World Cancer Research Journal wcrj 2017; 4 (2): e880

# UNDERSTANDING THE LINK BETWEEN DEPRESSION AND PAIN PERCEPTION IN IRANIAN CANCER PATIENTS



A.H. GOUDARZIAN<sup>1</sup>, A. JAFARI<sup>2</sup>, M. BAGHERI-NESAMI<sup>3</sup>, F. ZAMANI<sup>2</sup>

<sup>1</sup>BSc of Nursing, Student Research Committee, Mazandaran University of Medical Sciences, Sari, Iran <sup>2</sup>Nursing Student, Student Research Committee, Mazandaran University of Medical Sciences, Sari, Iran <sup>3</sup>Associate Professor, Infectious Diseases Research Center with Focus on Nosocomial Infection, Mazandaran University of Medical Sciences, Sari, Iran

**Abstract – Objective:** This study aimed to investigate the relationship between depression and pain perception in cancer patients.

Patients and Methods: In this cross-sectional study (from October to December 2015) 380 cancer patients were admitted to one of our hospitals associated with the Mazandaran University of Medical Sciences (Sari, Iran), and were entered to the study using accessible sampling. Data was collected by demographic questionnaire, depression scale of Center for Epidemiological studies and McGill pain questionnaire. The statistical package for social sciences, version 20.0 (SPSS Inc., Chicago, IL, USA) was utilized for data analysis by descriptive and inferential statistic tests, including Spearman's correlation and Generalized Linear models.

**Results:** Males (48.39±13±39; CI 95: 46.41-50.38) were older than females (45.33±18.44; CI 95: 42.79-47.87). There is a positive and significant correlation between pain perception and depression and there is a negative and significant correlation between depression and stages of cancer (p<0.05).

**Conclusions:** Considering the adverse effects of depression on the treatment process and secondary problems, more comprehensive studies should be conducted about the effects of depression on pain perception in order to take effective steps towards intervention and health promotion of these patients.

**KEYWORDS:** Depression, Cancer, Pain perception, Iran, Cross-sectional study.

# **INTRODUCTION**

Cancer can reduce a person's ability to play his/her role, leading to a feeling of lack of competence, thus reducing the individual's self-esteem and, eventually, causing psychological reactions such as depression, anxiety, and stress<sup>1</sup>. Several studies have acknowledged that depression is a common psychiatric disorder in cancer patients<sup>2-4</sup>. Depression is very detrimental to cancer patients because it reduces the individual's resistance against the disease<sup>5</sup>. The prevalence of advanced depression in cancer patients has been reported to be 1.5% to 30%<sup>6,7</sup>. Furthermore, can-

cer, as the most common chronic disease, is often accompanied with severe pain<sup>8</sup>. Cancer pain can have various causes, including tumor growth and spread, side effects of therapies such as chemotherapy, radiotherapy, surgery, and underlying diseases<sup>9</sup>. The prevalence of pain varies from 5% in patients with leukemia to 85% in patients with primary bone tumors<sup>10</sup>. There are several ways to treat and relieve the pain. One of the more popular methods involves the prescription of analgesics, but these drugs often have considerable side effects on patients, either physical or mental<sup>11</sup>. Besides conventional pharmacological methods of pain relief, a number of behavioral techniques

# World Cancer Research Journal

have also attracted a degree of academic scrutiny of late<sup>12,13</sup>, but researchers need to find and explain the various ways of pain relief in this group of patients<sup>10</sup>. Studies suggest a link between pain and depression in cancer patients. Pain and depression interact each other and aggravate the conditions<sup>14</sup>. The mechanism of the relation of pain and depression has not been clearly stated<sup>15,16</sup>, but a study on the possible mechanisms of this relationship speculates on the sharing of the neural pathways and behavioral cycle change<sup>17</sup>. The results show that an increase in depression impairs part of the neural circuitry in the brain that regulates emotions, thus intensifying the perception of pain. In other words, the brain processes pain more accurately when a person is in a depressed mood, thus making pain unbearable<sup>10</sup>. Despite the high prevalence of cancer in Iran<sup>16</sup> and the need to investigate the complications associated with it, no studies have been conducted until now on the relationship between depression and pain in cancer patients. Therefore, the present study aimed to investigate the relationship between depression and pain perception in cancer patients.

