Published online 2022 March 15.

**Research Article** 

# Anxiety Disorders in Children with Functional Constipation: A Case-Control Study

Reza Saber Sangari<sup>1</sup>, Mojtaba Hashemi<sup>2,\*</sup>, Bahman Salehi<sup>3</sup>, Parsa Yousefichaijan <sup>1</sup>, Mohamad Rafiei<sup>5</sup> and Masoud Rezagholizamenjany <sup>1</sup>,<sup>\*\*</sup>

<sup>1</sup>School of Medicine, Arak University of Medical Sciences, Arak, Iran

<sup>2</sup>Department of Pediatric Gastroenterologist, Amir Kabir Hospital, Arak University of Medical Sciences, Arak, Iran

<sup>3</sup>Department of Psychiatry, Arak University of Medical Sciences, Arak, Iran

<sup>4</sup> Department of Pediatric Nephrology, Amir Kabir Hospital, Arak University of Medical Sciences, Arak, Iran

<sup>5</sup>Department of Biostatistics, Arak University of Medical Sciences, Arak, Iran

Corresponding author: Department of Pediatric Gastroenterologist, Amir Kabir Hospital, Arak University of Medical Sciences, Arak, Iran. Email: mojtaba53hashemi@yahoo.com

"Corresponding author: School of Medicine, Arak University of Medical Sciences, Arak, Iran. Email: masoudrezagholi074@gmail.com

Received 2021 October 02; Revised 2022 January 18; Accepted 2022 January 30.

## Abstract

**Background:** Functional constipation as a common disorder in children, may be induce by psychiatric disorders. The present study aimed to evaluate and compared anxiety disorders in children with and without functional constipation.

**Methods:** This case-control study was conducted on 200 children (100 cases with functional constipation and 100 children without functional constipation) aged 15 - 8 years. The participants' age and gender were evaluated and compared in two groups. Moreover, the Spence Children Anxiety Scale (SCAS) (parent form) for children was completed. Finally, the collected data were analyzed using SPSS software version 24.

**Results:** Of 200 evaluated cases in the case and control groups, there were 46 (46.0%) and 59 (59.0%) males, respectively (P = 0.113). The mean  $\pm$  SD of age in total, the case, and the control groups were 7.51  $\pm$  1.91, 7.61  $\pm$  0.91, and 7.41  $\pm$  2.49 years, respectively (P = 0.585). Furthermore, SCAS in the case group was significantly higher (33.06  $\pm$  14.4 vs. 24.8  $\pm$  14.9, P = 0.001). Furthermore, the mean of separation anxiety disorder (P = 0.001), generalized anxiety (P = 0.003), and obsessive-compulsive disorder (OCD) were significantly higher in the functional constipation group (P = 0.001).

**Conclusions:** Anxiety was higher in children with functional constipation than in children without functional constipation (ie, control group), among which separation anxiety, generalised anxiety, and OCD were significantly different in the two groups.

Keywords: Anxiety, Children, Functional Constipation, Obsessive-Compulsive Disorder, Separation Anxiety, Generalized Anxiety

#### 1. Background

Functional constipation (FC), as a prevalent condition in children, is estimated to have a 9.5% prevalence rate worldwide (1). A clinical-based approach, which is currently based on the pediatric diagnostic Rome IV criteria, is used to diagnose FC in children (2). The etiology and pathophysiology of constipation in pediatrics is multifactorial, and many factors, including low-socioeconomic status, genetic predisposition, immobility, inadequate fluid, and daily fiber intake, may affect this condition (3-5). However, there are still many unknown risk factors. The most common and well-known etiological factor in children is withholding behavior, which usually happens after experiencing the frightening or painful defecation of stools, which may lead to and exacerbated by an anxiety disorder (1). This behavior is associated with incomplete feces defecation, fecal impaction, dyssynergic evacuation, decreased rectal sensation, and overflow fecal incontinence (2, 6).

