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# Alexithymia and Breast Cancer Surgery: A Systematic Review

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# Authors' contributions

This work was carried out in collaboration between all authors. All authors designed the study, wrote the protocol and wrote the first draft of the manuscript. Authors DDB, SM and MC managed the literature searches. All authors wrote, read and approved the final manuscript.

## Article Information

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Review Article

#### **ABSTRACT**

The term alexithymia is a complex multidimensional construct that literally means "no words for mood". There are two kind of alexithymia: primary and secondary or otherwise called trait and state alexithymia. There is still an unresolved debate about the nature of alexithymia as a trait or a state. The etiology of alexithymia has not been completely determined because of the numerous factors influencing its development. Alexithymia has been observed among patients with a variety of psychiatric disorders and medical conditions. The aim of this systematic review was to review studies investigating the correlations between alexithymia, depression and anxiety in breast cancer surgery women. Literature search was conducted in January, 2016. PubMed and Scopus databases were used to find studies for inclusion in the systematic review. Correlations between alexithymia, depression and anxiety evidenced that alexithymia may be considered a stable personality trait and an important factor to the development of anxiety symptoms. However, limitations of these studies must be considered and further investigations are needed.

Keywords: Alexithymia; breast cancer; surgery; women; state; trait; anxiety; depression.

#### 1. INTRODUCTION

The term Alexithymia, which literally means "no words for mood", was introduced by Sifneos during the early 1970s [1]. Alexithymia is a complex multidimensional construct characterized by the following features: 1. difficulty identifying and describing subjective feelings; 2. difficulty distinguishing between feelings and bodily sensations of emotional arousal; 3. constricted imaginal capacities; 4. an externally oriented cognitive style [2]. A "classic" alexithymic subject is mainly focused on somatic sensations or symptoms and cannot give a word to emotions. In a case observed by one of the authors, a patient with irritable bowel syndrome described an emotion without recognizing it ("rage" in this occasion) as follows: "...When my boss scolded me for my carelessness without a true reason. I felt an huge muscular tension, my heartbeat was accelerated and I started to sweat as an adolescent! Obviously, I experienced, as usual in these occasions, spasms and abdominal pain. I cannot explain what it was happened to me, but these sensations are boring").

Bagby and Taylor [2] suggested that there may be two kinds of alexithymia, "primary alexithymia" which is an enduring psychological trait that does not alter over time, and "secondary alexithymia" which is state-dependent and disappears after the evoking stressful situation has changed. These two manifestations of alexithymia are otherwise called "trait" or "state" alexithymia.

There is still an unresolved debate about the nature of alexithymia as a trait or a state. Attempting to resolve that debate, a number of authors started to distinguish absolute (alexithymia scores may change across time because of symptoms changes) from relative (alexithymia scores have similar rank orders over time) stability [3,4,5].

The etiology of alexithymia has not been completely determined. Numerous authors reported a number of factors influencing the alexithymia development such as genetic factors [6,7], social environments, cultural factors, stressful situations, early childhood trauma, abuse, or neglect [8,9,10]. Recent studies revealed how alexithymia may overlap with other constructs such as autism spectrum disorders emotional intelligence, emotional [11,12], awareness and empathy deficits [13,14]. To date there are three neurobiological hypothesis of alexithymia, represented by lack of integration inter-hemispheric communication dysfunction of the right hemisphere and dysregulation of the prefrontal cortex and regions of the anterior areas during the evaluation of emotional stimuli. Moreover, a neurological damage may be, in some cases, the cause of a secondary alexithymia [16,17].

Alexithymia has been observed among patients with a variety of psychiatric disorders and medical conditions [18]. Authors reported correlations between alexithymia and depression

[4,8,19,20], schizophrenia and schizophrenia spectrum disorders [21,22], obsessivecompulsive disorders [23] anxiety disorders (panic disorder [24,25,26], post traumatic stress disorder [27,28], somatoform disorders [29], eating disorders [30,31], alcohol and drug abuse or dependence [32,33,34], body image disorder [35,36,37] and problematic gambling [38,39]. Among medical conditions, high prevalence of alexithymia has been found in essential hypertension [40], gastrointestinal diseases [41,42,43], kidney transplantation hemodialysis patients [44]. Few immune studies found that alexithymia was associated with poorer immune status [45]. Moreover, some authors suggested alexithymia as a possible vulnerability factor for medical and psychiatric disorders [46,47] and may be associated with increased mortality for any cause and suicide [48,49,50,51].