## **PATIENTS AND METHODS**

In this cross-sectional study conducted in 2015 (from October to December), 380 cancer patients were admitted to one of our hospitals associated with the Mazandaran University of Medical Sciences (Sari, Iran) and were recruited randomly. In four months, about 600 patients were admitted to the Oncology Unit of one of these hospitals; about 430 patients met the inclusion criteria (participation rate was 63.3%). Inclusion criteria included an age range of 18 years and above, cancer treatment with radiation, chemotherapy or surgery, and the ability to read and write Persian. The exclusion criteria included taking antidepressants in the last six months, the transfer of patients to other hospitals, and the occurrence of acute medical conditions (such as loss of consciousness).

## Ethical Issue

After explaining the purpose of the study and how to complete the questionnaire, an informed consent form was signed by the eligible patients. Then, they were given the details of the study objectives and the questionnaires were distributed. If a question was vague, explanations were offered to the patients to make things clear. It should be noted that these explanations were given only in order to avoid ambiguity and were without bias.

## Data Collection tools

Data was collected with the help of a socio-demographic questionnaire, the questionnaire of the Center for Epidemiologic Studies Depression Scale (CES-D) and McGill Pain Questionnaire (MPQ). The socio-demographic questionnaire sought data on age, sex, education level, economic status, history of drug use, family history of cancer, and the stage of cancer. CES-D is a 20-item tool used to assess symptoms associated with depression experienced in the past week<sup>18</sup>. Each of the 20 items available in this tool was given a score as per the Likert scale: 0 = rarely or never (less than one day), 1 = occasionally or in few cases (1 to 2 days), 2 =occasionally or a moderate amount of time (3 to 4 days), and 3 = most of the time or all the time (5 to 7 days)<sup>14</sup>. The total scores ranged from 0 to 60. Knight et al14 have reported the reliability of this tool to be 0.88 using Cronbach's alpha. In this study, the reliability of the instrument in assessing Iranian cancer patients was calculated to be 0.741 by using Cronbach's alpha. MPQ includes 78 descriptive sentences out of 20 collections9. The patients were asked to select the most fitting description of their pain with just one word from each group. If all options in a group failed to provide an appropriate description of the pain, the patient was not free to choose an option from that group. When patients chose more than one option in each group, the highest rating (maximum pain) was chosen for the final analysis<sup>19</sup>. The score of each group was collected separately to calculate the final score. The MPQ total score is described as the "pain rating index based on the scores of the words," leading to a sum of the scores of each group<sup>20</sup>. The score range varies from 0 (when no word is selected) to 78 (maximum pain when selected in each group)<sup>19</sup>. Dworkin et al<sup>21</sup> calculated the reliability of this tool to be 0.77 using Cronbach's alpha. In the present study, the reliability of this tool was calculated to be 0.94 using Cronbach's alpha.

# Statistical Analysis

The statistical package for social sciences, version 20.0 (SPSS Inc., Chicago, IL, USA), was used for data analysis. First, descriptive statistics for the continuous variables were shown as means with standard deviation (SD) and n (%) for the categorical variables. Spearman's correlations were used to probe the relationship between depression and pain perception. Finally, the predictors associating with pain perception were determined using the generalized linear models (GLM). Statistical significance was set at p < 0.05.

## **RESULTS**

The demographic characteristics of 380 cancer patients are summarized in Table I. Men's age (48.39±13.93; CI 95%: 46.41-50.38) was higher than the age of women (45.33±18.44; CI 95%: 42.79-47.87). About 69.7% of cases were at an early stage of the disease (Stage I-II) and 31.3% of them were in an advanced stage of cancer (Stage III-IV). It was found that 29.5% of them had a family history of cancer, 38.7% were with a history of a particular disease, and 18.7% had a history of drug use. The mean score of depression was (24.14±5.45; CI 95%: 23.59-24.69) in cancer patients. Moreover, the perception of pain was equal to (21.61+13.55; CI 95%: 20.36-22.17). The results of the Spearman correlation analysis to determine the association between depression and pain in cancer patients, showed that there was a significant positive correlation between the perception of pain and depression (r = 0.106; p<0.05), and this correlation was negative with respect to the stage of cancer (r = -0.337; p < 0.05). According to the results of the GLM, there was a significant relationship between depression and the four stages of cancer in cancer patients (B = 0.046; p = 0.032). Also, according to Table II, there was a significant relationship between depression and level of education (p<0.05), average economic status (B = 1.6; p = 0.027), age (B = -0.048; p = 0.046), sex (B = 2.44; p < 0.001), marital status (B = 3.8; p<0.001) and history of drug use (B = -2.36; p = 0.001) in cancer patients.