FC puts a heavy burden on children and their parents (7), and this condition is associated with a decrease in quality of life, decreased academic performance, and psychological disorders including depression, aggression, emotional reactivity, and anxiety (7, 8). Moreover, FC is more common in children with psychological problems such as autism spectrum disorders (ASDs), attention deficit hyperactivity disorder (ADHD), and different types of anxiety (2, 9, 10). Another critical disease in such children is less frequent urinary tract dysfunctional disorders such as enuresis, which can simultaneously be observed with FC, thereby arousing further anxiety in children and parents (11, 12). In these cases, where psychological problem is associated

Copyright © 2022, Author(s). This is an open-access article distributed under the terms of the Creative Commons Attribution-NonCommercial 4.0 International License (http://creativecommons.org/licenses/by-nc/4.0/) which permits copy and redistribute the material just in noncommercial usages, provided the original work is properly cited.

with FC, and there is two-way communication between the two variables, as stress can also affect FC, symptoms may persist by adulthood. Although such cases have received adequate laxative treatment, no improvement in the cases is observed (13).

# 2. Objectives

The present study aimed to evaluate and compare anxiety disorders in children with and without FC.

## 3. Methods

# 3.1. Study Setting

This case-control study was conducted on patients with FC in Amir-Kabir hospital.

#### 3.2. Study Population

The study population encompassed patients with FC in the case group, and children without FC in the control group. The samples were selected using the nonrandom convenience sampling method. In the case of selection, the matching method was also considered as such the children in the two groups were homogenous in terms of age and gender. One hundred seventy-two cases were included in the case group, and there were 145 children in the control group. Considering inclusion and exclusion criteria and according to the sample size, 100 children with FC and 100 children without FC participated in this study.

The control group was selected from children without FC, who referred to the pediatric clinic of Amir Kabir Hospital on an outpatient basis, and the case group included children who were diagnosed with FC considering their medical history and medical records, and a pediatric gastroenterologist approved the diagnosis.

#### 3.3. Inclusion Criteria

Children aged 8 - 15 years, a history of FC with confirmed diagnosis, and informed consent from the children's parents.

#### 3.4. Exclusion Criteria

- Positive history of chronic and prevalent psychiatric disorders such as depression, anxiety disorders (for the case group before constipation), bipolar disorder, ADHD, and others,

- Mental retardation (having an intelligence quotient (I.Q) < 70 based on previous diagnosis documents or an I.Q test if needed),

- Positive history of chronic medical diseases such as epilepsy, diabetes, congenital syndromes, immunodeficiency, malignancies, and others,

- History of major psychiatric illnesses in the grade 1 of families,

- History of developmental disorders in infancy and childhood,

- History of moderate to severe head trauma,

- History of chronic gastrointestinal disorders and congenital anomalies in the gastrointestinal tract such as anorectal anomalies and Hirschsprung's disease, and

- A history of parents' death or separation.

According to the aforementioned criteria, 200 children were selected, compared, and analyzed.

#### 3.5. Measurements

FC in children was defined and confirmed as follows (14):

- Bowel movements two or fewer times a week or defeating complicated and bulky stools for at least two weeks as constipation definition.

- Presence of the following in the history and clinical examination to diagnose FC: withholding behavior with a certain position, increased stool diameter and painful bowel defecation, overflow incontinency, and rectal ampulla containing stools.

- Rejection of organic causes of constipation and other differential diagnoses (to confirm FC), including congenital anomalies of the gastrointestinal tract such as anorectal anomalies, Hirschsprung's disease, hypothyroidism, hypocalcemia, poisoning, and celiac disease. It should be noted that FC was diagnosed and confirmed according to pediatric gastroenterologist diagnosis.

Then the parents and children of the two groups were interviewed to evaluate their anxiety disorders using SCAS. This scale of anxiety disorders in children addresses:

- (1) Panic agoraphobia
- (2) Separation anxiety
- (3) Fear of physical injury
- (4) Social phobia
- (5) OCD, and
- (6) Generalized anxiety.

