

## PSYCHOPATHOLOGY IN OFFSPRING OF PARENTS WITH BIPOLAR DISORDER IN AN EGYPTIAN SAMPLE

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### ABSTRACT:

**Background:** *Offspring of parents with Bipolar affective disorder are at risk for a spectrum of future psychiatric disorders. Although the link between family functioning and children's development remained well-established, there is a small research examining whether family factors play a role in helping children with behavior problems outgrow their difficulties.*

**Aim of the work:** *The current study was to identify the risk factors that contributes to the development of psychiatric morbidity, to identify the nature of psychiatric morbidity that affects these children.*

**Patients and methods:** *A sample consist of 200 participants was recruited and divided into 100 offspring of parents diagnosed with bipolar affective disorder using SCID-I, patients were selected from either new or follow up cases, who were attending the outpatient clinics, psychiatry department, Ain Shams University Hospitals) And 100 offspring of parents with irrelevant psychiatric history. Children were also evaluated by The Child Behavior Checklist (CBCL) to screen for psychopathology.*

**Results:** *offspring of parents with Bipolar affective disorder are at increased risk of developing a wide range of psychiatric disorders and accompanying dysfunction than offspring of healthy or non-BP parents. the most prevalent behavior among cases was attention problems 75%, aggression 63%, social problem 48%*

**Conclusion:** *These findings demonstrated the Offspring of BD are at significantly higher risk of developing a broad range of affective and non-affective psychopathology when compared to control offspring.*

**Keywords:** *bipolar disorder, psychopathology, CBCL, offspring.*

### INTRODUCTION:

Bipolar affective disorder (BD) is a neuropsychiatric disorder characterized by extreme fluctuations mood shifts alternating between episodes of mania and depression<sup>(1)</sup>. Depressive episodes are associated with sad mood, suicidality, impaired cognition, and anhedonia whereas Manic symptoms include decreased sleep, high-risk actions, and impulsivity. So, the estimated lifetime

prevalence of BD is 1 to 3 percent worldwide<sup>(2,3)</sup>. There is evidence of genetic influences explaining 60–85% of risk inherited of mood disorder from family members referring a potentially large genetic contribution to the disorder<sup>(4)</sup>.

Genetic factors in bipolar affective disorder tends to be familiar, about half the people with Bipolar affective disorder have family member with a mood disorder, such as depression, ADHD. The person who has one parent with bipolar disorder has a 15 to

25 % chance of having the incidence. A person who has a non-identical twin with the disorder has a 25% chance of disorder, the same risk as if both parents have BD. A person who has an identical twin with Bipolar affective disorder has an even higher risk of developing the disorder about an eight-fold higher risk than a non-identical twin<sup>(5)</sup>.

It is generally held that the offspring of parents with Bipolar affective disorder are at risk for a spectrum of future psychiatric disorders. The degree of risk is an important question for both clinicians and parents. These risks involving conduct disorders, learning problems, neuropsychological deficits, and high rates of internalizing problems. Although the link between family functioning and children's development remained well-established, there is a small research examining whether family factors play a role in helping children with behavior problems outgrow their difficulties. Even rarer are studies that examine factors which predict how well children with behavior problems develop emotionally, socially, and academically, over time. Since children with behavior problems are at risk for a host of negative outcomes. It is critical to identify factors that might increase risk among these children. Parents' psychopathology is one aspect of family functioning that is thought to play an important role in children's development<sup>(6)</sup>.

Compared with the offspring of both psychiatric and non-psychiatric comparison parents, the offspring of bipolar parents showed elevated rates of anxiety disorders, involving anxiety disorder, separation anxiety disorder, and overanxious disorder. Moreover, anxiety disorders in the high-risk offspring were significant predictors of the subsequent onset of diagnosable mood disorders. Given that the BD typically debuts as depressive disorders in high-risk offspring, the finding that anxiety disorders did not exactly predict BD is not unexpected

That is, the cohort are still in the period of risk for manifesting the index activated episode<sup>(7,8)</sup>.

