Associations between Metabolic Syndrome and Bone Mineral Density and Trabecular Bone Score in Postmenopausal Women with Non-Vertebral Fractures

Vladyslav Povoroznyuk, Larysa Martynyuk, Iryna Syzonenko, Liliya Martynyuk

Abstract—Medical, social, and economic relevance of osteoporosis is caused by reducing quality of life, increasing disability and mortality of the patients as a result of fractures due to the low-energy trauma. This study is aimed to examine the associations of metabolic syndrome components, bone mineral density (BMD) and trabecular bone score (TBS) in menopausal women with non-vertebral fractures. 1161 menopausal women aged 50-79 year-old were examined and divided into three groups: A included 419 women with increased body weight (BMI - 25.0-29.9 kg/m²), B – 442 females with obesity (BMI >29.9 kg/m²)i and C – 300 women with metabolic syndrome (diagnosis according to IDF criteria, 2005). BMD of lumbar spine (L1-L4), femoral neck, total body and forearm was investigated with usage of dual-energy X-ray absorptiometry. The bone quality indexes were measured according to Med-Imaps installation. All analyses were performed using Statistical Package 6.0. BMD of lumbar spine (L1-L4), femoral neck, total body, and ultradistal radius was significant higher in women with obesity and metabolic syndrome compared to the pre-obese ones (p<0.001). TBS was significantly higher in women with increased body weight compared to obese and metabolic syndrome patients. Analysis showed significant positive correlation between waist circumference, triglycerides level and BMD of lumbar spine and femur. Significant negative association between serum HDL level and BMD of investigated sites was established. The TBS (L1-L4) indexes positively correlated with HDL (high-density lipoprotein) level. Despite the fact that BMD indexes were better in women with metabolic syndrome, the frequency of non-vertebral fractures was significantly higher in this group of patients.

Keywords—Bone mineral density, trabecular bone score, metabolic syndrome, fracture.

I. INTRODUCTION

OSTEOPOROSIS and metabolic syndrome are important public health problems, due to the decreasing of quality and reducing life expectancy of patients as a result of lowtrauma fractures in a case of osteoporosis and the possibility of cardiovascular, endocrine and other complications in a case of metabolic syndrome development [1], [5]. The frequency of

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both diseases increases with age of patient and duration of menopausal period as a result of slowdown in metabolism and estrogen deficiency development [8], [11].

Traditionally, osteoporosis is diagnosed according to the history of low-energy fractures or the results of BMD (T-score), which are determined by using X-ray densitometry [10]; however, BMD provides only 70-75% of bone strength [13]. Other factors that affect it include state of cortical bone macro-geometry and trabecular bone micro-architecture, presence damages and cracks in it, which can be calculated by index TBS, patented by MED-I maps (m. Bordeaux, France) in 2006 [3]. In our opinion, evaluation of TBS is important to perform this work.

Scientists paid much attention to the study of the relationships between metabolic syndrome and osteoporosis. Abdominal obesity, high glucose (as a result of insulin deficiency or insulin resistance), high triglycerides and low high density lipoproteins which are the main components of metabolic syndrome have significant impact on bone tissue and fractures development but published research results are contradictory [4], [7], [12], [14], [16]. The discrepancy of opinions prompted this investigation.

The aim of our study was to evaluate the relationships between metabolic syndrome components and BMD, TBS in postmenopausal women with low-trauma non-vertebral fractures.

II. MATERIALS AND METHODS

The study involved 1161 postmenopausal women aged 50-79 years (mean age - 63.977±7.961 years; mean body mass index (BMI) - 31.587 \pm 4.739 kg/m²; mean waist circumference - 92.524±11.466 cm; mean duration of menopause period - 13.858±8.014). Patients were divided into three groups: A - 419 women with increase body weight (preobese) defined on the basis of WHO criteria, [15] BMI 25.0-29.9 kg/m² (mean age $- 63.983 \pm 8.283$ years; mean BMI - 27.547 ± 1.906 kg/m²; mean waist circumference - $79.995\pm$ 4.511 cm; mean duration of menopause period - 13.809± 8004), B - 442 women with obesity - BMI \geq 30.0 kg/m² (mean age - 63.884±7.619 years; mean BMI - 34.418±3.864 kg/m²; mean waist circumference -100.464 ± 6.726 cm; mean duration of menopause period - 13.627±7.847), C - 300 females with metabolic syndrome (diagnosed according to the

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International Diabetic Federation criteria of 2005) [6] (mean age $- 64.163\pm8.017$; mean BMI $- 33.058\pm4.812$ kg/m²; mean waist circumference $- 98.323\pm8.244$ cm; mean duration of menopause period $- 14.267\pm8.280$). Additionally, groups were divided according to the presence of low-trauma non-vertebral fractures (202 female had them in history (NVF) and 959 women were without fractures (WF) in the past).