After completing the questionnaires and identifying children with anxiety disorders using SCAS, they were referred to the psychiatrist to confirm their diagnosis based on the DSM-IV-TR anxiety diagnostic criteria. Finally, anxiety and the subcategories of this disorder were compared in the two groups.

## 3.6. Ethical Considerations

The study group observed the principles of medical ethics introduced by the Health Ministry and the Declaration of Helsinki and legislation in the medical ethics committee of the Arak University of Medical Sciences. Moreover, the ethical committee of the Arak University of Medical Sciences approved the research protocols (Ethical code: IR.ARAKMU.REC.1393.171.21). Research resources bore the cost of using the drug. This investigation had no effect on the patients' treatment process, and the assessment results were only reported to the patients later.

#### 3.7. Statistical Analysis

The collected data were analyzed by SPSS software version 24 at a 95% confidence level, and descriptive statistical methods were used to determine the frequency of the research variables. Further, Student's *t*-test was used to analyze the quantitative and qualitative variables. Spearman's test was also used to examine the correlation between constipation duration and anxiety disorders. In this study, P < 0.05 was considered as the significance level.

# 4. Results

#### 4.1. Demographic Characteristics

Of 200 cases, in the case and control groups, there were 46 (46.0%) and 59 (59.0%) males, respectively (P=0.113). The mean  $\pm$  SD of age in total, the case, and the control groups were 7.51  $\pm$  1.91, 7.61  $\pm$  0.91, and 7.41  $\pm$  2.49 years, respectively (P=0.585) (Table 1).

#### 4.2. Anxiety Disorder

The SCA score was significantly higher in the case group than in the control group ( $33.06 \pm 14.4 \text{ vs.} 24.8 \pm 14.9$ , P = 0.001). Although panic agoraphobia (P = 0.854), fear of physical injury (P = 0.962), and social phobia (P = 0.471) were not significantly different between the two groups, the mean of separation anxiety (P = 0.001), generalized anxiety (P = 0.003), and OCD (P = 0.001) were significantly higher in children with FC in the case group than children without FC as the control group (Table 2).

Moreover, the results of the Spearman's test for the correlation between constipation duration and anxiety disorders indicate that constipation duration was significantly associated with panic agoraphobia (P = 0.041), social anxiety (P = 0.037), and OCD (P = 0.0001) (Table 3).

# 5. Discussion

The findings revealed that anxiety was higher in children with FC than children without FC as the control group. Of the concerned indices, separation anxiety, generalized anxiety, and OCD were significantly different in the two groups, and they were significantly higher in the case group than the control group. Hosseinzadeh et al. reported that depression and anxiety had a prevalence rate of 23.5% and 34.6% in chronic constipation, based upon screening tools, MINI, and the HADS questionnaire (15); however, the methods in this study were different, thereby justifying the differences in the findings.

Among studies examining psychological conditions in gastrointestinal disorder cases, Nehra et al. (11) retrospectively reviewed the patients with evacuation disorders, who had referred for biofeedback retraining to a tertiary referral center. They investigated their cases for psychological disorders considering the DSM-IV criteria and observed that patients with evacuation disorder and constipation had a 65% psychological impairment, and the most common disorders were anxiety and depression (10). Mason et al. investigated the relationships between psychological morbidity and idiopathic constipation. They evaluated psychological disorders using the General Health Questionnaire (GHQ-28) and documented that constipated cases had significantly higher levels of anxiety (P < 0.022) compared to children without FC as the control group (16). In another study, Rezakhaniha et al. found out that the flaccid measurement could be as helpful as stretched measurement if it is done from pubic bone to tip of glans (17). Furthermore, Rezakhaniha et al. mentioned that penile length was positively correlated with height and testosterone but not with weight and estradiol in non-obese and obese children (18). These two studies addressed anthropometric data, which are different from the present study.