In sum, these parent-child relationships may protect against the impact of genetic vulnerability for BD. Family communication training, problem-solving and behavioral parenting strategies have shown positive consequences in symptom remission in high-risk offspring<sup>(9)</sup>

Additionally, mood liability, anxiety, attention difficulties, hyper-arousal, depression, somatic complaints, and school problems display more symptoms in offspring of bipolar disorder parents than controls, also dysthymic, cyclothymic, or hyperthymic temperaments may presage eventual in bipolar disorder offspring<sup>(10)</sup>

Thus, BD offspring were estimated to be 2.5 times more at risk to develop any psychiatric disorder and four times more at risk to have a mood disorder but, perspective studies in preschool children submitted a prevalence eight times higher for the presence of ADHD, as well as increased frequency of sub-threshold manic and depressive BD<sup>(11)</sup>

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## **PATIENTS AND METHODS:**

### **1. Site of the study**

Participants were selected from the outpatient clinics at the Institute of Psychiatry, Ain Shams University Hospitals. The study was conducted in accordance with the guidelines of the Research and Ethics Committee of the Institute of Psychiatry, Ain Shams University.

### **2. Study design**

It is a Case- Control study comparative study by convenience sampling.

### **3. Subjects:**

A total number of 200 hundred children from participants were recruited and divided

into: One hundred offspring of parents diagnosed with bipolar affective disorder according to diagnostic and statistical Manual of Mental disorders, 4th edition (DSM IV) was recruited from 8- 12 years old, Egyptians only. Another One hundred offspring of selected volunteer healthy parents without any history suggestion of any psychiatric symptoms from 8- 12 years old, Egyptians only.

#### **4. Procedures**

- 1- Informed consent: A written consent was obtained from them after explaining the objectives of the study.
- 2- Arabic version of Structured Clinical Interview for DSM-IV-TR Axis I Disorders (SCID-I)<sup>(12)</sup>: for clinical assessment and diagnosis of substance use disorder and exclude other psychiatric disorders.
- 3- The Child Behavior Checklist (CBCL)<sup>(13)</sup>. Arabic version by<sup>(14)</sup>. to assess of The Children's Psychopathology.

#### **5. Measurements**

##### **1) Structured Clinical Interview for DSM-IV (SCID I)<sup>(12)</sup>.**

The study used the Arabic version of the structured clinical interview for DSM-IV axis I diagnosis (SCID-I). It is a semi-structured diagnostic interview which has been updated for DSM-IV<sup>(15)</sup>. It begins with a section on demographic information and clinical background. Then there are 7 diagnostic modules, focused on different diagnostic groups: mood, psychotic, substance abuse, anxiety, somatoform, eating and adjustment disorders. It is applied to the case group to diagnose drug Dependence and exclude other Axis I diagnosis.

##### **2) The Child Behavior Checklist (CBCL)<sup>(13,14)</sup>.**

The Child Behavior Checklist are questionnaires to be completed, respectively,

by parents, and adolescents themselves and can be scored on eight syndrome scales (Withdrawn, Somatic Complaints, Anxious/Depressed, Social Problems, Thought Problems, Attention Problems, Delinquent Behavior and Aggressive Behavior), and two broad-band groupings of syndromes: Internalizing (consisting of the first three scales), and Externalizing (consisting of the last two scales) The Child Behavior Checklist-school age version (CBCL) was completed by the primary caregiver. The scale includes 118 problem behavior items rated on a Likert scale from zero (not true) to two (true or frequently true). We used the two broadband subscales (Internalizing and Externalizing) and the six DSM-IV oriented subscales (Affective Problems, Anxiety Problems, Somatic Problems, Attention Deficit/Hyperactivity Problems, Oppositional Defiant Problems and Conduct Problems)

##### **3) Statistical analysis.**

All data was tabulated, grouped, and statistically analyzed using (SPSS 20) (Statistical Package for the Social Sciences) on a compatible PC (personal computer). Data was presented, and suitable analysis was done according to the type of data obtained for each parameter. For descriptive statistics: Mean, Standard deviation ( $\pm$  SD) and range for parametric numerical data. while Median and Interquartile range (IQR) for non-parametric numerical data. Frequency and percentage of non-numerical data. Analytical statistics: Student T Test was used to assess the statistical significance of the difference between two study group means. For analytical statistics: Pearson Chi Square Test ( $\chi^2$ ): was used to assess the statistical significance of the difference between two study group means. Fisher's exact test was used to examine the relationship between two qualitative variables when the expected count is less than 5 in more than 20% of cells and **P value**: Used to indicate the level of significance:  $P > 0.05$ : Nonsignificant,  $P <$

0.05: Significant, P < 0.01: Highly significant and P < 0.001: Very highly significant.

significant difference between both groups for age (P-value >0.05). Most of the parents in the study were mothers in both cases and control groups (72%, 78% respectively). There appeared to no statistically significant difference between any of the general characteristics of both cases & control, only the sex of parent showed a statistical difference between the parent was a mother or a father which can be attributed to the fact the mothers exceed the father's number in the study's sample.