#### **DISCUSSION**

This study aimed to investigate the relationship between depression and pain perception in cancer patients. Based on our findings, the average score of cancer in patients was from moderate to high, a feature that has been reported in many studies<sup>22-24</sup>. Cancer can reduce a person's ability to play his/her role and create a feeling of lack of competence, thus reducing a person's self-esteem and, eventually, causing psychological reactions such as depression, anxiety and stress<sup>25</sup>. Also, the perception of pain in the present study was from moderate to low in cancer patients. These findings were also consistent with the findings of several studies<sup>26-30</sup>. It should be noted that the perception of pain, in the context of the cancer type and the stage of the disease, was different in various studies 31. In most cases, factors such as lack of knowledge or fear of strong analgesics and their side effects, prevented the effective treatment pain, resulting the accentuation of the pain perception<sup>32</sup>. One of the most important results of the present study was finding a signifi-

**TABLE 1.** Sample characteristics of cancer patients in the study.

study.				
Characteristic	N (%)			
Sex				
– Male	175 (46.1) 205 (53.9)			
– Female				
Economic situation	, ,			
– Weak	110 (28.9) 204 (53.7) 66 (17.4)			
- Average				
- Good				
Education	,			
– Illiterate	210 (55.3)			
– Diploma	138 (36.3)			
-BS	22 (5.8) 10 (2.6)			
– MSc and above				
Material	- ( )			
- Single	51 (13.4) 329 (86.6)			
– Married				
Cancer stage	( )			
- One - Two - Tree	132 (34.7)			
	133 (35)			
	92 (24.2)			
– Four	23 (6.1)			
Family history of cancer				
Family history of cancer  - Yes	112 (29.5) 268 (70.5)			
- No				
Depression	, ,			
– Down	261 (68.7) 119 (31.3)			
– Up				
History of drug use	,			
- Yes	71 (18.7)			
- No	309 (81.3)			
Past medical history	, ,			
- Yes	147 (38.7)			
- No	233 (61.3)			
Characteristic	Mean (SD)			
Age	46.74 (16.328)			
Pain				
Depression	21.61 (13.55) 24.14 (5.45)			
_	` '			

cant positive relationship between depression and pain perception in cancer patients. These results are consistent with the results of Farnam et al<sup>33</sup>, Mollazade et al<sup>34</sup>, Kroenk et al<sup>35</sup>, Sullivan-Singh et al<sup>36</sup> and Mirghaforvand et al<sup>37</sup>. Among the possible mechanisms associated with this relationship, one can refer to Shared neural pathways and change of behavioral cycle<sup>17</sup>. The results show that an increase in depression impairs a part of the neural circuit in the brain that regulates emotions, intensifying the pain perception. In other words, the brain processes pain more accurately while a person is in a depressed mood and, as result, the perception of pain can become more acute<sup>10</sup>. In the present study, there was a significant difference between depression and the level of education, economic status, age, sex, marital status and a history of drug use in cancer patients, a trend consistent with other stud-



# World Cancer Research Journal

**TABLE 2.** Relationship between depression and its covariates in Iranian cancer patients.

Variable	В	SE	95%	CI	р	
Education						
– Illiterate	7.486	1.5964	4.357	10.615	$0.000^{\rm b}$	
– Diploma	6.715	1.5244	3.727	9.703	$0.000^{\rm b}$	
– BŜ	3.324	1.6747	0.042	6.607	$0.047^{b}$	
<ul> <li>MSc and upper</li> </ul>	$0^{a}$	_	_	_	_	
Economic situation						
– Weak	1.390	148	-0.148	2.927	0.077	
– Average	1.609	-3.032	-3.032	-0.186	$0.027^{b}$	
- Good	O <sup>a</sup>	_	_	_	_	
Cancer stage						
– One	10.934	8.590	8.590	13.277	$0.000^{\rm b}$	
– Two	10.706	8.593	8.593	12.818	$0.000^{\rm b}$	
- Tree	8.796	6.425	6.425	11.168	$0.000^{\rm b}$	
– Four	$0^{a}$	_	_	_	_	
Age	-0.048	-0.095	-0.095	-0.001	$0.046^{b}$	
Sex	2.442	1.241	1.241	3.643	$0.000^{\rm b}$	
Marital	3.795	2.032	2.032	5.558	$0.000^{\rm b}$	
History of drug use	-2.366	-3.808	-3.808	-0.923	$0.001^{b}$	
Family history of cancer	0.395	-0.852	-0.852	1.641	0.535	
PMH**	0.880	-0.167	-0.167	1.927	0.100	
Pain	0.046	0.004	0.004	0.087	$0.032^{b}$	

