

Journal of Medical Sciences

ISSN 1682-4474









JMS (ISSN 1682-4474) is an International, peer-reviewed scientific journal that publishes original article in experimental & clinical medicine and related disciplines such as molecular biology, biochemistry, genetics, biophysics, bio-and medical technology. JMS is issued eight times per year on paper and in electronic format.

For further information about this article or if you need reprints, please contact:

Maryam Izadi-Mazidi Department of Clinical Psychology, Shahed University, Tehran, Iran

Tel : 09171346040 Fax : 02144159668 J. Med. Sci., 15 (5): 235-240 1st July, 2015 DOI: 10.3923/jms.2015.235.240

Anxiety and Quality of Life in patients with Irritable Bowel Syndrome

¹Shahnaz Mohammadi, ²Maryam Izadi-Mazidi and ¹Mahsa Akbarian-Firoozabadi

Irritable Bowel Syndrome (IBS) is a prevalent gastroenterological problem that has no identified pathology. There is evidence to suggest that patients with IBS suffer from mental disorders and impairment in quality of life. The aim of this study was to compare the anxiety and quality of life in iranian individuals with and without irritable bowel syndrome. A total of 60 patients with Irritable Bowel Syndrome (IBS) and 60 healthy individuals participated in the study. The data was gathered using Cattle anxiety scale, Short Form-36 Health Survey (SF-36), Rome III diagnostic questionnaire and demographic forms and were analyzed through multivariate analyses of variance (MANOVA), Chi-square test and Fisher's exact test. The results revealed that patients with IBS have more anxiety (F = 287.48, p<0.001) and less quality of life (F = 151.7, p = 0.002) compared to healthy individuals. IBS was significantly more prevalent in females and individuals who was living in urban areas (p<0.05). But, there was no significant difference in prevalence of IBS among individuals with different levels of education (p = 0.53). The IBS is associated with higher rates of anxiety and lower quality of life. This highlights the importance of management of these patients and assessing the impact of psychological interventions.

Key words: Irritable bowel syndrome, Cattle anxiety scale, Rome III criteria, alexithymia, gastrointestinal specific anxiety



¹Department of Psychology, Kharazmi University, Tehran, Iran ²Department of Clinical Psychology, Shahed University, Tehran, Iran

INTRODUCTION

Irritable Bowel Syndrome (IBS) is a prevalent gastroenterological problem that causes suffering from abdominal pain, bloating and altered bowel habits (Canavan *et al.*, 2014; Dainty *et al.*, 2014; Roberts *et al.*, 2013).

This chronic and relapsing disorder (Lee et al., 2009) occurs in all age groups (Canavan et al., 2014). Existing estimates indicate that global prevalence of IBS is about 4-30% (Ghoshal and Srivastava, 2014) and patients tend to be young and middle-aged women (Canavan et al., 2014; Lorenzo-Zuniga et al., 2014). Irritable bowel syndrome interfere with the affected individuals' health related quality of life (Lorenzo-Zuniga et al., 2014; DiBonaventura et al., 2011; Li et al., 2003; Creed et al., 2001; El-Serag et al., 2002), in spite of its benign long term prognosis (Lorenzo-Zuniga et al., 2014). And there is evidence to suggest that the costs of somatic diagnostic procedures and medical treatment attempts on IBS are enormous (Lorenzo-Zuniga et al., 2014; Levy et al., 2001; Nyrop et al., 2007). It has also indirect costs related to deceased productivity and efficacy at work (Lorenzo-Zuniga et al., 2014).

The exact cause of IBS is poorly understood (Mykletun et al., 2010; Folks, 2004). And beside different hypothesis about physical factors such as colonic motility disturbances, gastrointestinal hypersensitivity, gastrointestinal inflammation (Lorenzo-Zuniga et al., 2014; Muscatello et al., 2014; Hauser et al., 2014), abnormalities in the gut flora (Khanna and Tosh, 2014), small intestinal bacterial overgrowth (Ohman and Simren, 2010) and disruption of the brain-gut axis (Lin, 2004), psychological conditions have been reported to be an important contributor to production or exacerbation of IBS (Lee et al., 2009; Mykletun et al., 2010; Walker et al., 1995; Osterberg et al., 2000; Gulewitsch et al., 2013; Lackner et al., 2013). More than 90% of patients with IBS experience at least one mental disorder during their lifetime (Liss et al., 1973; Fossey and Lydiard, 1990; Irwin et al., 1996; Lydiard, 2001).