In three studies, Taylor et al. (1), Hutson et al. (19), and van den Heuvel et al. (20) showed that FC in children could lead to psychological disorders, low self-esteem, academic failure, and behavioral disorders. Yacob et al. also observed that children with depression and anxiety are at higher risk of FC (21). Waters et al. evaluated FC in children with anxiety disorders. They showed that FC and irritable bowel syndrome with a prevalence rate of 40.7%, were more frequent in children with anxiety, than the control group with the prevalence rate of 5.9% (22). In addition to anxiety, some authors have reported a significant association between other psychiatric disorders and FC in children. In this regard, Peeters et al. (9) and Mckeown et al. (23) observed a significant relationship between FC and autism and ADHD in children, respectively.

Regarding recent psychodynamic models, there

Fable 1. Participants	' Age and Gender in T	wo Groups <sup>a</sup>
-----------------------	-----------------------	------------------------

Variables		Groups		Statistical Value
	Case	Control	Total	Statistical value
Age	$7.61\pm0.91$	$7.41 \pm 2.49$	$7.51 \pm 1.91$	0.585 <sup>b</sup>
Gender				0.113 <sup>c</sup>
Male	46(46.0)	59 (59.0)	105 (52.5)	
Female	54 (54.0)	41 (41.0)	95 (47.5)	
<sup>a</sup> Values are expressed as mean SD or No. (%).				

<sup>b</sup> Independent sample *t*-test

<sup>c</sup> Chi-square test

Table 2. Participants' SCAS in Two Groups

Variables	Groups		P.Value <sup>a</sup>
	Functional Constipation	Control	1-value
Panic agoraphobia	$4.54\pm2.28$	$2.94 \pm 3.06$	0.845
Separation anxiety	$6.72\pm3.77$	$5.22\pm3.02$	0.001
Fears of physical injury	$6.40\pm2.43$	$6.09 \pm 4.00$	0.962
Social phobia	$4.80\pm4.06$	$3.72\pm2.92$	0.471
OCD	$5.22\pm3.43$	$3.30\pm4.08$	0.001
Generalized anxiety	$5.90\pm3.33$	$3.70\pm2.70$	0.003
Total	$33.06 \pm 14.40$	$24.8\pm14.9$	0.001

<sup>a</sup> Independent sample *t*-test

Table 3. Correlation Coefficient of Functional Constitution Duration and Anxiety Disorders

Constipation Duration	Panic Agoraphobia	Separation Anxiety	Fears of Physical Injury	Social Phobia	OCD	Generalized Anxiety	Total
P-value <sup>a</sup>	0.041	0.631	0.143	0.037	0.001	0.087	0.633
Correlation coefficient	0.2	0.04	-0.15	0.2	0.4	0.017	0.04

<sup>a</sup> Spearman test

are three different ways to express anxiety (11): cognitive/perceptual distortions, smooth muscles, and striated muscles. Dysfunctions of smooth muscle-like constipation, diarrhea, shortness of breath, IBS, and so on may be related to anxiety that is so strong that it inhibits the function of the striated muscles. Accordingly, a patient with smooth muscle disorders in response to anxietyprovoking situations shows more problems with adapting to stressful situations as such, helping patients with constipation to express their emotions and anxiety and eliminate their stressful condition and anxiety using psychotherapeutic methods could reduce constipation symptoms. Given the lack of studies on anxiety disorders in children with FC, further studies in this field are recommended. However, based on the findings of the present study, further studies on children with FC (especially

long-term cases of this disorder) is recommended. In this study, the mutual communication of FC and anxiety was similar to those reported in previous studies.

# 5.1. Conclusions

Our study showed an association between anxiety disorders and FC. Accordingly, anxiety was higher in children with FC than children without FC as the control group. Of these indices, separation anxiety, general anxiety, and OCD were significantly different in the two groups.

