

## ORIGINAL ARTICLE / ОРИГИНАЛНИ РАД

# Association between bipolar affective disorder, use of antidepressants and osteoporosis

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## SUMMARY

**Introduction/Objective** Osteoporosis is one of the most common comorbid disorders in depressive mood disorder. The aim of this study was to assess the association between the use of antidepressants and osteoporosis in patients with bipolar affective disorder (BPAD).

**Methods** The study included 73 inpatients, aged 50–72 years, male and female, hospitalized with a depressive episode of BPAD from 2016 to 2020 at the Clinic of Psychiatry, Clinical Centre of Vojvodina, divided into two groups: a) the first group (40) was treated with selective serotonin reuptake inhibitors (SSRIs) in combination with mood stabilizer (lithium carbonate/lamotrigine); b) the second group (33) was treated with mood stabilizer only. Study included two control groups as well. Clinical measurements of bone mineral density at lumbar spine and hip was made using dual energy X-ray absorptiometry. CrossLaps and levels of calcium and vitamin D were collected from blood samples. The data was analyzed by the analysis of variance and the Kruskal–Wallis test.

**Results** Osteoporosis was registered in 25% of patients in the first group and in 18% of patients in the second group, while osteopenia was observed within 40% of patients in the first group and in 37% of patients in the second group. There was significant difference in value of CrossLaps, and the level of 25(OH)D vitamin between the control groups and the first two groups, as well as in prevalence of osteoporosis and osteopenia.

**Conclusion** Depressive episodes in BPAD is connected with higher prevalence of osteoporosis. Patients treated with SSRIs have higher prevalence of osteoporosis than patients treated with mood stabilizers only.

**Keywords:** bone mineral density; depression; selective serotonin reuptake

## INTRODUCTION

Depression has been reported as the most common mental disorder in the 21st century based on its incidence and prevalence that have been increasing constantly over the last decades, not only in Serbia but in the majority of other countries with valid health statistics [1, 2]. Besides the incidence rate, depression is significant because numerous metabolic disorders can result from an untreated or inadequately treated mental disorder. Osteoporosis is one of the most common comorbid disorders, especially in bipolar and unipolar mood disorders. Nowadays, there is a risk of osteoporosis becoming a “silent epidemic,” just like mood disorders, which are both metabolic disorders. According to the official statistics, every third woman and every sixth man over 60 is affected by osteoporosis [3]. The risk of these health problems is growing substantially considering the common comorbidity of mood disorders and osteoporosis [4, 5]. Well-known risk factors are biological predisposition, sex, age, positive family history, low body weight and bad habits (smoking, consuming alcohol, fast food, greasy and poor-quality food, lack of physical activity) as well as the use of certain medications (e.g.,

corticosteroids) [6, 7]. In addition to those risk factors, it is not clear whether the same pathophysiological processes take place in mood disorders and osteoporosis [1, 2].

Common facts that prove the existence of common pathophysiological processes in both disorders are considered to be hypercortisolemia, increased activity of the hypothalamic-pituitary-adrenal axis, increased cytokine activity (interleukin-6 (IL-6) and tumor necrosis factor (TNF)), and a decrease in anti-inflammatory interleukin activity (IL-10, IL-13) as well as an increase in oxidative stress factors, an increase in parathyroid hormone levels with consecutive decrease of 25(OH)D vitamin, and the decrease in estrogen levels in plasma [6, 8–11].

Although the association between depression and osteoporosis has not been clearly explained, recent studies suggest that depression should be considered as an official risk factor for osteoporosis [12]. Depressive disorder can be represented as unipolar (depressive episodes only), or within bipolar disorder, when depressive symptoms are replaced or can overlap with manic/hypomanic symptoms. Few clinical studies have been focused on monitoring the risk for osteoporosis within bipolar patients [13].

**Received • Примљено:**

August 11, 2020

**Revised • Ревизија:**

August 13, 2021

**Accepted • Прихваћено:**

August 17, 2021

**Online first:** November 17, 2021

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The aim of this study was to assess the association between developing osteoporosis and taking antidepressants in patients with the diagnosis of bipolar affective disorder.