Contrasting data are available regarding the correlations between alexithymia and prognosis of patients in medical, psychiatric, or behavioral treatments. Poorer outcome has been reported for the treatment of anxiety and somatoform disorders [52], depression [53], alcoholism [54], functional gastrointestinal disorders [55]. Some studies reported beneficial effects of alexithymia on surgical treatment outcomes [56,57,58].

For women, breast cancer is the one cause of death due to cancer [59]. Treatments required for the disease, could alter the physical integrity and bodily image of women and can led patients to develop psychological disorders [60] mainly anxiety and mood disorders [61]. The prevalence of alexithymia in cancer patients has been reported to be about 8-fold greater than in the general population of Italy [62,63]. perception was found to be closely related to alexithymia, maladaptative coping and negative affect [64]. Tulipani et al. demostrated that patients who did not cancer received psychological treatment, both pain and alexithymia scores were higher than patients received psychological intervention Porcelli et al. [65] reported that psychological intervention in cancer patients didn't alter the relative stability of alexithymia, but altered the absolute stability of the construct. This study confirmed the relative stability of alexithymia and that the lack of absolute stability of the construct was influenced by psychological intervention in cancer patients.

Some authors hypothesized that alexithymia could be a risk factor for breast and uterine

carcinoma [66,67], but it also might be a psychological consequence of stress and suffering [68]. Two studies investigated the association of affect dysregulation neoplastic colonic disease, but they provided inconsistent results [69,70]. As a significant correlation has been found between alexithymia and some lymphocyte clusters as well as some inflammatory markes (such as C-Reactive Protein, CRP, and cytokines), one hypothesize that alexithymia seems to favor the development of breast cancer (and other forms of cancers) through influence on the immune system, dysblancing the cytokine network toward an increase of proinflammatory ones, generating a chronic tissue inflammation (even if often subclinical) that may contribute to cancer development [71]. However, proven data are lacking and further studies are needed.

There are several methologies to assess the alexithymia construct: self reported questionnaires and diagnostic interview. However, the self reported questionnaires represents the most used methodologies for alexithymia assessments. Among these, the 20items Toronto Alexithymia Scale (TAS) is the most used self-rating scale in the large majority of the studies [72]. The TAS-20 consists of 20 items and has a three-factor structure: Factor I assesses capacity to identify feelings and to distinguish between feelings and bodily sensations of emotional arousal (difficulty in identifying feelings [DIF]); Factor 2 reflects inability to communicate feelings to other people (difficulty in describing feelings [DDF]); and Factor 3 assesses externally-oriented thinking (EOT). the Schalling-Sifneos Personality Scales (SSPS) the Bermond-Vorst Alexithymia Questionnaire (BVAQ) [73] and the Minnesota Multiphasic Personality Inventory Alexithymia Scale (MMPI-A) [74] are the most used self reported scales. Among the most widely used diagnostic interviews we have: the Beth Israel Hospital Psychosomatic Questionnaire (BIQ) [1], the Toronto Structured Interview for Alexithymia (TSIA) [75] and the Diagnostic Criteria for Psychosomatic Research (DCPR) [76]. New assessments are represented by the Rorschach Alexithymia Scale (RAS) [77] and the California Q-Set Alexithymia Prototype (CAQ-AP).

# 2. MATHERIALS AND METHODS

## 2.1 Literature Search

A literature search was conducted in January, 2016. PubMed and Scopus databases were used

to find studies for inclusion in the systematic review. Keywords used for the search were: alexithymia, toronto alexithymia scale, breast cancer and breast cancer patients. In each search, keywords were used together with logical operators: "and", "in". Keywords used in the literature selection criteria are summarized in Table 1.

A total of ninety-nine (n = 99) articles were retrieved using PubMed and Scopus databases. Each study was required to meet all of the following criteria in order to be included in the systematic review: a) diagnosis of breast cancer; b) 20 items Toronto Alexithymia Scale scores; c) no current psychiatric treatment; d) no psychological interventions; e) breast cancer surgery; f) anxiety and depression scores. Two

(n = 2) potentially relevant studies were ultimately obtained (Fig. 1).

Table 1. Keywords used in the literature selection criteria

Keywords	Logical operators	Keywords
Alexithymia	And, in	Breast cancer
Toronto		Breast cancer
Alexithymia		patients
Scale		

SM and DDB screened each abstract and copies of any potentially relevant article were obtained. Other authors independently reviewed the articles and any disagreements in selecting the studies were resolved by discussion.