# Footnotes

Authors' Contribution: All authors performed equally in manuscript preparation and submission. Conflict of Interests: We declare no conflict of interests.

**Ethical Approval:** The study group adhered to the principles of medical ethics introduced by the Health Ministry and the Declaration of Helsinki and legislation in medical ethics committee of Arak University of Medical Sciences. In addition, the ethical committee of Arak University of Medical Sciences approved the protocols of the study. The cost of using the drug was borne by research resources. This investigation did not have any effect on the patients treatment process, and the result of the assessment was only reported to the patient later. (Ethical code: IR.ARAKMU.REC.1393.171.21)

**Funding/Support:** Our study was funded by the Arak University of Medical Sciences.

**Informed Consent:** Informed consent was obtained from all patients.

## References

- Taylor E, Sandberg S, Thorley G, Giles S. *The Epidemiology of Childhood Hyperactivity. Maudsley Monographs*. Oxford University Press; 1991. p. 60–8.
- Kuizenga-Wessel S, Koppen IJN, Vriesman MH, Di Lorenzo C, van Dijk M, Beelen MLR, et al. Attention Deficit Hyperactivity Disorder and Functional Defecation Disorders in Children. *J Pediatr Gastroenterol Nutr.* 2018;66(2):244–9. doi: 10.1097/MPG.000000000001695. [PubMed: 28742722].
- Mugie SM, Di Lorenzo C, Benninga MA. Constipation in childhood. Nat Rev Gastroenterol Hepatol. 2011;8(9):502–11. doi: 10.1038/nrgastro.2011.130. [PubMed: 21808283].
- Duan S, Liu L, Li G, Wang J, Hu Y, Zhang W, et al. Altered Functional Connectivity Within and Between Salience and Sensorimotor Networks in Patients With Functional Constipation. *Front Neurosci.* 2021;**15**:147. doi: 10.3389/fnins.2021.628880. [PubMed: 33776637]. [PubMed Central: PMC7991789].
- Liu L, Hu C, Hu Y, Zhang W, Zhang Z, Ding Y, et al. Abnormalities in the thalamo-cortical network in patients with functional constipation. *Brain Imaging Behav*. 2021;15(2):630–42. doi: 10.1007/s11682-020-00273y. [PubMed: 32314199].
- Piccoli de Mello P, Eifer DA, Daniel de Mello E. Use of fibers in childhood constipation treatment: systematic review with meta-analysis. *J Pediatr (Rio J)*. 2018;**94**(5):460–70. doi: 10.1016/j.jped.2017.10.014. [PubMed: 29474804].
- Kaugars AS, Silverman A, Kinservik M, Heinze S, Reinemann L, Sander M, et al. Families' perspectives on the effect of constipation and fecal incontinence on quality of life. *J Pediatr Gastroenterol Nutr.* 2010;**51**(6):747–52. doi: 10.1097/MPG.0b013e3181de0651. [PubMed: 20706148].
- Williams KC, Rogers LK, Hill I, Barnard J, Di Lorenzo C. PEG 3350 Administration Is Not Associated with Sustained Elevation of Glycol Levels. *J Pediatr.* 2018;195:148–153 e1. doi: 10.1016/j.jpeds.2017.11.028. [PubMed: 29395181].
- 9. Peeters B, Noens I, Philips EM, Kuppens S, Benninga MA. Autism spectrum disorders in children with functional defecation disorders. J

*Pediatr.* 2013;**163**(3):873-8. doi: 10.1016/j.jpeds.2013.02.028. [PubMed: 23522863].

