85.
5. Watson D, Tellegen A. Toward a consensual structure of mood. *Psychological Bulletin* 1985; 98(2):219-35.
6. Clark LA, Watson D. Tripartite model of anxiety and depression: psychometric evidence and taxonomic implications. *J Abnorm Psychol* 1991; 100(3):316-36.
7. Mineka S, Watson DW, Clark LA. Comorbidity of anxiety and unipolar mood disorders. *Annu Rev Psychol* 1998; 49:377-412.
8. Krueger RF, Chentsova-Dutton YE, Markon KE, Goldberg D, Ormel J. A cross-cultural study of the structure of comorbidity among common psychopathological syndromes in the general health care setting. *Journal of Abnormal Psychology* 2003; 112(3):437-77.
9. Krueger RF, Markon KE. Reinterpreting comorbidity: a model-based approach to understanding and classifying psychopathology. *Annu Rev Clin Psychol* 2006; 2:111-33.
10. Gray JA. Framework for A Taxonomy of Psychiatric Disorder. In: Van Goozen SH, Van de Poll NE, editors. *Emotions: Essays on Emotion Theory*. New York: Psychology Press; 1994.
11. Gray JA, McNaughton N. The neuropsychology of anxiety: Reprise. In: Hope DA. (editor). *Nebraska Symposium on Motivation. Perspectives on anxiety, panic, and fear*. Lincoln: Nebraska University; 1996: 61-134.
12. Corr PJ. Reinforcement sensitivity theory and personality. *Neurosci Biobehav Rev* 2004; 28(3):317-32.
13. Bijttebier P, Beck I, Claes L, Vandereycken W. Gray's reinforcement sensitivity theory as a framework for research on personality-psychopathology associations. *Clin Psychol Rev* 2009; 29(5):421-30.
14. Corr PJ, editor. *The Reinforcement Sensitivity Theory of Personality*. Swansea: University of Wales; 2008.
15. Colder CR, O'Connor RM. Gray's reinforcement sensitivity model and child psychopathology: laboratory and questionnaire assessment of the BAS and BIS. *J Abnorm Child Psychol* 2004; 32(4):435-51.
16. Alloy B, Bender RE, Wanger CA, Whitehouse WG, Abramson LY, Hogan ME, et al. Bipolar spectrum-substance use co-occurrence: behavioral approach system sensitivity and impulsivity as shared personality vulnerability. *J Pers Soc Psychol* 2009; 97(3):549-65.
17. Quilty LC, Mackew L, Bagby RM. Distinct profiles of behavioral inhibition and activation system sensitivity in unipolar vs. bipolar mood disorders. *Psychiatry Res* 2014; 219(1):228-31.



18. Gomez R, Corr PJ. Attention-deficit/hyperactivity disorder symptoms: associations with gray's and Tellegen's models of personality. *Personality and Individual Differences*. 2010; 49(8):902-6.
19. Heym N, Kantini E, Checkley HL, Cassaday HJ. Gray's revised reinforcement sensitivity theory in relation to attention-deficit/hyperactivity and tourette-like behaviors in the general population. *Pers Individ Dif* 2015; 78:24-8.
20. Pawluk EJ, Koerner N. A preliminary investigation of impulsivity in generalized anxiety disorder. *Pers Individ Dif* 2013; 54(6):732-7.
21. Pinto-Meza A, Caseras X, Soler J, Puigdemont D, Pèrez V, Torrubia R. Behavioral inhibition and behavioral activation systems in current and recovered major depression participants. *Pers Individ Dif* 2006; 40(2):215-26.
22. Kasch KL, Rottenberg J, Arnow BA, Gotlib IH. Behavioral activation and inhibition systems and the severity and course of depression. *J Abnorm Psychol* 2002; 111(4):589-97.