Anxiety disorders (Lee *et al.*, 2009; Yi *et al.*, 2014; Yacob *et al.*, 2013), depression (Lee *et al.*, 2009; Yi *et al.*, 2014; Yacob *et al.*, 2013; Walker *et al.*, 1995), panic disorder (Paras *et al.*, 2009), childhood sexual abuse (Yacob *et al.*, 2013; Van Tilburg *et al.*, 2010; Mahvi-Shirazi *et al.*, 2013) PTSD (Hauser *et al.*, 2014), phobia (Van Tilburg *et al.*, 2010) and somatoform disorders (Surdea-Blaga *et al.*, 2012) are psychological conditions that has been reported to be associated with IBS. And mood disorders, anxiety disorders and somatoform disorders are most prevalent (Surdea-Blaga *et al.*, 2012). Gastrointestinal Specific Anxiety (GSA) and alexithymia also are two psychological constructs that may contribute to severity of irritable bowel syndrome (Porcelli *et al.*, 2014). In the study conducted by Niesten *et al.* (2014), a childhood history of non-sexual abuse and a family history of IBS were predictors of IBS diagnosis in women with bipolar disorder (Niesten *et al.*, 2014). Some results however challenge the plausibility of the association between IBS and psychological problems (Talley *et al.*, 2001).

Because of these controversial findings and little knowledge about psychological conditions and health related quality of life of patients with IBS in Iran, the present study aimed to compare anxiety and quality of life in patients with IBS and healthy individuals in Iranian population.

Since poor treatment outcome in patients with IBS may partly due to undetected comorbid psychological disorders (Lee *et al.*, 2009), identifying mental disorders help clients in management of their patients.

MATERIALS AND METHODS

Subjects: Research was conducted over a period of 12 months from 2013-2014. A total of 60 patients with Irritable Bowel Syndrome (IBS) and 60 healthy individuals participated in the study. The patients selected from referral centers in Tehran, Iran. The diagnosis of Irritable Bowel Syndrome (IBS) was performed by gastroenterologist. Healthy individuals were selected from general population and had no history of gastroenterological problem.

After describing the procedures and purposes of the study, written informed consents were obtained. Each subject was asked to complete the Persian version of Cattle anxiety scale, World Health Organization Quality of Life (WHOQOL) and Irritable Bowel Syndrome (IBS) symptom checklist as well as a questionnaire to obtain demographic data about gender, age, level of education and marital status.

Instruments

Cattell's anxiety scale: The instrument consists of 40 items. The items are divided into 2 sub-scales including state anxiety and trait anxiety. The range of the test-retest reliability coefficient is reported to be from 0.65-0.75, with a 2 month interval between tests. Construct and concurrent validity of the scale has also been performed and supports its validity (Spielberger, 1989; Spielberger *et al.*, 1983). Cattle anxiety scale has been shown to be reliable and valid in an Iranian population (Salarifar and Etemad, 2011).

World Health Organization Quality of Life (WHOQOL):

This questionnaire consists of 36 items. It measures eight health subscales including physical functioning, role

limitations because of physical problems, bodily pain, general health perception, vitality, social functioning, role limitations because of emotional problems and mental health.

Physical functioning, role limitations because of physical problems, bodily pain, general health perception grouped in to physical component and vitality, social functioning, role limitations because of emotional problems and mental health are subscales of mental component. Each subscale graded from 0-100. Higher scores are considered better status (Stull *et al.*, 2014).

This scale has been used in many researches in Iran and the questionnaire validity and reliability has been confirmed (Montazeri *et al.*, 2005).