- Kuizenga-Wessel S, Di Lorenzo C, Nicholson LM, Butter EM, Ratliff-Schaub KL, Benninga MA, et al. Screening for autism identifies behavioral disorders in children functional defecation disorders. *Eur J Pediatr*. 2016;**175**(10):1371–8. doi: 10.1007/s00431-016-2775-x. [PubMed: 27624626]. [PubMed Central: PMC5031734].
- Nehra V, Bruce BK, Rath-Harvey DM, Pemberton JH, Camilleri M. Psychological disorders in patients with evacuation disorders and constipation in a tertiary practice. *Am J Gastroenterol*. 2000;**95**(7):1755–8. doi:10.1111/j.1572-0241.2000.02184.x. [PubMed: 10925980].
- Li G, Zhang W, Hu Y, Wang J, Li J, Jia Z, et al. Distinct Basal Brain Functional Activity and Connectivity in the Emotional-Arousal Network and Thalamus in Patients With Functional Constipation Associated With Anxiety and/or Depressive Disorders. *Psychosom Med.* 2021;83(7):707-14. doi: 10.1097/PSY.0000000000000958. [PubMed: 34117157].
- Bongers ME, van Wijk MP, Reitsma JB, Benninga MA. Long-term prognosis for childhood constipation: clinical outcomes in adulthood. *Pediatrics*. 2010;**126**(1):e156–62. doi: 10.1542/peds.2009-1009. [PubMed: 20530072].
- Marcdante K, Kliegman RM, Jenson H, Behrman R. Essentials of pediatrics. *Elsevier*. 2015;11(14):231.
- Hosseinzadeh ST, Poorsaadati S, Radkani B, Forootan M. Psychological disorders in patients with chronic constipation. *Gastroenterol Hepatol Bed Bench*. 2011;4(3):159–63. [PubMed: 24834176]. [PubMed Central: PMC4017427].
- Mason HJ, Serrano-Ikkos E, Kamm MA. Psychological morbidity in women with idiopathic constipation. *Am J Gastroenterol.* 2000;95(10):2852-7. doi: 10.1111/j.1572-0241.2000.02313.x. [PubMed: 11051359].
- Rezakhaniha S, Rezakhaniha B, Siroosbakht S. Which of the Anthropometric Parameters and Hormonal Variables in Prepubertal Children Are Correlated to True Micropenis? *Iran J Pediatr.* 2021;**31**(1). doi: 10.5812/ijp.108534.
- Rezakhaniha S, Rezakhaniha B, Aarabi N, Siroosbakht S. Is It Necessary to Weight Loss in Obese Boys with Small Penile Length? A Case-Control Study. J Compr Pediatr. 2020;11(4). doi: 10.5812/compreped.107272.
- Hutson JM, O'Brien M, Beasley SW, Teague WJ, King SK, editors. Jones' clinical paediatric surgery. John Wiley & Sons; 2015.
- van den Heuvel E, Starreveld JS, de Ru M, Krauwer V, Versteegh FG. Somatic and psychiatric co-morbidity in children with attention deficit hyperactivity disorder. *Acta Paediatr.* 2007;96(3):454–6. doi: 10.1111/j.1651-2227.2006.00145.x. [PubMed: 17407478].
- Yacob D, Di Lorenzo C, Bridge JA, Rosenstein PF, Onorato M, Bravender T, et al. Prevalence of pain-predominant functional gastrointestinal disorders and somatic symptoms in patients with anxiety or depressive disorders. *J Pediatr.* 2013;163(3):767-70. doi: 10.1016/j.jpeds.2013.02.033. [PubMed: 23522860].
- Waters AM, Schilpzand E, Bell C, Walker LS, Baber K. Functional gastrointestinal symptoms in children with anxiety disorders. J Abnorm Child Psychol. 2013;41(1):151–63. doi: 10.1007/s10802-012-9657-0. [PubMed: 22773359].
- McKeown C, Hisle-Gorman E, Eide M, Gorman GH, Nylund CM. Association of constipation and fecal incontinence with attentiondeficit/hyperactivity disorder. *Pediatrics*. 2013;**132**(5):e1210–5. doi: 10.1542/peds.2013-1580. [PubMed: 24144702]. [PubMed Central: PMC4530301].