23. Hundt NE, Nelson-Gray RO, Kimbrel NA, Mitchell JT, Kwapil TR. The interaction of reinforcement sensitivity and life events in the prediction of anhedonic depression and anxiety symptoms. *Pers Individ Dif* 2007; 43(5):1001-12.
24. Johnson SL, Turner RJ, Iwata N. BIS/BAS levels and psychiatric disorder: an epidemiology study. *J Psychopathol Behav Assess* 2003; 25(1):25-36.
25. Ly C, Gomez R. Unique associations of reinforcement sensitivity theory dimensions with social interaction anxiety and social observation anxiety. *Pers Individ Dif* 2014; 60:20-4.
26. Vervoort L, Wolters LH, Hogendoorn SM, de Haan E, Boer F, Prins PJM. Sensitivity of gray's behavioral inhibition system in clinically anxious and non-anxious children and adolescents. *Personality and Individual Differences* 2010; 48(5):629-33.
27. Kimbrel NA, Nelson-Gray RO, Mitchell JT. Reinforcement sensitivity and maternal style as predictors of psychopathology. *Pers Individ Dif* 2007; 42(6):1139-49.
28. Muris P, Meesters C, de Kanter E, Timmerman PE. Behavioural inhibition and behavioural activation system scales for children: Relationships with Eysenck's personality traits and psychopathological symptoms. *Pers Individ Dif* 2005; 38(4):831-41.
29. Zinbarg RE, Yoon KL. RST and Clinical Disorders: Anxiety and Depression. In: Corr, PJ, editor. *The Reinforcement Sensitivity Theory of Personality*. New York: Cambridge University Press; 2008.
30. Schofield CA, Coles ME, Gibb BE. Retrospective reports of behavioral inhibition and young adults' current symptoms of social anxiety, depression, and anxious arousal. *J Anxiety Disord* 2009; 23(7):884-90.
31. Brown TA. Temporal course and structural relationships among dimensions of temperament and DSM-IV anxiety and mood disorder constructs. *J Abnorm Psychol* 2007; 116(2):313-28.
32. Lahey BB. Public health significance of neuroticism. *Am Psychol* 2009; 64(4):241-56.
33. Kovacs M, Lopez-Duran N. Prodromal symptoms and atypical affectivity as predictors of major depression in juveniles: implication for prevention. *Journal of child psychology and psychiatry and allied disciplines* 2010; 51(4):472-96.
34. Erdfelder E, Faul F, Buchner A. GPOWER: a general power analysis program. *Behavior*

- Research Methods, Instruments, & Computers 1996; 28(1):1-11.
35. Beak AT, steer RA, Brown Gk. Beck Depression Inventory for measuring depression. San Antonio 1996; 78(2):490-8.
  36. Fata L, Birashk B, Atefvahid MK, Dabson KS. Meaning assignment structures/ schema, emotional states and cognitive processing of emotional information: comparing two conceptual frameworks. Iranian Journal of Psychiatry and Clinical Psychology 2003; 11(3):312-26. [In Persian].
  37. Dozois DG, Dobson KS, Ahnberg JL. A psychometric evaluation of the beck depression inventory-ii. Psychological Assessment 1998; 10(2):83-9.
  38. Smarr KL, Keefer AL. Measures of depression and depressive symptoms: Beck Depression Inventory-II (BDI-II), Center for Epidemiologic Studies Depression Scale (CES-D), Geriatric Depression Scale (GDS), Hospital Anxiety and Depression Scale (HADS), and Patient Health Questionnaire-9 (PHQ-9). Arthritis Care Res (Hoboken) 2011; 63 (Suppl 11):S453-66.
  39. Beck AT, Epstein N, Brown G, Steer RA. An inventory for measuring clinical anxiety: psychometric properties. J Consult Clin Psychol 1988; 56(6):893-7.
  40. Jackson CJ. Jackson-5 scales of revised Reinforcement Sensitivity Theory (r-RST) and their application to dysfunctional real world outcomes. Journal of Research in Personality 2009; 43(4):556-69.