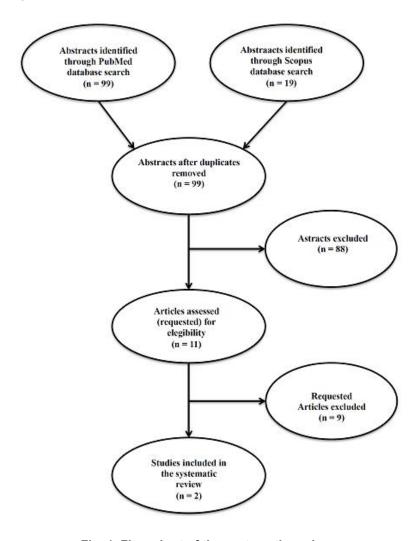


Fig. 1. Flow chart of the systematic review

Table 2. Studies evaluating alexithymia, depression and anxiety in breast cancer surgery women

Authors	Year	Participants	Mean age	Type of surgery	Parameters studied	Time since surgery	Aim of the study	Main findings
Mantani et al.	2007	n = 46 postsurgical ambulatory women with breast cancer	52.3±10.5	Mastectomy or lumpectomy	-SAS -SDS -FAD -TAS-20	Months elapsed after surgery 18.3±11.7	To explore how alexithymia and family functioning might affect anxiety and depression levels	Alexithymia is important for the development of anxiety symptoms and perceptions of inappropriate affective responsiveness among family members correlated with high degree of depression
Luminet et al.	2007	n = 122 breast cancer surgery women	56.2±10.3	Not reported	-HADS -TAS-20	The day before surgery and 6 months later	To examine stability for alexithymia factor scores	Correlations between alexithymia, depression and anxiety scores at baseline and follow-up evidenced a strong argument in favor of the relative stability alexithymia, as a stable personality trait rather than a state dependent phenomenon

SAS: Zung self-rating anxiety scale, SDS: Zung self-rating depression scale, TAS-20: 20-item Toronto Alexithymia Scale, FAD: Family Assessment Device, HADS: Hospital Anxiety and Depression Scale

#### 3. DISCUSSION

There are only two (n = 2) studies in literature meeting the selection criteria of the systematic review. Both of these studies correlate alexithymia, breast cancer surgery, depression and anxiety (Table 2).

Mantani et al. [78] explored how alexithymia, family functioning and other factors could be related to anxiety and depression in fourty six (n = 46) outpatients, represented by women who had undergone surgery for breast cancer at least three months before the interview. High anxiety levels were significantly associated to high TAS-20 total score and to communication and affective responsiveness subscales of the Family Assessment Device (FAD). Depression levels were associated with much more variables: high TAS-20 total score, high scores on the communication, affective responsiveness, behavior control, and general functioning subscale scores of the FAD, and the presence of pain. Multiple regression analysis reported that only high TAS-20 total score and only high score on the affective responsiveness subscale of the FAD were correlated respectively with high degree of anxiety and high depression levels. In other words, alexithymia were important for the development of anxietv symptoms and perceptions of inappropriate affective responsiveness among family members correlated with high degree of depression.

Luminet et al. [79] evaluated one hundred and twenty two (n = 122) women in acute (Time 1) and in chronic (Time 2) stress periods: the day before surgery and six months later. The aims of the study were to evaluate the relationship between alexithymia, depression and anxiety and to examin absolute and relative stability for the three components. Depression was positively correlated with both total and factors scores of alexithymia ("difficulty indentifying feeling" (DIF), "externally oriented thinking" (EOT), "difficulty describing feelings" (DDF)), both at baseline and at follow-up. Correlations between anxiety and total alexithymia scores were also significant. However, the relationship varied across factors. DIF was highly related to anxiety both at baseline and follow-up; DDF correlated to anxiety only at follow-up; correlations were not significant between anxiety and EOT. Concerning absolute stability changes in scores were reported from Time 1 to Time 2 only for anxiety and alexithymia. Decrease was found for anxiety. About alexithymia increased scores

were reported regarding total scores, the factors DIF and EOT, while DDF was completely stable. Changes were not significant for depression. Correlations between alexithymia, depression and anxiety scores at baseline and follow-up evidenced a strong argument in favor of the relative stability alexithymia.

Limitations of the studies investigated in this systematic review include: a) small sample sizes; b) not reported adjuvant therapy; c) unknown time elapsed since diagnosis and surgery; d) unknown breast cancer stages. Further studies in breast cancer surgery women should: a) identify risk factors for alexithymia; b) better understand the nature of alexithymia on the lack of emotional expression; c) identify possible correlations between alexithymia and adjuvant therapies; d) investigate relationships between alexithymia and quality of life; e) search correlations between alexithymia and breast cancer stage; f) find differences between patients receiving palliative treatment and those undergoing breast caner surgery.