## METHODS

The study included 73 patients, aged 50–72 years, both male and female, hospitalized with diagnosis of bipolar affective disorder [middle, recurrent, depressive episode, more than 20 on the Hamilton Depression Scale (HAMD)], 2016–2020 at the Clinic of Psychiatry, Clinical Centre of Vojvodina. Also, two control groups were included: a) the first control group (30) included depressive patients without psychopharmacotherapy; b) the second control group (30) was formed of healthy volunteers.

Patients with psychopharmacotherapy (73) were divided into two groups: a) the first group (40) was treated with selective serotonin reuptake inhibitors (SSRIs) antidepressants (escitalopram, fluoxetine, sertraline) in combination with mood stabilizers (lithium carbonate and/or lamotrigine). Antidepressants were added to therapy due to non-reactive depression; b) the second group (33) was treated with mood stabilizers only, without antidepressants.

Sociobiographic and sociodemographic data were collected, including information about all medication. Diabetes mellitus, hypothyroidisms, hyperlipidemia, and hypertension were the most common comorbidities. The patients were under medication control. The diagnosis of osteoporosis was based on the ICD 10 code. Clinical measurements and assessment of BMD at lumbar spine (L2–L4) and hip was made, using dual energy X-ray absorptiometry (DEXA). Also, CrossLaps and vitamin D and calcium levels were ascertained from blood samples [14].

All the patients signed the consent according to the Declaration of Helsinki. The study was approved by the competent ethics committee of the Clinical Center of Vojvodina, and conforms to the legal standards. For the purpose of statistical analysis, IBM SPSS Statistics, version 21.0 (IBM Corp., Armonk, NY, USA) and JASP, version

0.14.1.0 (University of Amsterdam, Amsterdam, the Netherlands) were used. Results were statistically analyzed by one-way analysis of variance – ANOVA and Tukey's Honest Significant Difference post-hoc test as parametric method of testing and by Kruskal–Wallis test with post-hoc Mann–Whitney U-tests as a non-parametric method of testing. The presence of osteoporosis or osteopenia was ascertained using the analysis of contingency tables. A p-value less than 0.05 was considered statistically significant.

## RESULTS

The average age in the first group was  $56.2 \pm 7.8$  years, and it was  $55.9 \pm 4.3$  years in the second group. In both control groups, the average age was similar ( $55 \pm 3.3$  years in the first control group and  $56.1 \pm 4.1$  years in the second control group). Women were represented in a larger percentage in all groups: 66% vs. 34% and 67% vs. 33% (control groups: 68% vs. 32% and 75% vs. 25%). Mood stabilizers such as lithium carbonate and lamotrigine were included:  $7.8 \pm 3.6$  years in the first group, and  $8.4 \pm 3.2$  years in the second group. Patients with comorbid physical disorders (hypothyroidism, diabetes mellitus, and arterial hypertension) were also included in this study. All comorbidities were adequately treated. Forty patients were administered with SSRI antidepressants, which are indicated in the treatment of moderate depressive episodes in combination with a mood stabilizer. Thirty-three patients were treated with mood stabilizers as monotherapy. The following SSRI antidepressants were administered: sertraline in 68% of patients in an average dose of 75 mg/day, escitalopram in 20% of patients in an average dose of 10 mg/day, and fluoxetine in 12% of patients in an average dose of 20 mg/day. The average length of SSRI was 72 days (Table 1).

In Table 2 are presented values of indicators of osteoporosis in the patients of all four groups: osteoporosis was registered in 25% of patients treated with mood stabilizers and SSRI antidepressants and in 18% of patients treated with mood stabilizers, while osteopenia was observed