**RESULTS:**

**Socio-demographic characteristics:**

Descriptive analysis of the sociodemographic and clinical data of the parents of the children in the whole sample: The general characteristics of the studied participants showed no statistically

Table (1): General characteristics of the studied participants (n=200).

Participants		case		Control		test of significance		
		N	%	N	%	Value	p value	sig.
Child (male/female)	Male	55	55.00%	53	53.00%	X <sup>2</sup> = 0.08	0.777	NS
	Female	45	45.00%	47	47.00%			
Birth-order	1	44	44.00%	44	44.00%	Fisher exact test	>0.999	NS
	2	41	41.00%	41	41.00%			
	3	14	14.00%	14	14.00%			
	4	1	1.00%	1	1.00%			
Marital status	Married	76	76.00%	78	78.00%	Fisher exact test	0.867	NS
	Divorced	23	23.00%	22	22.00%			
	widow	1	1.00%	0	0.00%			
Mother Education	illiterate	18	18.00%	19	19.00%	Fisher exact test	0.711	NS
	Primary school	20	20.00%	19	19.00%			
	secondry school	14	14.00%	18	18.00%			
	Graduate	28	28.00%	30	30.00%			
	Postgraduate	19	19.00%	14	14.00%			
Father Education	illiterate	8	8.00%	10	10.00%	Fisher exact test	0.766	NS
	Primary school	13	13.00%	11	11.00%			
	secondry school	25	25.00%	26	26.00%			
	Graduate	29	29.00%	31	31.00%			
	Postgraduate	25	25.00%	22	22.00%			
Parent (father/mother)	father	28	28.00%	24	24.00%	X <sup>2</sup> = 4.20	0.041	S
	mother	72	72.00%	78	78.00%			

Table (2): Sub-type of the bipolar disorder among parents.

Participants	Type	Bipolar disorder parents	
		N	%
Bipolar disorder type	I	40	40.0%
	II	60	60.0%
duration of disorder by years		11.7	3.3

Table (2) shows that most of the bipolar parents (60%) had bipolar type II, while (40%) had type I. The mean disease duration was 11.7±3(±SD).

Diagram (1): The type of bipolar among the studied parents (N=200).

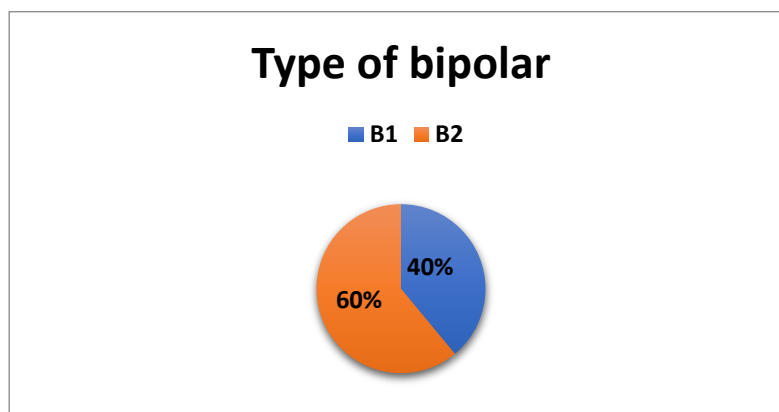


Diagram 1: shows that 60% of recruited cases has bipolar type II while 40% had bipolar type I.

**Comparing the Child Behavior Checklist scores of the children among both groups.**

The Child Behavior Checklist (CBCL) showed no statistically significant difference between both cases and control groups for withdrawn domains (P-value

>0.05), while statistically significant differences were observed between both groups (P-value <0.05) in the other domains, as shown in table (3) and figure (2).