#### 4. CONCLUSIONS

For women breast cancer is the one cause of death due to cancer. Alexithymia is hypothesized to be a risk factor for the development of breast cancer. The prevalence of alexithymia in cancer patients has been reported to be about 8-fold greater than in the general population of Italy. There are numerous factors influencing the development of the alexithymia. Recently many neuroimaging studies demostrated dysfunction of the right hemisphere and dysregulation of the prefrontal cortex and regions of the anterior areas during the evaluation of emotional stimuli. The studies we investigated in this review report two findings: a) support the role of the alexithymia in the development of anxiety symptoms among family members correlated with high degree of depression; b) demostrate the relative stability of alexithymia the day before surgery and after 6 months even if the construct were closely associated with the severity and improvement of cancer-related pain. Limitations of the studies investigated in this systematic review must be considered in further investigations.

However, on the basis of these studies and in the light of what has emerged from the present systematic review, it is possible to hypothesize that the prevention or the treatment of primary or secondary alexithymia may be beneficial to

reduce the incidence of breast cancer or alleviate the post-surgical complications [80,81]. No pharmacological therapies for alexithymia exist. Psychotherapy may be effective in some cases [82]. As pointed out by Vanheule et al. [83], ...alexithymic subjects should focus distressing situations, starting from which a three-step logic can be deployed. During therapy, mental representations on difficult situations in patients' lives need to be constructed by (1) putting into words the chain of events that makes up the distressing situation; (2) making the patient's appraisal of the difficult situation explicit: and (3) addressing affective responses and discussing the patient's way of dealing with the difficult situation". This method may be applied to alexithymic women with at risk of or affected by breast cancer, bu to date no studies have been conducted on this topic.

#### CONSENT

It is not applicable.

## ETHICAL APPROVAL

It is not applicable.

## **DISCLOSURE**

This manuscript was entirely funded by the authors, and no pharmaceutical companies were informed of or were involved in the review. All authors contributed to this review with equal efforts. The authors have no potential conflict of interest directly relevant to the contents of the manuscript.

#### **COMPETING INTERESTS**

Authors have declared that no competing interests exist.

# **REFERENCES**

- Sifneos PE. The prevalence of alexithymic characteristics in psychosomatic patients. Psychother Psychosom. 1973;22(2):255-62
- Taylor G, Bagby R. An overview of the alexithymia construct. In The handbook of emotional intelligence: Bar-On R, Parker JDA, editors.. San Francisco Jossey Bass. 2000;40–67.
- 3. Luminet O, Bagby RM, Taylor GJ. An evaluation of the absolute and relative

- stability of alexithymia in patients with major depression. Psychother Psychosom. 2001;70(5):254-60.
- Saarijarvi S, Salminen JK, Toikka T. Temporal stability of alexithymia over a five-year period in outpatients with major depression. Psychother Psychosom. 2006; 75(2):107-12.
- 5. Tolmunen T, Heliste M, Lehto SM, Hintikka J, Honkalampi K, Kauhanen J. Stability of alexithymia in the general population: An 11-year follow-up. Compr Psychiatry. 2011;52(5):536-41.
  - DOI: 10.1016/j.comppsych.2010.09.007 Epub 2010 Nov 16.
- Fantini-Hauwel C, Dauvier B, Arciszewski T, Antoine P, Manouvrier S. Genetic testing for hereditary cancer: Effects of alexithymia and coping strategies on variations in anxiety before and after result disclosure. Psychol Health. 2011;26(7): 855-73.
  - DOI: 10.1080/08870446.2010.506575 Epub 2011 May 24.
- 7. Taylor GJ, Bagby RM. Genetic contributions to alexithymia. Psychosom Med. 2011;73(7):633;author reply 633-5. DOI: 10.1097/PSY.0b013e31822ed5ba Epub 2011 Aug 26.
- 8. De RA, Vanheule S. The relationship between perceived parenting, adult attachment style and alexithymia in alcoholic inpatients. Addict Behav. 2006, 31:1265–1270.
- Joukamaa M, Kokkonen P, Veijola J, Laksy K, Karvonen JT, Jokelainen J, Jarvelin MR: Social situation of expectant mothers and alexithymia 31 years later in their offspring: A prospective study. Psychosom Med. 2003;65:307–312.
- Kokkonen P, Veijola J, Karvonen JT, Laksy K, Jokelainen J, Jarvelin MR, Joukamaa M. Ability to A speak at the age of 1 year and alexithymia 30 years later. J Psychosom Res. 2003;54:491–495.
- Fitzgerald M, Molyneux G: Overlap between alexithymia and Asperger's syndrome. Am J Psychiatry. 2004;161: 2134–2135.
- Berthoz S, Hill EL. The validity of using self-reports to assess emotion regulation abilities in adultswith autism spectrum disorder. Eur Psychiatry. 2005;20:291– 298.
- Pino MC, De Berardis D, Mariano M, Vellante F, Serroni N, Valchera A, Valenti M, Mazza M. Two systems for empathy in