Rome III diagnostic questionnaire: It is a self-report questionnaire that has been developed for the diagnosis of functional gastrointestinal disorders (Wang *et al.*, 2012). According to Rome III diagnostic criteria, IBS is characterized by abdominal pain or discomfort in at least 3 of the previous 6 months, with 2 or more of the following items:

- Pain or discomfort improved after defecation
- Pain or discomfort associated with a change in frequency of stool
- Pain or discomfort associated with a change in the form of stool (Dong *et al.*, 2013)

Statistical tests: Data was analyzed using multivariate analyses of variance (MANOVA), Chi-square test and Fisher exact test. The probability level of 0.05 was accepted as statistically significant. Statistical analyses were carried out using SPSS version 16.

RESULTS

Demographic characteristics (sex, residence and level of education) of both patients and healthy individuals are detailed in Table 1.

The anxiety and quality of life of patients and healthy individuals were compared using multiple univariate

Table 3: Prevalence of IBS based on demographic features

analyses of variance (ANOVAs). There were significant differences between patients and healthy individuals in anxiety (F = 287.48, p<0.001) and quality of life (F = 151.7, p = 0.002). It means patients with IBS suffering from higher level of anxiety and have lower level of quality of life compared to healthy individuals. A comparison of two variables is shown in Table 2.

Distributions of IBS according to demographic variables are listed in Table 3. The IBS was found in 65% (39 of 60) of female and 35% (21 of 60) of male ($\chi^2 = 5.57$, p = 0.006). Twenty four percent of patients had grade school level of education, 35% had 12 years/high school education and 21.6% of patients had university degree ($\chi^2 = 19$, p = 0.53). Forty five percent of patients were living in urban areas and 25% of them were from rural areas ($\chi^2 = 7.68$, p = 0.001). Hence, there are significant differences between patients and healthy individuals with different gender and education subgroups in prevalence of IBS.

Table 1: Demographic characteristics of the patient and control group	Table 1	: Demographic	characteristics (of the	patient :	and control	groups
---	---------	---------------	-------------------	--------	-----------	-------------	--------

	Frequency					
	Patients	with IBS	Healthy individuals			
Demographic variables	 No.	%	No.	%		
Gender						
Female	39	65	33	55		
Male	21	35	27	45		
Education						
Grade school	24	40	17	28.33		
12 years/high school	21	35	27	45		
University degree	13	21.6	16	26.66		
Residence						
Urban	45	75	52	86.66		
Rural	15	25	8	13.33		

IBS: Irritable bowel syndrome

Table 2: Comparison using multivariate analysis of variance (MANOVA) of anxiety and quality of life of patients and healthy individuals

	Patients	5	Healthy	individual	s	
					-	
Variables	Mean	SD	Mean	SD	$F_{1,118}$	p-value
Anxiety	50.25	±7.60	26.76	±7.82	287.48	< 0.001
Quality of life	57.46	± 8.12	37.68	± 7.95	151.7	0.002

	Frequency						
	Patients with IBS		Healthy individuals				
Demographic variables	 No.	%	No.	%	χ^2	df	p-value
Gender							
Male	39	65	33	55.0	5.57	1	0.006
Female	21	35	27	45.0			
Education							
Grade school	24	40	17	28.33	19.0	2	0.53
12 years/high school	21	35	27	45.0			
University degree	13	21.6	16	26.66			
Residence							
Urban	45	75	52	86.66	7.68	1	0.001
Rural	15	25	8	13.33			

IBS: Irritable bowel syndrome, df: Degree of freedom

DISCUSSION

The present study aimed to compare anxiety and quality of life in patients with Irritable Bowel Syndrome (IBS) and healthy individuals. Our findings showed that patients with IBS suffered from higher level of anxiety and poorer quality of life compared to healthy individuals.

The findings replicate the results of studies conducted by Lee *et al.* (2009), Mykletun *et al.* (2010), Porcelli *et al.* (2014) and Yi *et al.* (2014) that showed IBS was significantly related to anxiety. Our finding also is congruent with result of Lackner *et al.* (2013), who reported anxiety sensitivity and perceived stress as some of predictors of fatigue in patients with IBS (Lackner *et al.*, 2013).