Table (3): The Child Behavior Checklist (CBCL) among the whole sample of participants (n=200)

Participants	Case		control		test of significance		
	N	%	N	%	Value	p value	sig.
Anxious	42	42.00%	20	20.00%	$X^2 = 11.31$	0.001	S
Withdrawn	11	11.00%	8	8.00%	$X^2 = 0.52$	0.469	NS
Somatic Complaints	0	0.00%	4	4.00%	Fisher exact test	0.121	NS
Social Problems	48	48.00%	15	15.00%	$X^2 = 25.24$	<0.001	S
Thought Problems	28	28.00%	4	4.00%	$X^2 = 21.43$	<0.001	S
Attention Problems	75	75.00%	30	30.00%	$X^2 = 40.60$	<0.001	S
Rule Breaking Problems	37	37.00%	11	11.00%	$X^2 = 18.53$	<0.001	S
Aggressive	63	63.00%	24	24.00%	$X^2 = 30.94$	<0.001	S
Other problems	47	47.00%	9	9.00%	$X^2 = 35.81$	<0.001	S

Table (3) showed that, the cases vs control groups showed (42% and 4% respectively) in somatic complaints. moreover, the case group showed higher

scores in anxious symptoms. For social problems, the cases group had more social problem scores (48%) than the control group (15%).

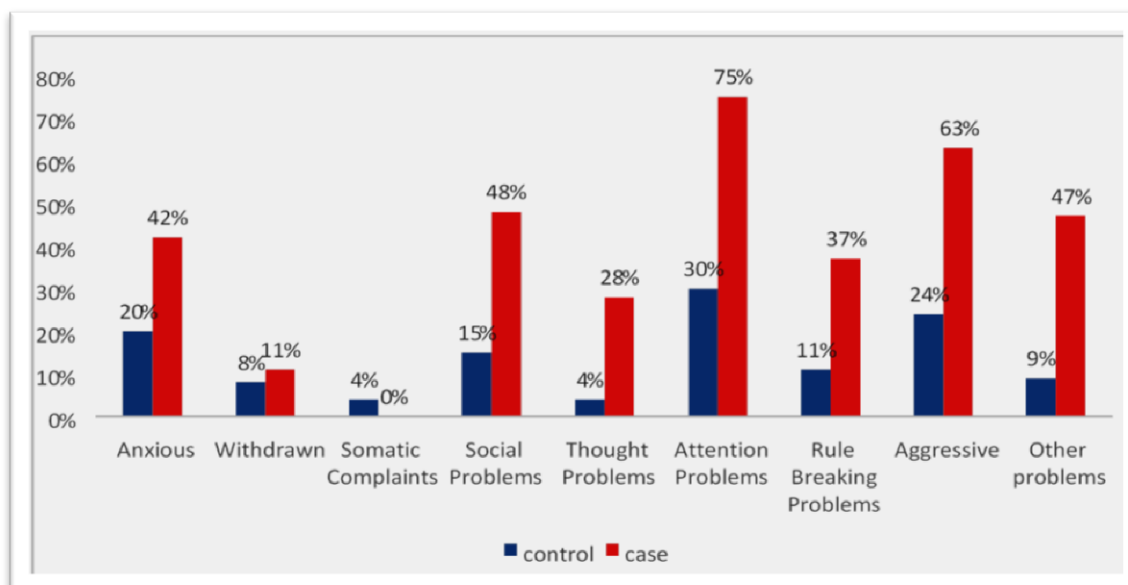


Diagram (2): The comparison in Child Behavior Checklist (CBCL) among the studied participants and control (N=200)

Diagram (2): shows that the most prevalent behavior among both cases and controls was attention problems (75%, 30% respectively) and least was somatic problems which was absent among cases and present in 4% of controls.

## DISCUSSION:

When studying the socio-demographic of the study sample, our findings revealed that, no statistically significant difference in marital status between the 2 groups the bipolar parent's group were married 76.0%, divorced 23.0%, widow 1.0% while the control parent's group married 78.0%, divorced 22.0%. Most of the bipolar parents (60%) had bipolar type II, while (40%) had type I. The disease duration was  $11.7 \pm 3$ . Also, **Ranning and colleagues** <sup>(16)</sup> study showed that the OBP with BP-I had significantly higher rates of BP-I. And offspring of parents with BP-II had significantly higher rates of BP-II in offspring, although this difference did not reach statistical significance in entire age group. The etiology and clinical course of BP are determined by genetic and environmental factors. The same in our present study showed that the older mother's or father's age does not affect the incidence of disease ( $P$ -value  $< 0.05$ ). but there appears to be a significant correlation between age of