- obsessive-compulsive disorder: Mentalizing and experience sharing. Rev Bras Psiquiatr; 2016. (In press)
- Bird G, Silani G, Brindley R, White S, Frith U, Singer T. Empathic brain responses in insula are modulated by levels of alexithymia but not autism. Brain. 2010; 133:1515–1525.
- 15. Parker JD, Keightley ML, Smith CT, Taylor GJ. Interhemispheric transfer deficit in alexithymia: An experimental study. Psychosom Med. 1999;61(4):464-8.
- Redaktsiia, Antonova NA, Kuznetsova EB, Sholomov II. Alexithymia in patients with left-hemispheric cerebral infarction in the late recovery period. Zh Nevrol Psikhiatr Im S S Korsakova. 2015;115:3-6.
- Neumann D, Zupan B, Malec JF, Hammond F. Relationships between alexithymia, affect recognition, and empathy after traumatic brain injury. J Head Trauma Rehabil. 2014;29:18-27.
- Taylor GJ, Bagby RM, Parker JDA. Disorders of affect regulation alexithymia in medical andpsychiatric illness. Cambridge University Press, Cambridge; 1997.
- De Berardis D, Serroni N, Campanella D, Carano A, Gambi F, Valchera A, Conti C, Sepede G, ScaliM, Fulcheri M, Salerno RM, Ferro FM. Alexithymia and its relationships with C-reactive protein and serum lipid levels among drug naïve adult outpatients with major depression. Prog Neuropsychopharmacol Biol Psychiatry. 2008;32(8):1982-6.
  - DOI: 10.1016/j.pnpbp.2008.09.022 Epub 2008 Oct 7.
- Honkalampi K, Hintikka J, Laukkanen E, Lehtonen J, Viinamaki H. Alexithymia and depression: A prospective study of patients with major depressive disorder. Psychosomatics. 2001;42:229–234.
- Marasco V, De Berardis D, Serroni N, Campanella D, Acciavatti T, Caltabiano M, Olivieri L, Rapini G, Cicconetti A, Carano A, La Rovere R, Di Iorio G, Moschetta FS, Di Giannantonio M. Alexithymia and suicide risk among patients with schizophrenia: Preliminary findings of a cross-sectional study. Riv Psichiatr. 2011; 46(1):31-7.
- van 't Wout M, Aleman A, Bermond B, Kahn RS. No words for feelings: Alexithymia in schizophrenia patients and first-degree relatives. Compr Psychiatry. 2007;48(1):27-33.