These findings indicate that psychological factors, including anxiety, may collaborate to physical factors to trigger or intensify functional gastrointestinal disorders. Enteric nervous system is very sensitive to the emotional states so that negative emotions such as stress, change in bowel movement activities and can cause bowel symptoms such as IBS (Hauser *et al.*, 2014). Patients with IBS have more often reported themselves as experiencing stressful and major life events before the onset of illness (Surdea-Blaga *et al.*, 2012).

The relationship between psychosocial factors and symptom onset might also be explained, in part, by specific biological mechanisms. The main stress-response axis of the body i.e., hypothalamic-pituitary-adrenal axis has been reported to be related to chronic widespread pain. This axis may also be involved in functional gastrointestinal disorders including IBS (Nicholl *et al.*, 2008).

It is also suggested that excess of corticotropin releasing hormone may be related to both anxiety and IBS-like symptoms (Lee *et al.*, 2009). Other comments stated that psychological problems in patients with IBS may be reaction to stress of the chronic illness (Whitehead *et al.*, 1988). But lower rate of mental disorders in patients with other gastrointestinal diseases such as inflammatory bowel disease, rejected the idea (Walker *et al.*, 1995).

Some studies have reported that psychological distress is unrelated to irritable bowel syndrome but influence on medical care-seeking (Whitehead *et al.*, 1988). But some investigators didn't find difference in mental health status between healthcare seekers and healthcare non-seekers (Mykletun *et al.*, 2010; Tang *et al.*, 2008) and even community sample with IBS symptoms report stress and poorer mental health (Gralnek *et al.*, 2000).

However, the study of young adults from New Zealand reported that IBS was not significantly associated with psychiatric disorder including anxiety disorders (Talley *et al.*, 2001). This gap might be somewhat due to features of subjects, methodological differences in assessment of psychological factors or cultural differences in somatic presentations of psychological problem. In the present study we found that patients with IBS suffered from poor quality of life compared to healthy individuals. Similarly Creed *et al.* (2001) reported that, both abdominal pain and psychological symptoms were associated with impaired health-related quality of life in patients with severe IBS. And results of the study conducted by DiBonaventura *et al.* (2011) indicated that constipation predominant IBS was related to poorer HRQOL, less work productivity and high direct health care costs. Li *et al.* (2003) also found that people with IBS experience substantial impairment in health related quality of life. Persistent abdominal pain, other physical problems, fatigue (Monnikes, 2011) and psychological problems (Tkalcic *et al.*, 2014) that are often associated with IBS cause impairment in patient's quality of life.

For instance Lackner *et al.* (2013) found that generalized anxiety, agoraphobia, tension headache, insomnia, depression and back pain were related to greater illness and poorer quality of life.

With respect to gender, our results showed that IBS were significantly more prevalent in women than in men. We replicate the findings of the study conducted by Canavan *et al.* (2014). By contrast Lee *et al.* (2009) didn't find significant different between the genders in the prevalence of IBS (Lee *et al.*, 2009).

Williams *et al* (2006) showed that doctors are more likely to diagnose IBS in women than in men, even when men meet criteria for IBS (Tang *et al.*, 2008). And several studies (Williams *et al.*, 2006) reported that the Rome criteria are more sensitive in diagnosing IBS women than in men (Tang *et al.*, 2008).

We also found more prevalence of IBS in patients from urban areas compared to those of from rural areas. This maybe because complications and multiple stressors in urban areas make people prepared to different dieses. Consistent with Mykletun *et al.* (2010), no difference was found in prevalence of IBS infection between patients with different Levels of education.

In conclusion, patients with IBS are more likely to suffer from anxiety and burdens of illness on quality of life. Our results have implications for clinicians who dealing with patients with IBS. And support the claim of the need for psychological interventions in management of patients with IBS.

The strength of the present study is that diagnosis of IBS was made by gastroenterologist in addition to completing Rome III questionnaire. While in some previous studies the diagnosis was based on the participant's report of their condition. Some limitations should, however, be taken into account. The participants selected from referral centers, which means the findings should be generalized to patients who are not in care medical centers, with caution. Moreover, the study was cross sectional and could not demonstrate the causal relationship between variables.