<sup>&</sup>lt;sup>a</sup> Set to zero because this parameter is redundant; <sup>b</sup> statistically significant at  $p \le 0.05$ . \*\*Past medical history.

ies<sup>38-40</sup>. Although the results of the study by Farnam et al<sup>33</sup> showed a positive and significant relationship between gender, occupation, depression and pain, they did not report a relationship between age, intensity of pain, and depression. Rayner et al<sup>41</sup> study showed there was no difference between the age of depressed and non-depressed groups. Several possible causes of this paradox can be the type of pain, the sample size, and the instruments used to determine depression.

## Limitations

The most important limitation of this study was a lack of access to patients in other hospitals in the Mazandaran Province and Country. So, the small sample size makes it difficult to generalize the results correctly. The other limiting factor was cultural differences of the patients, a variable that was not controllable in this study. Impatience and imprecision of some of the patients in the completion of the questionnaire due to disease-related treatments could affect the results. It is suggested that, because of the importance of the issue, similar studies should be conducted more frequently in the future with a bigger sample size.

# Application of Results

Depression, as a consequence of cancer, must be one of the important nursing diagnoses in healthcare centers. Given the prevalence of depression in these patients, holding psychotherapy sessions for early diagnosis of depression and, if necessary, therapy and use of antidepressants, must be started. Moreover, considering the significant effect of depression in the perception of pain, it is expected that by controlling depression in these patients, the pain tolerance threshold can be significantly controlled.

# **CONCLUSIONS**

According to the results, a significant positive correlation was found between depression and the perception of pain in cancer patients. Also, there was a significant relationship between depression and level of education, average economic status, age, sex, marital status, and history of drug use in cancer patients. Regarding the adverse effects of depression and pain on the treatment process and secondary problems, more comprehensive studies must be done on the effects of depression on pain control in these patients, to take effective steps to promote the health of these patients.

## **ACKNOWLEDGEMENT:**

Nurses, doctors, patients and all those who helped in carrying out this study are sincerely appreciated. Special thanks also goes out to student Research Committee of the Mazandaran University of Medical Sciences for their financial support.

## AUTHOR'S ROLE:

AH. G contributed to the study design, data collection, and the writing of the draft of the manuscript, A. J contributed to data collection, and the writing and revision of the draft manuscript, M. BN contributed to data analysis and the revision of the manuscript, and F. Z contributed to data collection and the writing of the draft of manuscript. All the authors approved the final version of the manuscript.

## CONFLICT OF INTEREST:

There is no conflict of interest in the design of this study and the report of results.