the child (The mean  $\pm$ SD  $10.0 \pm 1.5$ ).and the development of anxiety disorders. Yet, it is not apparent to be due to the father's or the mother's age. It can be explained by considering that anxiety disorders can be particularly the prodrome of mood disorder in childhood. On evaluating predictive factors, we found by using The Child Behavior Checklist (CBCL) showed no statistically significant difference between both groups for anxious and withdrawn, while 43% of the sample were diagnosed as Social problems, but Aggressiveness was the most prevailing diagnosis with 82% followed by attention problems with 67%, generalized anxiety disorder with 33%, while Rule Breaking Problems with 31%. And, as regarding somatic complaints all the control group was normal while 17% of the study group were borderline.

**Limaa and colleagues** <sup>(17)</sup> found that the offspring of bipolar parents are at serious risk of being affected by mood disorders. Additionally, data from existing longitudinal studies of the offspring of bipolar parents

have proved that the first manic phase is generally preceded by depressive symptoms. So, Studies focusing on the offspring of affected parents, frequently referred to as the high-risk study design, utilize the well-established familial aggregation of mood disorders as a powerful tool for the identification of risk factors, early clinical manifestations and prodromes of mood disorders in these offspring.

Of note, **Bobo (2017)**<sup>(18)</sup> study's findings from high-risk studies specific to BD pro-bands, which controlled for potential confounding factors have shown that the offspring of BD pro-bands are at much higher risk for MDD, BD, and ADHD than controls. Parents' sociodemographic factors and Bipolar affective disorder history in relation to their offspring's psychopathology. Breaux and colleagues found that the offspring of parents with BP-I were higher risk for BP-I, with a morbidity risk of 13.8% by age 17. These findings are like those of other studies (20.7%), supporting the hypothesis that relatives of pro-bands BP-I have increased risk of suffering BP-I.

**Colorations in anxious symptoms between BP offspring and offspring of control McDonald and Schwanz**<sup>(19)</sup> assumed that the subgroup of offspring of BD who will develop major affective disorders in adulthood will have inherited a tendency for high levels of neuroticism. So, in our present study, we found that the cases of our sample (42,0%, n=42) had anxious symptoms on CBCL scale while the control was (20,0%, n=20) and this finding is interesting and statistically significant. Also, the results of our study supported the hypothesis that diathesis of bipolar disorder itself includes a risk of anxiety. This finding is particularly noticeable given studies in both adults and children with bipolar affective disorder that observed high rates of comorbidity with anxiety disorders, especially panic disorder, GAD, and obsessive-compulsive disorder. Also, we

found that the only factor significant among cases is the age of OBD. Additionally, **Ellenbogen & colleagues**<sup>(20)</sup> found that BD offspring that present with symptoms of depression, anxiety and/or ADHD at a young age appear to be at highest risk for developing BD. Also, **Moreno and colleagues**<sup>(21)</sup> found early onset of an anxiety disorder remained to be the most consistent and reliable indicator of subsequent affective disorder in BD high-risk subjects. A pre-existing anxiety disorder among high-risk offspring more than doubled the risk of having a major affective disorder approximately 8 years after the initial diagnosis, even after adjusting for parental anxiety and behavioral disorders. But, in **Merikukka et al.**<sup>(22)</sup> study showed that there were no statistically significant differences between the two groups for anxiety and withdrawal (P-value > 0.05). This finding was proven by life pressures and increased level of anxiety in the non-bipolar offspring as the bipolar offspring. Also, in **Zavaleta-Ramírez et al.**<sup>(23)</sup> study Anxiety disorders occurred more frequently in children of non-bipolar parents. There finding was 71% of offspring of non-bipolar parents' verses 58% of offspring of bipolar parents that was related to an increased sensitivity to parents' recognition of psychopathology when accepting that their children were assessed. **Biederman and colleagues**<sup>(24)</sup> noted that, the presence of anxiety disorder in biological parents was associated with psychopathology in the offspring of BP, but this meant that it was lost after controlling for bipolar diagnosis. **Chorot and colleagues**<sup>(25)</sup> observed that, offspring with anxiety diagnoses were more likely to have a history of both BD mothers and BD fathers. Although these results were not statistically significant (i.e., P-value < 0.05) for offspring of BD fathers, this may be a result of limited power given the similar effect sizes between offspring of BD mothers and BD fathers. A relationship between offspring control cases and

subsequent offspring anxiety remained observed in preceding studies of offspring of parents with and without psychopathology.