  41. Hasani J, Salehi S, Azad MR. Psychometric properties of Jackson's five factor questionnaire: scales of revised Reinforcement Sensitivity Theory (r-RST). Journal of Research in psychological Health 2012; 6(3):60-73. [In Persian].
  42. Fayazi M, Hasani J. Structural relations between brain-behavioral systems, social anxiety, depression and internet addiction: with regard to revised Reinforcement Sensitivity Theory (r-RST). Comput Human Behav 2017; 72:441-8.
  43. Li Y, Xu Y, Chen Z. Effects of the Behavioral Inhibition System (BIS), Behavioral Activation System (BAS), and emotion regulation on depression: a one-year follow-up study in Chinese adolescents. Psychiatry Res 2015; 230(2):287-93.
  44. Gray JA. Neurobiology of learning, emotion and affect. New York: Raven Press; 1991.
  45. Takahashi Y, Roberts BW, Yamagata S, Kijima N. Personality traits show differential relations with anxiety and depression in a nonclinical sample. Psychologia 2015; 58(1):15-26.
  46. Struijs SY, Lamers F, Vroling MS, Roelofs K, Spinhoven P, Penninx BW. Approach and avoidance tendencies in depression and anxiety disorders. Psychiatry Res 2017; 256:475-81.
  47. Gable SL, Reis HT, Elliot AJ. Behavioral activation and inhibition in everyday life. Journal of Personality and Social Psychology 2000; 78(6):1135-49.
  48. Corr PJ. Anxiety: splitting the phenomenological atom. Pers Individ Dif 2011; 50(7):889-97.
  49. Ly C. The relevance of reinforcement sensitivity theory to social anxiety and response to cognitive behavioural therapy for social anxiety disorder [dissertation]. Tasmania, Australian: University of Tasmania; 2011.
  50. Mellick W, Sharp C, Alfano C. The role of BIS/BAS in the vulnerability for depression in adolescent girls. Pers Individ Dif 2014; 69:17-21.

51. Hewig J, Hagemann D, Seifert J, Naumann E, Bartussek D. The relation of cortical activity and BIS/BAS on the trait level. *Biol Psychol* 2006; 71(1):42-53.
52. Walkup JT, Albano A, Piacentini J, Birmaher B, Compton SN, Sherrill JT, et al. Cognitive behavioral therapy, sertraline, or a combination in childhood anxiety. *N Engl J Med* 2008; 359(26):2753-66.
53. Levita L, Bois C, Healey A, Smyllie E, Papakonstantinou E, Hartley T, et al. The behavioural inhibition system, anxiety and hippocampal volume in a non-clinical population. *Biol Mood Anxiety Disord* 2014; 4(1):4.
54. Pourmohammad-Rezai-Tajrishi M, Mirzamani-Bafghi M. The relationship between the activity of brain-behavioral systems, social support and depression. *Social Welfare Quarterly* 2007; 7(26):223-46. [In Persian].
55. Mansuri A, Bakhshipour Roudsari A. Relationship between behavioral inhibition and behavioral activation systems with pathological and non-pathological worries. *Journal of Babol University of Medical Sciences* 2010; 12(1):59-64 [In Persian].
56. Basharpour S, Mozafari S. The role of behavioral activation/inhibition systems in prediction of the state/trait anxiety in high school students. *Journal of School Psychology* 2014; 3(4):158-66. [In Persian].
57. Karsazi H, Hashemi Nosratabad T. Structural relationship of brain - behavior systems and difficulty in regulation with social anxiety disorder and depression. *Journal of Thought & Behavior in Clinical Psychology* 2015; 10(37):77-89. [In Persian].
58. Spielberg JM, Heller W, Siltan RL, Stewart JL, Miller GA. Approach and avoidance profiles distinguish dimensions of anxiety and depression. *Cognitive Therapy and Research* 2011; 35(4):359-71.