## **REFERENCES**

- 1. Sharif Nia H, Pahlevan Sharif S, Lehto RH, Boyle C, Yaghoobzadeh A, Kaveh O, Goudarzian AH. Development and psychometric evaluation of a Persian version of the death depression scale-revised: a cross-cultural adaptation for patients with advanced cancer. Jpn J Clin Oncol 2017: 1-7.
- 2. Burgess C, Cornelius V, Love S, Graham J, Richards M, Ramirez A. Depression and anxiety in women with early breast cancer: five year observational cohort study. BMJ 2005; 330: 702.
- 3. Bussing A, Fischer J, Ostermann T, Matthiessen PF. Reliance on God's help, depression and fatigue in female cancer patients. Int J Psychiatry Med 2008; 38: 357-372.
- 4. So WK, MARSH G, LING WM, LEUNG FY, LO JC, YEUNG M, LI GK. The symptom cluster of fatigue, pain, anxiety, and depression and the effect on the quality of life of women receiving treatment for breast cancer: a multicenter study. Oncol Nurs Forum 2009; 36: 205-214.
- 5. Sharif Nia H, Sharif SP, Esmaeili R, Goudarzian AH, Tahmasbi B, Yaghoobzadeh A, O. Factors influencing the level of death depression in patients with cancer: a path analysis. Journal of Mazandaran University of Medical Sciences 2017; 26: 318-331.
- 6. Currier MB, Nemeroff CB. Depression as a risk factor for cancer: from pathophysiological advances to treatment implications. Annu Rev Med 2014; 65: 203-221.
- GOUDARZIAN AH, NESAMI MB, ZAMANI F, NASIRI A, BEIK S. Relationship between depression and self-care in Iranian patients with cancer. Asian Pac J Cancer Prev 2017; 18: 101.
- 8. Rose L, SMITH O, GÉLINAS C, HASLAM L, DALE C, LUK E, BURRY L, McGILLION M, MEHTA S, WATT-WATSON J. Critical care nurses' pain assessment and management practices: a survey in Canada. Am J Crit Care 2012; 21: 251-259.
- 9. Gauthier LR, Young A, Dworkin RH, Rodin G, Zimmermann C, Warr D, Librach SL, Moore M, Shepherd FA, Pillai Riddell R, Macpherson A, Melzack R, Gagliese L. validation of the short-form mcgill pain questionnaire-2 in younger and older people with cancer pain. J Pain 2014; 15: 756-770.
- GOESLING J, CLAUW DJ, HASSETT AL. Pain and depression: an integrative review of neurobiological and psychological factors. Curr Psychiatry Rep 2013; 15: 1-8.

- SHARIF NIA H, PAHLEVAN SHARIF S, YAGHOOBZADEH A, YEOH KK, GOUDARZIAN AH, SOLEIMANI MA, JAMALI S. Effect of acupressure on pain in Iranian leukemia patients: a randomized controlled trial study. Int J Nurs Pract 2017; 23: doi: 10.1111/ijn.12513.
- CHAO LF, ZHANG AL, LIU HE, CHENG MH, LAM HB, LO SK. The efficacy of acupoint stimulation for the management of therapy-related adverse events in patients with breast cancer: a systematic review. Breast Cancer Res Treat 2009; 118: 255-267.
- HEIDARI GORJI A, TAEBEI M, RANJBAR M, BABAEI HATKEHLOUEI M, GOUDARZIAN AH. Effect of distraction technique and hypnosis in pain of bone marrow aspiration in children: a narrative review. Int J Pediatr 2017; 5: 4521-4531.
- 14. KNIGHT RG, WILLIAMS S, McGEE R, OLAMAN S. Psychometric properties of the Centre for Epidemiologic Studies Depression Scale (CES-D) in a sample of women in middle life. Behav Res Ther 1997; 35: 373-380.
- BAIR MJ, ROBINSON RL, KATON W, KROENKE K. Depression and pain comorbidity. Arch Intern Med 2003; 163: 2433-2445.
- 16. DOLATKHAH R, SOMI MH, BONYADI MJ, ASVADI KERMANI I, FARASSATI F, DASTGIRI S. Colorectal cancer in Iran: molecular epidemiology and screening strategies. J Cancer Epidemiol 2015; 2015: 10.
- 17. Вомса J. Cancer pain: a major national health problem. Cancer Nurs 1978; 1: 313-316.
- RADLOFF LS. The CES-D Scale: a self-report depression scale for research in the general population. Appl Psychol Meas 1977; 1: 385-401.
- Berger MB, Damico NJ, Haefner HK. Responses to the McGill pain questionnaire predict neuropathic pain medication use in women in with vulvar lichen sclerosus. J Low Genit Tract Dis 2015; 19: 135-139.
- 20. Melzack R. The McGill pain questionnaire: major properties and scoring methods. Pain 1975; 1: 277-299.
- 21. DWORKIN RH, TURK DC, TRUDEAU JJ, BENSON C, BIONDI DM, KATZ NP, KIM M. Validation of the Short-form Mc-Gill Pain Questionnaire-2 (SF-MPQ-2) in acute low back pain. J Pain 2015; 16: 357-366.
- Breitbart W. Identifying patients at risk for, and treatment of major psychiatric complications of cancer. Support Care Cancer 1995; 3: 45-60.
- 23. Наднідні **F.** Correlation between religious coping and depression in cancer patients. Psychiatr Danub 2013; 25: 236-240.
- Linden W, Vodermaier A, Mackenzie R, Greig D. Anxiety and depression after cancer diagnosis: prevalence rates by cancer type, gender, and age. J Affect Disord 2012; 141: 343-351.
- NOGHANI FATEMEH MZ, BOHRANI NASER, GHODARTI VIDA. The comparison of self esteem between male and female cancer patients. HAYAT 2006; 12: 33-41.
- 26. Bradley N, Davis L, Chow E. Symptom distress in patients attending an outpatient palliative radiotherapy clinic. J Pain Symptom Manage 2005; 30: 123-131.
- 27. Breivik H, Cherny N, Collett B, de Conno F, Filbet M, Foubert AJ, Cohen R, Dow L. Cancer-related pain: a pan-European survey of prevalence, treatment, and patient attitudes. Ann Oncol 2009; 20: 1420-1433.
- 28. Gupta M, Sahi MS, Bhargava AK, Talwar V. The prevalence and characteristics of pain in critically III cancer patients: a prospective nonrandomized observational study. Indian J Palliat Care 2015; 21: 262-267.
- 29. STRÖMGREN AS, GROENVOLD M, PETERSEN MA, GOLDSCHMIDT D, PEDERSEN L, SPILE M, IRMING-PEDERSEN G, SJOGREN P. Pain characteristics and treatment outcome for advanced cancer patients during the first week of specialized palliative care. J Pain Symptom Manage 2004; 27: 104-113.