**Table 1.** Type of psychopharmacotherapy

Patients	First group	Second group	Third group	Fourth group	p	
	SSRI and mood stabilizers in therapy ( $\pm$ SD)	Mood stabilizers in therapy ( $\pm$ SD)	Control group of patients with depressive disorder, without SSRI or mood stabilizers in therapy ( $\pm$ SD)	Control group of healthy volunteers ( $\pm$ SD)		
n = 133	n = 40	n = 33	n = 30	n = 30	No significant difference	
Age	$56.2 \pm 7.8$	$55.9 \pm 4.3$	$55 \pm 4.3$	$54.2 \pm 3.2$	No significant difference	
Sex	Male	34%	33%	32%	25%	No significant difference
	Female	66%	67%	68%	75%	No significant difference
Duration of therapy with mood stabilizers (in years)	$7.8 \pm 3.6$	$8.4 \pm 3.9$	-	-	No significant difference	
Dose of SSRIs: sertraline (68%) escitalopram (20%) fluoxetine (12%)	75 mg/day 10 mg/day 20 mg/day	-	-	-	-	

SSRI – selective serotonin reuptake inhibitor

**Table 2.** Parameters of bone metabolism

Bone metabolism parameters	First group	Second group	Third group	Fourth group	p
	SSRI and mood stabilizers in therapy (n = 40) (± SD)	Mood stabilizers in therapy (n = 33) (± SD)	Control group of patients with depressive disorder, without SSRI or mood stabilizers in therapy (n = 30) (± SD)	Control group of healthy volunteers (n = 30) (± SD)	
25(OH)D vitamin (ng/ml)	24.5 ± 6.7 <sup>a</sup>	29.1 ± 8.5 <sup>a</sup>	37.5 ± 7.2 <sup>b</sup>	51.1 ± 11.4	< 0.001*
Ca++ (mmol/l)	1.1 ± 0.1 <sup>c</sup>	1 ± 0.1 <sup>a</sup>	1.1 ± 0.1	1.1 ± 0.1	< 0.001*
β-CrossLaps (ng/l)	760.9 ± 129.4 <sup>a</sup>	780.5 ± 84 <sup>a</sup>	583.7 ± 48.3 <sup>b</sup>	380 ± 84.9	< 0.001*
Osteopenia	40% <sup>d</sup>	37%	32%	20%	< 0.01*
Osteoporosis	25% <sup>e</sup>	18%	16%	10%	< 0.01*

SSRI – selective serotonin reuptake inhibitor;

<sup>a</sup>p < 0.001 compared with the third and fourth group;

<sup>b</sup>p < 0.001 compared with the fourth group;

<sup>c</sup>p < 0.001 compared with the second group;

<sup>d</sup>p < 0.01 compared with the fourth group;

<sup>e</sup>p < 0.01 compared with the second, third, and fourth group;

\*significant difference

within 40% of patients treated with the combination of SSRI and mood stabilizers and in 37% of patients treated with mood stabilizers only. In the first control group, osteoporosis was detected in 16% of patients and osteopenia was present in 32% of the patients. In the group of healthy volunteers, osteoporosis was presented in 10% of examinees, while osteopenia was detected in 20% of examinees. There was no significant difference in the value of CrossLaps and 25(OH)D vitamin between the first two groups of patients. There was a significant difference between both control groups and other two groups in the prevalence of osteoporosis and osteopenia as well as in the value of CrossLaps and 25(OH)D vitamin.

Smokers were highly represented in all the patient groups (98% in patients treated with SSRI vs. 99% in patients treated just with mood-stabilization medicaments). In the control groups, smokers represented 86.5% of the sample.

## DISCUSSION

Numerous results indicate strong relationship between use of antidepressants and osteoporosis. In addition to undoubtedly common comorbidity of depression and osteoporosis, the measures for prevention and early diagnosis need to be taken in order to have an early detection, treatment and lowering of the complication and mortality rate of both disorders [1, 4, 10, 11, 14].

According to the World Health Organization reports, mood disorders represent a huge health problem in comorbidity with osteoporosis and there is no doubt that unrecognized and untreated symptoms of depression, as well as anxiety symptoms, extensive use of certain types of antidepressants have been associated with an increased risk of osteoporosis in the last decade [4, 15, 16]. Osteoporosis is related to higher incidence of hip fracture in women over 60 years old treated with therapeutic doses of SSRI in comparison with depressive patients treated with other antidepressants [17, 18]. The most supported assumption in literature nowadays is that depression, as a separate mood disorder, triggers the development of osteoporosis

through neuroendocrine and immune mechanisms as well as through bad habits (poor nutrition, alcohol intake, smoking, lack of physical activity) [3–5, 7, 15, 16]. The results have been controversial up to now, even though there is still a large number of studies that do not support the assumption that SSRI antidepressants play neuroendocrine role in bone metabolism. However, all the data require adequate choice of antidepressants for each individual patient together with consideration of all comorbidities and received therapies.