REFERENCES

- Canavan, C., J. West and T. Card, 2014. The epidemiology of irritable bowel syndrome. Clin. Epidemiol., 6: 71-80.
- Creed, F., J. Ratcliffe, L. Fernandez, B. Tomenson and S. Palmer *et al.*, 2001. Health-related quality of life and health care costs in severe, refractory irritable bowel syndrome. Ann. Intern Med., 134: 860-868.
- Dainty, A.D., M. Fox, N. Lewis, M. Hunt and E. Holtham *et al.*, 2014. A mixed methods feasibility study to evaluate the use of a low-intensity, nurse-delivered cognitive behavioural therapy for the treatment of irritable bowel syndrome. Br. Med. J. Open, Vol. 4. 10.1136/bmjopen-2014-005262.
- DiBonaventura, M., S.X. Sun, S.C. Bolge, J.S. Wagner and R. Mody, 2011. Health-related quality of life, work productivity and health care resource use associated with constipation predominant irritable bowel syndrome. Curr. Med. Res. Opin., 27: 2213-2222.
- Dong, Y.Y., F.X. Chen, Y.B. Yu, C. Du, Q.Q. Qi, H. Liu and Y.Q. Li, 2013. A school-based study with Rome III criteria on the prevalence of functional gastrointestinal disorders in Chinese college and university students. PloS One, Vol. 8. 10.1371/journal.pone.0054183
- El-Serag, H.B., K. Olden and D. Bjorkman, 2002. Health-related quality of life among persons with irritable bowel syndrome: A systematic review. Alimentary Pharmacol. Therapeut., 16: 1171-1185.
- Folks, D.G., 2004. The interface of psychiatry and irritable bowel syndrome. Curr. Psychiatry Rep., 6: 210-215.
- Fossey, M.D. and R.B. Lydiard, 1990. Anxiety and the gastrointestinal system. Psychiatr. Med., 8: 175-186.
- Ghoshal, U.C. and D. Srivastava, 2014. Irritable bowel syndrome and small intestinal bacterial overgrowth: Meaningful association or unnecessary hype. World J. Gastroenterol., 20: 2482-2491.
- Gralnek, I.M., R.D. Hays, A. Kilbourne, B. Naliboff and E.A. Mayer, 2000. The impact of irritable bowel syndrome on health-related quality of life. Gastroenterology, 119: 654-660.
- Gulewitsch, M.D., P. Enck, J. Schwille-Kiuntke, K. Weimer and A.A. Schlarb, 2013. Mental strain and chronic stress among university students with symptoms of irritable bowel syndrome. Gastroenterol. Res. Pract. 10.1155/ 2013/206574
- Hauser, G., S. Pletikosic and M. Tkalcic, 2014. Cognitive behavioral approach to understanding irritable bowel syndrome. World J. Gastroenterol., 20: 6744-6758.
- Irwin, C., S.A. Falsetti, R.B. Lydiard, J.C. Ballenger, C.D. Brock and W. Brener, 1996. Comorbidity of posttraumatic stress disorder and irritable bowel syndrome. J. Clin. Psychiatry, 57: 576-578.
- Khanna, S. and P.K. Tosh, 2014. A clinician's primer on the role of the microbiome in human health and disease. Mayo Clin. Proc., 89: 107-114.