- De Berardis D, Campanella D, Gambi F, Sepede G, Salini G, Carano A, La Rovere R, Pelusi L, Penna L, Cicconetti A, Cotellessa C, Salerno RM, Ferro FM. Insight and alexithymia in adult outpatients with obsessive-compulsive disorder. Eur Arch Psychiatry Clin Neurosci. 2005; 255(5):350-8. Epub 2005 Feb 21.
- 24. Parker JD, Taylor GJ, Bagby RM, Acklin MW. Alexithymia in panic disorder and simple phobia: A comparative study. Am J Psychiatry. 1993;150:1105–7.
- De Berardis D, Campanella D, Serroni N, Moschetta FS, Di Emidio F, Conti C, Carano A, Acciavatti T, Di Iorio G, Martinotti G, Siracusano A, Di Giannantonio M. Alexithymia, suicide risk and serum lipid levels among adult outpatients with panic disorder. Compr Psychiatry. 2013 Janpii: S0010-440X(12) 00269-6.
  - DOI: 10.1016/j.comppsych.2012.12.013 [Epub ahead of print].
- 26. De Berardis D, Campanella D, Gambi F, La Rovere R, Sepede G, Core L, Canfora G, Santilli E, Valchera A, Mancini E, Salerno RM, Moschetta FS, Ferro FM. Alexithymia, fear of bodily sensations, and somatosensory amplification in young outpatients with panic disorder. Psychosomatics. 2007;48(3):239-46.
- Frewen PA, Pain C, Dozois DJ, Lanius RA. Alexithymia in PTSD: psychometric and FMRI studies. Ann N Y Acad Sci. 2006;1071:397-400.
- 28. Badura AS. Theoretical and empirical exploration of the similarities between emotional numbing in posttraumatic stress disorder and alexithymia. Journal of Anxiety Disorders. 2003;17:349–360.
- Waller E, Scheidt CE. Somatoform disorders as disorders of affect regulation: a study comparing the TAS-20 with nonself-report measures of alexithymia. J Psychosom Res. 2004;57:239–47.
- 30. de Groot JM, Rodin G, Olmsted MP. Alexithymia, depression, and treatment outcome in bulimia nervosa. Compr Psychiatry. 1995;36:53–60.
- Carano A, De Berardis D, Campanella D, Serroni N, Ferri F, Di Iorio G, Acciavatti T, Mancini L, Mariani G, Martinotti G, Moschetta FS, Di Giannantonio M. Alexithymia and suicide ideation in a sample of patients with binge eating disorder. J Psychiatr Pract. 2012;18:5-11.

- 32. Loas G, Otmani O, Lecercle C, Jouvent R. Relationships between the emotional and cognitive components of alexithymia and dependency in alcoholics. Psychiatry Res 2000:96:63–74.
- 33. El Rasheed AH. Alexithymia in Egyptian substance abusers. Substance Abuse. 2001;22:11–21.
- 34. Haviland MG, Hendryx MS, Shaw DG, Henry JP. Alexithymia in women and men hospitalized forpsychoactive substance dependence. Comprehensive Psychiatry 1994;35:124–128.
- 35. De Berardis D, Campanella D, Gambi F, Sepede G, Carano A, Pelusi L, La Rovere R, Di Matteo D, Salini G, Cotellessa C, Salerno RM, Ferro FM. Alexithymia and body image disturbances in women with Premenstrual Dysphoric Disorder. J Psychosom Obstet Gynaecol. 2005;26(4): 257-64.
- 36. Carano A, De Berardis D, Gambi F, Di Paolo C, Campanella D, Pelusi L, Sepede G, Mancini E, La Rovere R, Salini G, Cotellessa C, Salerno RM, Ferro FM. Alexithymia and body image in adult outpatients with binge eating disorder. Int J Eat Disord. 2006;39(4):332-40.
- 37. De Berardis D, Carano A, Gambi F, Campanella D, Giannetti P, Ceci A, Mancini E, La Rovere R, Cicconetti A, Penna L, Di Matteo D, Scorrano B, Cotellessa C, Salerno RM, Serroni N, Ferro FM. Alexithymia and its relationships with body checking and body image in a non-clinical female sample. Eat Behav. 2007;8:296-304.
- 38. Lumley MA, Roby KJ. Alexithymia and pathological gambling. Psychotherapy & Psychosomatics. 1995;63:201–206.
- Parker JD, Wood LM, Bond BJ, Shaughnessy P. Alexithymia in young adulthood: A risk factor for pathological gambling. Psychotherapy & Psychosomatics 2005;74:51–55.
- Jula A, Salminen JK, Saarijarvi S. Alexithymia: A facet of essential hypertension. Hypertension. 1999;33: 1057–61.
- 41. Porcelli P, Taylor GJ, Bagby RM, De Carne M. Alexithymia and functional gastrointestinal disorders A comparison with inflammatory bowel disease. Psychother Psychosom. 1999;68:263–9.
- 42. Jones MP, Schettler A, Olden K, Crowell MD. Alexithymia and somatosensory