Bone architecture and the risk of osteoporosis can be assessed in a timely manner during therapy by measuring the bone density, determining vitamin D status in bones, as well as ionized calcium and other parameters of osteoporosis. Vitamin D is very important for physical and mental health. It is one of the key hormones in the regulation of bone metabolism. Vitamin D deficiency could increase the risk for low bone mineral density, or osteoporosis. Over recent years there have been studies that point out that vitamin D plays an important role in depression vulnerability [10, 15, 17]. It is interesting that an insufficient level of 25(OH)D (less than 50 nmol/l) could be associated with depressive disorder. Interestingly, lower level of vitamin D is found in 40–50% of depressive patients. This is probably due to lifestyle in depression. Depressive patients are very often heavy smokers, they sometimes abuse alcohol or other psychoactive substances. Such behavior may lead to hypovitaminosis [6, 19]. On the other hand, according to some studies, people with lower level of vitamin D could be at a greater risk of developing depressive disorders [20, 21]. Until now, the relationship between hypovitaminosis D and depression remains unclear. The association between low level of D vitamin and depression probably lies in homeostatic, trophic, and immunomodulatory effects of vitamin D [22, 23, 24]. New investigations also show that vitamin D receptors are identified in the same area of the brain associated with depression. In any case, hypovitaminosis D may represent an underlying biological vulnerability for depression [1, 17, 18, 23]. Although there is still no evidence that treatment with vitamin D supplements can reveal depressive symptoms, there is a possibility that some subgroup of depression may greatly

benefit from the treatment with vitamin D. Be as it may, it is clear that vitamin D has a prominent role in the treatment of depressive patients with low level of vitamin D. In such patients, long-term supplementation of vitamin D and calcium can increase bone mass and prevent fracture and long-term invalidity. Depressive patients with osteopenia/osteoporosis, benefit the most from the combination of adequate antidepressant therapy, psychotherapy, and, if indicated, therapy with vitamin D supplements. Also, there are studies suggesting that more intensive depression is associated with a lower level of vitamin D [23, 24]. In the present study, there was no relationship between the level of vitamin D and the severity of depressive symptoms measured with HAMD. In the future, an additional study should be made on a larger sample of patients to investigate if there is a connection between the low level of vitamin D in the blood and the severity of depressive symptoms.

The benefits as well as possible adverse effects of received therapy should be examined in every patient and useful advice should be given to them on their way of lifestyle and physical activity. In case it is necessary, bisphosphonates can be used for treating osteoporosis without any risk of interaction with SSRI antidepressants. Schweiger et al. [25] argue that osteoporosis of the spine has been diagnosed in almost 15% of the patients who suffer from depression – unipolar or bipolar – while numerous recent studies show a considerably higher incidence of depressive symptoms in women with vertebral and hip fractures [2, 8, 9, 25]. In the adolescent population, girls who suffer from anorexia and mood disorders are significantly more affected by osteoporosis than the general population. For the purpose of screening for osteoporosis, the National Osteoporosis Foundation recommends using the DEXA technique in women, aged 65 years and above and men aged 70 years and above or in people who are older than 50 years and are at an increased risk of fracture [1, 3, 4, 9, 10].