- Lackner, J.M., C.X. Ma, L. Keefer, D.M. Brenner and G.D. Gudleski *et al.*, 2013. Type, rather than number, of mental and physical comorbidities increases the severity of symptoms in patients with irritable bowel syndrome. Clin. Gastroenterol. Hepatol., 11: 1147-1157.
- Lee, S., J. Wu, Y.L. Ma, A. Tsang, W.J. Guo and J. Sung, 2009. Irritable bowel syndrome is strongly associated with generalized anxiety disorder: A community study. Alimentary Pharmacol. Therapeut., 30: 643-651.
- Levy, R.L., M. von Korff, W.E. Whitehead, P. Stang and K. Saunders *et al.*, 2001. Costs of care for irritable bowel syndrome patients in a health maintenance organization Costs of Care for IBD Patients in an HMO. Am. J. Gastroenterol., 96: 3122-3129.
- Li, F.X., S.B. Patten, R.J. Hilsden and L.R. Sutherland, 2003. Irritable bowel syndrome and health-related quality of life: A population-based study in Calgary, Alberta. Can. J. Gastroenterol., 17: 259-263.
- Lin, H.C., 2004. Small intestinal bacterial overgrowth: A framework for understanding irritable bowel syndrome. J. Am. Med. Assoc., 292: 852-858.
- Liss, J.L., D. Alpers and R.A. Woodruff, 1973. The irritable colon syndrome and psychiatric illness. Dis. Nervous Syst., 34: 151-157.
- Lorenzo-Zuniga, V., E. Llop, C. Suarez, B. Alvarez, L. Abreu, J. Espadaler and J. Serra, 2014. I.31, a new combination of probiotics, improves irritable bowel syndrome-related quality of life. World J. Gastroenterol., 20: 8709-8716.
- Lydiard, R.B., 2001. Irritable bowel syndrome, anxiety and depression: What are the links? J. Clin. Psychiatry, 62: 38-45.
- Mahvi-Shirazi, M., E. Ahmadi and M.S. Ali, 2013. Mental health levels in IBS, IBD and healthy people: A comparative study. J. Sci. Today's World, 2: 549-558.
- Monnikes, H., 2011. Quality of life in patients with irritable bowel syndrome. J. Clin. Gastroenterol., 45: 98-101.
- Montazeri, A., A. Goshtasebi, M. Vahdaninia and B. Gandek, 2005. The short form health survey (SF-36): Translation and validation study of the Iranian version. Qual. Life Res., 14: 875-880.
- Muscatello, M.R.A., A. Bruno, G. Scimeca, G. Pandolfo and R.A. Zoccali, 2014. Role of negative affects in pathophysiology and clinical expression of irritable bowel syndrome. World J. Gastroenterol., 20: 7570-7586.
- Mykletun, A., F. Jacka, L. Williams, J. Pasco and M. Henry *et al.*, 2010. Prevalence of mood and anxiety disorder in self reported Irritable Bowel Syndrome (IBS). An epidemiological population based study of women. BMC Gastroenterol., Vol. 10. 10.1186/1471-230X-10-88
- Nicholl, B.I., S.L. Halder, G.J. Macfarlane, D.G. Thompson, S. O'Brien, M. Musleh and J. McBeth, 2008. Psychosocial risk markers for new onset irritable bowel syndrome-results of a large prospective population-based study. J. Pain, 137: 147-155.

- Niesten, I.J.M., E. Karan, F.R. Frankenburg, G.M. Fitzmaurice and M.C. Zanarini, 2014. Prevalence and risk factors for irritable bowel syndrome in recovered and non-recovered borderline patients over 10 years of prospective follow-up. Personality Mental Health, 8: 14-23.
- Nyrop, K.A., O.S. Palsson, R.L. Levy, M. von Korff, A.D. Feld, M.J. Turner and W.E. Whitehead, 2007. Costs of health care for irritable bowel syndrome, chronic constipation, functional diarrhoea and functional abdominal pain. Alimentary Pharmacol. Therapeut., 26: 237-248.
- Ohman, L. and M. Simren, 2010. Pathogenesis of IBS: Role of inflammation, immunity and neuroimmune interactions. Nature Rev. Gastroenterol. Hepatol., 7: 163-173.
- Osterberg, E., L. Blomquist, I. Krakau, R.M. Weinryb, M. Asberg and R. Hultcrantz, 2000. A population study on irritable bowel syndrome and mental health. Scand J. Gastroenterol., 35: 264-268.
- Paras, M.L., M.H. Murad, L.P. Chen, E.N. Goranson and A.L. Sattler *et al.*, 2009. Sexual abuse and lifetime diagnosis of somatic disorders. A systematic review and meta-analysis. J. Am. Med. Assoc., 302: 550-561.
- Porcelli, P., M. de Carn and G. Leandro, 2014. Alexithymia and gastrointestinal-specific anxiety in moderate to severe irritable bowel syndrome. Comprehensive Psychiatry, 55: 1647-1653.
- Roberts, L.M., D. McCahon, R. Holder, S. Wilson and F.D.R. Hobbs, 2013. A randomised controlled trial of a probiotic functional food in the management of irritable bowel syndrome. BMC Gastroenterol., Vol. 13. 10.1186/ 1471-230X-13-45
- Salarifar M. and H.P. Etemad, 2011. Relationship between metacognitive beliefs and depression and anxiety. J. Yafteh, 13: 29-38.
- Spielberger, C.D., R.L. Gorsuch, R. Lushene, P.R. Vagg and G.A. Jacobs, 1983. Manual for the State-Trait Anxiety Inventory. Consulting Psychologists Press, Palo Alto, CA.
- Spielberger, C.D., 1989. State-Trait Anxiety Inventory: A Comprehensive Bibliography. 2nd Edn., Consulting Psychologists Press, Palo Alto, CA.
- Stull, D.E., R. Wasiak, N. Kreif, M. Raluy, A. Colligs, C. Seitz and C. Gerlinger, 2014. Validation of the SF-36 in patients with endometriosis. Q. Life Res., 23: 103-117.
- Surdea-Blaga, T., A. Baban and D.L. Dumitrascu, 2012. Psychosocial determinants of irritable bowel syndrome. World J. Gastroenterol., 18: 616-626.
- Talley, N.J., S. Howell and R. Poulton, 2001. The irritable bowel syndrome and psychiatric disorders in the community: Is there a link? Am. J. Gastroenterol., 96: 1072-1079.