- amplification in functional dyspepsia. Psycosomatics. 2004;45:508–516.
- Portincasa P, Moschetta A, Baldassarre G, et al. Pan-enteric dysmotility, impaired quality of life and alexithymia in a large group of patients meeting ROME II criteria for irritable bowel syndrome. World J Gastroenterol. 2003;9:2293-2299.
- 44. Fukunishi I. The influence of defense mechanisms on secondary alexithymia in hemodialysis patients. Psychother Psychosom. 1992;57:50–56.
- 45. De Berardis D, Conti C, Iasevoli F, Valchera A, Fornaro M, Cavuto M, Brucchi M, Perna G, Pompili M, Modabbernia A, Lucidi G, Mazza M, Martinotti G, Di Giannantonio M. Alexithymia and its relationships with acute phase proteins and cytokine release: An updated review. J Biol Regul Homeost Agents. 2014;28: 795-9.
- 46. Taylor GJ. Recent developments in alexithymia theory and research. Can J Psychiatry. 2000;45:134–42.
- 47. Kauhanen J, Kaplan GA, Cohen RD, Julkunen J, Salonen JT. Alexithymia and risk of death in middle-aged men. J Psychosom Res. 1996;41(6):541-9.
- 48. Tolmunen T, Lehto SM, Heliste M, Kurl S, Kauhanen J. Alexithymia is associated with increased cardiovascular mortality in middle-aged finnish men. Psychosom Med. 2010;72:187-91.
- 49. De Berardis D, Serroni N, Campanella D, Marini S, Rapini G, Valchera A, lasevoli F, Mazza M, Fornaro M, Perna G, Di Iorio G, Martinotti G, Di Giannantonio M. Alexithymia, suicide ideation, C-reactive protein, and serum lipid levels among outpatients with generalized anxiety disorder. Arch Suicide Res. 2015;1-13.
- 50. De Berardis D, Serroni N, Campanella D, Rapini G, Olivieri L, Feliziani B, Carano A, Valchera A, Iasevoli F, Tomasetti C, Mazza M, Fornaro M, Perna G, Di Nicola M, Martinotti G, Di Giannantonio M. Alexithymia, responsibility attitudes and suicide ideation among outpatients with obsessive-compulsive disorder: An exploratory study. Compr Psychiatry. 2015; 58:82-7.
- De Berardis D, Serroni N, Marini S, Rapini G, Carano A, Valchera A, Iasevoli F, Mazza M, Signorelli M, Aguglia E, Perna G, Martinotti G, Varasano PA, Pressanti GL, Di Giannantonio M. Alexithymia, suicidal ideation, and serum lipid levels

- among drug-naïve outpatients with obsessive-compulsive disorder. Rev Bras Psiquiatr. 2014;36:125-30.
- 52. De Berardis D, Campanella D, Serroni N, Sepede G, Carano A, Conti C, et al. The impact of alexithymia on anxiety disorders: a review of the literature. Curr Psychiatry Rev. 2008;4:80-6.
- 53. Ogrodniczuk JS, Piper WE, Joyce AS. Alexithymia as a predictor of residual symptoms in depressed patients who respond to short-term psychotherapy. Am J Psychother. 2004;58(2):150-61.
- 54. Loas G, Fremaux D, Otmani O, Lecercle C, Delahousse J. Is alexithymia a negative factor for maintaining abstinence? A follow-up study. Compr Psychiatry. 1997;38: 296-9.
- 55. Porcelli P, Bagby RM, Taylor GJ, De Carne M, Leandro G, Todarello O. Alexithymia as a predictor of treatment outcome in patients with functional gastrointestinal disorders. Psychosomatic Medicine. 2003;65:911–918.
- Tulipani C, Morelli F, Spedicato MR, Maiello E, Todarello O, Porcelli P. Alexithymia and cancer pain: the effect of psychological intervention. Psychother Psychosom 2010;79:156–163.
- Ripetti V, Ausania F, Bruni R, Campoli G, Coppola R. Quality of life following colorectal cancersurgery: the role of alexithymia. Eur Surg Res. 2008;41: 324–330.
- 58. Battista C, Angioli R, Cafa EV, Sereni MI, Vulcano E, Bruni R. Alexithymia-a disorder of the regulatory mechanism of the emotion elaboration-and quality of life in gynecologic surgery. J Minim Invasive Gynecol. 2009;16:63–67.
- 59. Coleman MP, Gatta G, Verdecchia A, Esteve J, Sant M, Storm H, Allemani C, Ciccolallo L, Santaquilani M, Berrino F. EUROCARE-3 summary: Cancer survival in Europe at the end of the 20<sup>th</sup> century. Ann Oncol. 2003;14(Suppl 5):128-49.
- Sellick SM, Crooks DL. Depression and cancer: an appraisal of the literature for prevalence, detection, and practice guideline development for psychological interventions. Psychooncology. 1999;8: 315-33.
- Compas BE, Stoll MF, Thomsen AH, Oppedisano G, Epping- Jordan JE, Krag DN. Adjustment to breast cancer: Agerelated differences in coping and emotional