Hypercortisolemia in depression is meant to be the possible neurobiological base for such hypothesis [8, 9, 19]. Depression causes the activation of the hypothalamic–pituitary–adrenal axis and this alteration, which could be the crucial factor for the increased risk of osteoporosis in depressed patients [1, 10]. Actual hypothesis considers that corticotropin-releasing hormone and persisting high level of cortisol in depressed patients lead to secondary hypogonadism, which present one of the crucial risk factors for bone loss [1, 8, 14, 15, 21]. Such negative influence could be responsible for higher incidence of osteoporosis in patients with depressive symptoms, both in unipolar and bipolar affective disorders compared to the general population [1, 15, 17, 22]. According to the definition provided by the World Health Organization, osteoporosis is a progressive systemic skeletal disease characterized by reduced bone mineral density and bone microarchitecture alteration, which contributes to the risk of fracture and disability [1, 2]. The incidence of osteoporosis is 8–10%, but it is 10 times more common in women who reach menopause – the osteoporosis has been diagnosed in almost 22 million women and about 5.5 million men in the European Union [6]. The reduced bone mass and demineralization

cause the bone to lose its strength and elasticity, which significantly increases the risk of fracture even with minimal trauma. According to the majority of the world's statistics, it is considered that almost 70% of fractures occur due to osteoporotic bones. Bone demineralization occurs as a result of bone remodeling process due to increased catabolic processes (increased osteoclast function and reduced osteoblast function). The osteoporosis is diagnosed by using the bone densitometry or DEXA scan and bone mineral density (BMD) described as a T-score and Z-score. A T-score represents standard deviation of a patient's BMD from the average value of BMD of a person of the same sex and constitution aged 20–30 years. The T-score between -1 and -2.5 is classified as osteopenia and the score lower than -2.5 is classified as osteoporosis [3, 20].

In addition to depression being considered a risk factor for osteoporosis, a considerable controversy has been caused by the results of certain studies in the last 10 years that show a possibility that certain types of antidepressants can cause osteoporosis as well. This assumption refers to the SSRIs type of antidepressants [12]. The mechanism of the action of these “newer” antidepressants is based on preventing the reuptake of serotonin at the presynaptic membrane, which leads to an increase in the level of serotonin in the synaptic cleft and its reuptake by receptors located at the postsynaptic membrane. The SSRIs achieve their antidepressant effect by binding to the serotonin transporter (SERT) in the central nervous system. The possibility that these antidepressants trigger osteoporosis is based on the discovery that functional serotonin receptors such as SERT (for which SSRIs are bound to at allosteric sites) are identified in osteoblasts, osteoclasts, and osteocytes. The second possibility is that SSRIs lead to a decrease in testosterone levels and an increase in prolactin level in both sexes, which represents a risk for osteoporosis. The following risk factors have a significant effect on the increased incidence of osteoporosis: chronic diseases – thyroid and parathyroid function disorders, hypogonadism, Cushing's and Addison's disease, insulin-dependent diabetes, neurological disorders and digestive disorders. [1, 14, 19, 21, 22].

At risk from osteoporosis are not only the middle age and elderly patients, but also the adolescent population – girls who suffer from anorexia and mood disorders are significantly more affected by osteoporosis than the general population. With regard to undoubtedly common comorbidity of depression and osteoporosis, the measures for prevention and early diagnosis need to be undertaken in order to have early detection, treatment, and lowering of the complication and mortality rate of both disorders [1, 4, 10, 21, 22, 26].

## CONCLUSION

The results of this investigation indicate that a middle intensity depressive episode in bipolar affective disorder is connected with a higher prevalence of osteoporosis. Patients treated with SSRIs have higher prevalence of osteoporosis than patients treated with mood stabilizers only,



without antidepressants. In the future, larger cohorts of patients should be included in this kind of study.

Prevention is undoubtedly better than treatment of osteoporosis, which, in case of treatment of depressive symptoms, implies the selection of adequate group of medicines with respect to the age, initial status of the patient's

skeleton, especially in patients already diagnosed with osteoporosis or other familiar risk factors pertaining to its development.