# World Cancer Research Journal

- 30. VAN DEN BEUKEN-VAN EVERDINGEN MH, DE RIJKE JM, KESSELS AG, SCHOUTEN HC, VAN KLEEF M, PATIJN J. High prevalence of pain in patients with cancer in a large population-based study in The Netherlands. Pain 2007; 132: 312-320
- 31. Bennett MI, Rayment C, Hjermstad M, Aass N, Caraceni A, Kaasa S. Prevalence and aetiology of neuropathic pain in cancer patients: a systematic review. Pain 2012; 153: 359-365.
- 32. JAIN PN, PAI K, CHATTERJEE AS. The prevalence of severe pain, its etiopathological characteristics and treatment profile of patients referred to a tertiary cancer care pain clinic. Indian J Palliat Care 2015; 21: 148-151.
- 33. FARNAM A, ARFAEI A, NOOHI S, AZAR M, SHAFIEE K, IMANI S, KARIMZADE A. Relationship between depression and degree of pain feeling in patients with migraine headache. J Behav Sci 2008; 2: 143-148.
- 34. MOLLAZADE J, TAGHAVI M, M N. Prediction of depression based on perception of pain and quality of sleep due to pain catastrophizing in elderly patient with rheumatoid arthritis. J Anesthesiol Pain 2015; 5: 69-80.
- 35. Kroenke K, Wu J, Bair MJ, Krebs EE, Damush TM, Tu W. Reciprocal relationship between pain and depression: a 12-month longitudinal analysis in primary care. J Pain 2011; 12: 964-973.

- 36. SULLIVAN-SINGH SJ, SAWYER K, EHDE DM, BELL KR, TEMKIN N, DIKMEN S, WILLIAMS RM, HOFFMAN JM. Comorbidity of pain and depression among persons with traumatic brain injury. Arch Phys Med Rehabil 2014; 95: 1100-1105.
- 37. Mirghaforvand M, Ahmadpur P, Salehinia P. The duration and severity of depression and anxiety associated with breast pain in women. Iranian journal of Obstetrics, Gynecology and Infertility 2016; 18: 1-7.
- 38. FENIX JB, CHERLIN EJ, PRIGERSON HG, JOHNSON-HURZELER R, KASL SV, BRADLEY EH. Religiousness and major depression among bereaved family caregivers: a 13-month follow-up study. J Palliat Care 2006; 22: 286-292.
- 39. KHEZRI L, BAHREYNI M, RAVANIPOUR M, MIRZAEE K. The Relationship between spiritual wellbeing and depression or death anxiety in cancer patients in Bushehr 2015. Nursing Journal of The Vulnerable 2015; 2: 15-28.
- PEARCE MJ, SINGER JL, PRIGERSON HG. Religious coping among caregivers of terminally ill cancer patients: main effects and psychosocial mediators. J Health Psychol 2006; 11: 743-759.
- RAYNER L, HOTOPF M, PETKOVA H, MATCHAM F, SIMPSON A, McCracken LM. Depression in patients with chronic pain attending a specialised pain treatment centre: prevalence and impact on health care costs. Pain 2016; 157: 1472-1479.