### **Colorations in withdrawal symptoms between BP offspring and control**

In our study showed no statistically significant difference between both groups for withdrawn with mean ( $\pm$ SD)  $60.3 \pm 7.2$  in cases and  $61.4 \pm 6.6$  in control. this finding may be demonstrated by the stressors of life was increased the technology lifestyle in non-bipolar offspring as bipolar offspring. Although, **Dilsaver and colleagues** <sup>(26)</sup> pointed out that aspects of the parent-child relationship are related to the ease with which the offspring infiltrates leaving the home leaving the home, and there is much less research examining how pathological psychology or parental impairment affects this process. Studies have documented that caregivers with mental health problems are more withdrawn from their offspring and show less and more hostile sensitivity to them. Additionally, parents with BD depend on their offspring to provide them with emotional and instrumental care-taking more than other caregivers. Thus, because parents with affective disorders may be more dependent on their children, they may be less likely to help guide and support the emerging adult's navigation of the leaving home transition. <sup>(27)</sup>

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## الأعراض النفسية المرضية لأبناء مرضى الأضطراب الوجداني في عينه من المصريين

**المقدمة:** تشير الدراسات أن هناك ٦٠٪ من أسر الأضطراب الوجداني مصابون بنفس الأضطراب وهذا يدل على المساهمات الجينية الكبيرة المحتملة في هذا الأضطراب و أيضًا يؤكد إلي أن العوامل الوراثية تكون مألوفة في الأضطراب الوجداني ثنائي القطب فأكثر من نصف الأشخاص المصابين بهذا الأضطراب لديهم أفراد من الأسرة يعانون من إضطراب من إضطرابات المزاج، مثل الاكتئاب. ويعتبر أولاد مرضى الأضطراب الوجداني لديهم فرصة بنسبة ١٥ للإصابة أما بالنسبة لشخص لديه توأم غير متطابق فتزيد النسبة إلي ٢٥٪ وهو نفس الخطر كما لو كان الوالدين مصابان بالأضطراب الوجداني كما يعد الشخص الذي لديه توأم متطابق لديه خطر أعلى من الإصابة بالأضطراب يبلغ ثمانية أضعاف أعلى من التوأم غير المتطابق.

**الهدف من العمل:** هذه الدراسة هو ربط عوامل الخطر المختلفة في الأطفال الذين يعانون من عوامل الخطر المختلفة في الأباء المصابين بإضطراب ثنائي القطب.

في الأضطراب الوجداني ثنائي القطب.

**المرضى و الأساليب:** شملت الدراسة مائة من أبناء مرضى الأضطراب الوجداني من عمر ٨ الي ١٢ سنة ومائه اخري من أبناء الاشخاص الذين ليس لا يعانون من اي إضطرابات نفسيه في نفس العمر وأجريت المقابلة الإكلينيكية المنشئة للتشخيص الإحصائي الأمريكي SCID-1 وأجريت مقياس سلوك الطفل باستخدام النسخة العربية المترجمة من اختبار سلوك الأطفال .

**النتائج:** أشارت النتائج باستخدام اختبار سلوك الطفل CBCL إلي عدم وجود دلالة إحصائية واضحة لإضطراب الانسحاب لكل من العينة الخاصة بالحالات والعينة الضابطة. ولكن يوجد دلالة احصائية في إضطراب نقص الانتباه وفرط الحركة بنسبة ٧٥٪ في عينة الحالات يليه العدوان بنسبة ٦٣٪ ثم إضطراب الخوف المرضي من المجتمعات فقد وُجد بنسبة ٤٨٪ في عينة الحالات بينما إضطرابات السلوك التخريبية فكان بنسبة ٣٧٪.

**الخاتمة:** توصلت هذه الدراسة إلى إرتفاع معدلات الاعراض النفسية المرضية في أبناء مرضى الاضطراب الوجداني مقارنة بنويعهم من أبناء العينة الضابطة ولذلك نحتاج إلى ضرورة تحديد المظاهر المحتملة للمرض ولتحديد استراتيجيات التدخل المبكر لأبناء المرضى لإنقاذ جيل من المزيد من الأضطرابات النفسية.