**Conflict of interest:** None declared.

## REFERENCES

1. Rauma PH, Pasco JA, Berk M, Stuart AJ, Koivumaa Honkanen H, Honkanen RJ, et al. The association between major depressive disorder, use of antidepressants and bone mineral density in men. *J Musculoskelet Neuronal Interact.* 2015;15(2):177–85.
2. Warden JS, Robyn KF. Do selective serotonin reuptake inhibitors (SSRIs) cause fractures? *Curr Osteoporos Rep.* 2016;14(5):211–8.
3. Gojković Z, Matijević R, Harhaji V, Ilinčić B, Barišić Lj, Kupusinac A, et al. Trends in bone mineral density among nutritional status categories of Vojvodina elderly population. *Srp Arh Celok Lek.* 2020;148(9–10):577–83.
4. Wei-Sheng Lee C, Chun-Hui L, Cheng-Li L, Ji-An L, Fung-Chang S, Chia-Hung K. Increased risk of osteoporosis in patients with depression, a population-based retrospective cohort study. *Mayo Clin Proc.* 2015;90(1):63–70.
5. Saraykar S, John V, Cao B, Hnatow M, Ambrose CG, Rianon N. Association of selective serotonin reuptake inhibitors and bone mineral density in elderly women. *J Clin Densitom.* 2018;21(2):193–9.
6. Weng SF, Hsu HR, Weng YL, Tien KJ, Kao HY. Health-related quality of life and medical resource use in patients with osteoporosis and depression: a cross-sectional analysis from the national health and nutrition examination survey. *Int J Environ Res Public Health.* 2020;17(3):1124.
7. Turcotte AF, O'Connor S, Morin S, Gibbs J, Willie B, Jean S, et al. Association between obesity and risk of fracture, bone mineral density and bone quality in adults: a systematic review and meta-analysis. *PLoS One.* 2021;16(6):e0252487.
8. Wadhwa R, Kumar M, Talegaonkar S, Vohora D. Serotonin reuptake inhibitors and bone health: a review of clinical studies and plausible mechanisms. *Osteoporos Sarcopenia.* 2017;3(2):75–81.
9. Wang CY, Fu SH, Wang CI, Chen PJ, Wu FL, Hsiao FY. Serotonergic antidepressant use and the risk of fracture: a population-based nested case-control study. *Osteoporos Int.* 2016;27(1):57–63.
10. Ham AC, Aarts N, Noordam R, Rivadeneira F, Ziere G, Zillikens MC, et al. Use of selective serotonin reuptake inhibitors and bone mineral density change: a population-based longitudinal study in middle-aged and elderly individuals. *J Clin Psychopharmacol.* 2017;37(5):524–30.
11. Jia-Sheng N, Kok-Yong Ch. Potential mechanisms linking psychological stress to bone health. *Int J Med Sci.* 2021;18(3):604–14.
12. Skowrońska-Józwiak E, Gałęcki P, Głowacka E, Wojtyła C, Biliński P, Lewiński A. Bone Metabolism in Patients Treated for Depression. *Int J Environ Res Public Health.* 2020;17(13):4756.
13. Li S, Qui Y, Teng Z, Chen J, Kang D, Tang H, et al. Association between bipolar disorder and low bone mass: a cross-sectional study with newly diagnosed, drug-naïve patients. *Front Psychiatry.* 2020;11:530.
14. Cvjetković Bošnjak M, Okanović M, Nedić A, Vasić V, Šakić B. Depression, antidepressants and risk of osteoporosis. *GJRA.* 2019;8(1):194–5.
15. Bradaschia Correa V, Josephson A, Mehta D, Mizrahi M, Neibarth S, Liu C, et al. Fluoxetine inhibits osteoblast differentiation and mineralization in fracture healing. *J Bone Miner Res.* 2017;32(4):821–33.
16. Yusuf AA, Hu Y, Chandler D, Crittenden DB, Barron RL. Predictors of imminent risk of fracture in Medicare-enrolled men and women. *Arch Osteoporos.* 2020;15(1):120.
17. Diem SJ, Blackwell TL, Stone KL, Yaffe K, Haney EM, Bliziotes MM, et al. Use of antidepressants and rates of hip bone loss in older women: the study of osteoporotic fractures. *Arch Intern Med.* 2007;167(12):1240–5.
18. Barnsley J, Buckland G, Chan PE, Ong A, Ramos AS, Baxter M, et al. Pathophysiology and treatment of osteoporosis: challenges for clinical practice in older people. *Aging Clin Exp Res.* 2021;33(4):759–73.
19. Kelly R, McDonald L, Jensen N, Sidles S, LaRue A. Impacts of psychological stress on osteoporosis: clinical implications and treatment interactions. *Front Psychiatry.* 2019;10:200.
20. Bahouq H, Soulaymani A. Depression, quality of life, and self-esteem of Moroccan postmenopausal women with osteoporosis before the occurrence of fractures. *J Menopausal Med.* 2020;26(2):121–9.
21. Boskovic K, Cvjetkovic-Bosnjak M, Tomasevic-Todorovic S, Soldatovic-Stajic B, Zvekic-Svorcan J. Effect of selective serotonin reuptake inhibitors on the incidence of osteoporosis in patients with depression. *Balneoclimatologia.* 2013;39(1):281–5. (Article in Serbian)
22. Lanteigne A, Sheu YH, Sturmer T, Pate V, Azrael D, Swanson SA, et al. Serotonin-norepinephrine reuptake inhibitor and selective serotonin reuptake inhibitor use and risk of fractures: a new-user cohort study among US adults aged 50 years and older. *CNS Drugs.* 2015;29(3):245–52.
23. Milaneshi Y, Hoogendijk W, Lips P, Heijboer A, Schoevers R, Van Hemert AM, et al. The association between low vitamin D and depressive disorders. *Mol Psychiatry.* 2014;19(4):444–51.
24. Menon V, Kumar Kar S, Suthar N, Nebhinani N. Vitamin D and depression: a critical appraisal of the evidence and future directions. *Indian J Psychol Med.* 2020;42(1):11–21.
25. Schweiger U, Deuschle M, Körner A, Lammers CH, Schmider J, Gotthardt U, et al. Low lumbar bone mineral density in patients with major depression. *Am J Psychiatry.* 1994;151(11):1691–3.
26. Boskovic K, Kovacev-Zaic B, Cvjetkovic Bosnjak M, Tomasev-Todorovic S, Knezevic A. Secondary osteoporosis in patients with depression treated with selective serotonin reuptake inhibitors. *Osteoporos Int.* 2015;26(2):464.