- distress. Breast Cancer Res Treat 1999;54:195–203.
- 62. Manna G, Foddai E, Di Maggio MG, et al. Emotional expression and coping style in female breast cancer. Ann Oncol. 2007; 18(suppl 6):vi77-80.
- 63. Messina A, Fogliani AM, Paradiso S. Alexithymia in oncologic disease: association with cancer invasion and hemoglobin levels. Ann Clin Psychiatry. 2011;23(2):125-30.
- Porcelli P, Tulipani C, Maiello E, Cilenti G, Todarello O. Alexithymia, coping, and illness behavior correlates of pain experience in cancer patients. Psycho-Oncology. 2007;16:644–650.
- 65. Porcelli P, Tulipani C, Di Micco C, Spedicato MR, Maiello E. Temporal stability of alexithymia in cancer patients following a psychological intervention. J Clin Psychol. 2011;67(12):1177-87. DOI: 10.1002/jclp.20839 Epub 2011 Nov 3.
- 66. Carta M, Orru W, Hardoy MC, Carpiniello B. Alexithymia and early diagnosis of uterine carcinoma: Results of a case-control study. Psychotherapy Psychosomatics. 2000;69:339–340.
- 67. Manna G, Foddai E, Di Maggio MG, Pace F, Colucci G, Gebbia N, et al. Emotional expression and coping style in female breast cancer. Annals of Oncology. 2007; 18:77–80.
- 68. Celikel FC, Saatcioglu O. Alexithymia and anxiety in female chronic pain patients. Ann Gen Psychiatry. 2006;5:13.
- 69. Kune GA, Kune S, Watson LF. Colorectal cancer risk, chronic illnesses, operations and medications: Case–control results from the melbourne colorectal cancer study. International Journal of Epidemiology. 2007;36:951–957.
- Nagano J, Kono S, Toyomura K, Mizoue T, Yin G, Mibu R, et al. Personality and colorectal cancer: The Fukuoka colorectal cancer study. Japan Journal of Clinical Oncology. 2008;38:553-561.
- Honkalampi K, Lehto SM, Koivumaa-Honkanen H, Hintikka J, Niskanen L, Valkonen-Korhonen M, Viinamäki H. Alexithymia and tissue inflammation. Psychother Psychosom. 2011;80:359-64.
- Taylor GJ, Bagby RM, Ryan DP, Parker JDA. Validation of the alexithymia construct: A measurement-based approach. Can J Psychiatry. 1990;35: 290-7.

- Bermond B, Vorst HC. Validity and reliability of the bermond-vorst alexithymia ques- tionnaire. Unpublished manuscript. The Netherland: University of Amsterdam; 1998.
- 74. Kleiger JH, Kinsman RA. The development of an MMPI Alexithymia scale. Psychother Psychosom. 1980;34:17-24.
- Haviland MG, Reise SP. Structure of the twenty-item Toronto Alexithymia scale. J Pers Assess. 1996b;66:116-25.
- 76. Mangelli L, Rafanelli C, Porcelli P, Fava GA. Interview for the Diagnostic Criteria for Psychosomatic Research (DCPR). In: Rafanelli C, Roncuzzi R, Finos L, et al. Psychological assessment in cardiac rehabilitation. Psychother Psychosom 2003;72:343-9.
- 77. Porcelli P, Mihura JL. Assessment of alexithymia with the rorschach comprehensive system: the Rorschach Alexithymia Scale (RAS). J Pers Assess 2010:92:128-36.
- 78. Mantani T, Saeki T, Inoue S, Okamura H, Daino M, Kataoka T, Yamawaki S. Factors related to anxiety and depression in

- women with breast cancer and their husbands: role of alexithymia and family functioning. Support Care Cancer. 2007; 15(7):859-68.
- Luminet O, Rokbani L, Ogez D, Jadoulle V. An evaluation of the absolute and relative stability of alexithymia in women with breast cancer. Journal of Psychosomatic Research. 2007;62:641–648.
- 80. Apfel RJ, Sifneos PE. Alexithymia: Concept and measurement. Psychother Psychosom. 1979;32:180-90.
- Ripetti V, Ausania F, Bruni R, Campoli G, Coppola R. Quality of life following colorectal cancer surgery: The role of alexithymia. Eur Surg Res. 2008;41: 324-30.
- 82. Reeves RR, Johnson-Walker D. Alexithymia: Should this personality disorder be considered during treatment of patients with mental illness? J Psychosoc Nurs Ment Health Serv. 2015;53:25-9.
- Vanheule S, Verhaeghe P, Desmet M. In search of a framework for the treatment of alexithymia. Psychol Psychother. 2011;84: 84-97.

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