- Tang, L.Y.L., A. Nabalamba, L.A. Graff and C.N. Bernstein, 2008. A comparison of self-perceived health status in inflammatory bowel disease and irritable bowel syndrome patients from a Canadian national population survey. Can. J. Gastroenterol., 22: 475-483.
- Tkalcic, M., S. Pletikosic and G. Hauser, 2014. Biological and psychological determinants of health related quality of life in irritable bowel syndrome. Proceedings of the 2nd Annual Conference of the European Association of Psychosomatic Medicine, June 25-28, 2014, Sibiu, Romania, pp: 516-516.
- Van Tilburg, M.A.L., D.K. Runyan, A.J. Zolotor, J.C. Graham and H. Dubowitz *et al.*, 2010. Unexplained gastrointestinal symptoms after abuse in a prospective study of children at risk for abuse and neglect. Ann. Family Med., 8: 134-140.
- Walker, E.A., A.N. Gelfand, M.D. Gelfand and W.J. Katon, 1995. Psychiatric diagnoses, sexual and physical victimization and disability in patients with irritable bowel syndrome or inflammatory bowel disease. Psychol. Med., 25: 1259-1267.
- Wang, Y.T., H.Y. Lim, D. Tai, T.L. Krishnamoorthy, T. Tan, S. Barbier and J. Thumboo, 2012. The impact of Irritable bowel syndrome on health-related quality of life: A Singapore perspective. BMC Gastroenterol., Vol. 12. 10.1186/1471-230X-12-104
- Whitehead, W.E., L. Bosmajian, A.B. Zonderman, P.T. Jr. Costa and M.M. Schuster, 1988. Symptoms of psychologic distress associated with irritable bowel syndrome. Comparison of community and medical clinic samples. Gastroenterology, 95: 709-714.
- Williams, R.E., C.L. Black, H.Y. Kim, E.B. Andrews, A.W. Mangel, J.J. Buda and S.F. Cook, 2006. Determinants of healthcare-seeking behaviour among subjects with irritable bowel syndrome. Alimentary Pharmacol. Therapeut., 23: 1667-1675.
- Yacob, D., C. Di Lorenzo, J.A. Bridge, P.F. Rosenstein, M. Onorato, T. Bravender and J.V. Campo, 2013. Prevalence of pain-predominant functional gastrointestinal disorders and somatic symptoms in patients with anxiety or depressive disorders. J. Pediatr., 163: 767-770.
- Yi, Z.H., Z.B. Yang, L. Kang, L. Feng and L. Yang, 2014. [Clinical features, quality of life and psychological health of patients with irritable bowel syndrome and functional dyspepsia]. Sichuan Da Xue Xue Bao, 45: 493-496.