## Повезаност биполарног афективног поремећаја, употребе антидепресива и остеопорозе

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### САЖЕТАК

**Увод/Циљ** Остеопороза је један од најчешћих коморбидних поремећаја међу болесницима са депресивним поремећајем.

Циљ овог истраживања је био да се процени повезаност употребе антидепресива са остеопорозом код болесника лечених од биполарног афективног поремећаја.

**Методе** Ова студија је обухватила 73 болесника, стара 50–72 године, оба пола, хоспитализована због депресивне епизоде биполарног афективног поремећаја, између 2016. и 2020. године, на Клиници за психијатрију Клиничког центра Војводине. Болесници су подељени у две групе: а) прва група (40) лечена је селективним инхибиторима поновног преузимања серотонина у комбинацији са стабилизатором расположења (литијум-карбонат/ламотригин), б) друга група (33) третирана је само стабилизатором расположења. Студија је такође обухватила и две контролне групе испитаника. Клиничка мерења минералне густине кости на лумбалној кичми и куку изведена су методом апсорпциометрије рендгенских зрака

двоструке енергије. Измерени су нивои калцијума и Де-витамина из узорака крви болесника. Подаци су статистички обрађени анализом варијансе и Краскал–Волисовим тестом.

**Резултати** Остеопороза је регистрована код 25% болесника у првој и код 18% болесника у другој групи, док је остеопенија установљена код 40% у првој и код 37% болесника у другој групи. Постоји статистички значајна разлика у вредностима нивоа *CrossLaps* и нивоа *25(OH)* Де-витамина, као и у заступљености остеопорозе и остеопеније у односу на контролне групе.

**Закључак** Депресивна епизода у биполарном афективном поремећају је повезана са већом преваленцом остеопорозе у односу на контролну групу. Болесници лечени селективним инхибиторима поновног преузимања серотонина имају већу преваленцу остеопорозе у односу на болеснике лечене само стабилизатором расположења.

**Кључне речи:** минерална густина кости; депресија; селективно преузимање серотонина