DXA method (Prodigy, GE Medical systems, Lunar, Madison, WI, USA, 2005) was used to investigate BMD of lumbar spine (L1-L4), femoral neck, total body and forearm.

TBS (L1-L4) was estimated by installation of TBS iNsight[®] software (Med-Imaps, Pessac, France) program on DXA machine.

The Statistical Package 6.0 ©StatSoft, Inc. was used for analyses. Continuous variables were reported as mean \pm SD. Pearson correlations examined the relationship between continuous variables, significance set at p<0.05.

III. RESULTS

We found that women with increased body weight have a significantly lower BMD of lumbar spine (A $- 0.986\pm0.178$ g/cm², B $- 1.109\pm0.181$ g/cm²; C $- 1.120\pm0.199$ g/cm²; F= 63.814; p<0.001); femoral neck (A $- 0.809\pm0.126$ g/cm², B $- 0.872\pm0.134$ g/cm²; C $- 0.880\pm0.149$ g/cm²;F=32.097; p< 0.001), total body (A $- 0.876\pm0.139$ g/cm², B $- 0.968\pm0.146$ g/cm²; C $- 0.978\pm0.160$ g/cm²;F=57.366; p<0.001) and ultradistal forearm (A $- 0.378\pm0.076$ g/cm², B $- 0.436\pm0.080$ g/cm²; C $- 0.435\pm0.088$ g/cm²;F=67.582; p<0.001) compared to women with obesity and metabolic syndrome. The bone tissue quality (TBS L1-L4) was significantly higher in women with increased weight in comparison with metabolic syndrome female (A $- 1.188\pm0.151$, B $- 1.169\pm0.163$; C $- 1.157\pm0.173$ g/cm²; F=3.479; p<0.05).

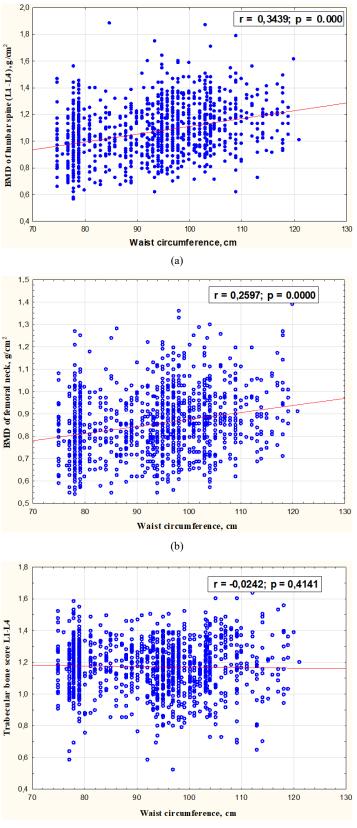
BMD of lumbar spine (L1-L4) was significantly higher in patients of groups B and C without fractures (Table I). BMD of femoral neck was significantly lower in female with obesity and non-vertebral fractures (Table II). BMD of total body and ultradistal radius significantly better in all groups of women without fractures compared to patients with non-vertebral fractures (Tables III and IV). TBS (L1-L4) was significantly higher in patients without fractures in the groups of women with increased body weight and obesity (p < 0.05) (Table V).

The analysis of the metabolic syndrome laboratory components (serum triglycerides and HDL indexes) was carried out. We established significantly higher triglycerides level (A $- 1.049\pm0.381$ g/cm², B $- 1.030\pm0.322$ g/cm²; C $- 1.605\pm0.703$ g/cm²; F=162.669; p<0.001) and significantly lower HDL level (A $- 1.531\pm0.372$ g/cm², B $- 1.509\pm0.314$ g/cm²; C $- 1.170\pm0.256$ g/cm²; F=126.832; p<0.001) in patients with metabolic syndrome. There was no difference of triglycerides level in female with non-vertebral fractures and without them in all investigated groups (Table VI). The level of HDL was significantly lower in patients with non-vertebral fractures and metabolic syndrome (Table VII).

BMD OF LUMBAR SPINE (L1-L4) IN GROUPS OF WOMEN				
Groups of	Subgroups of patients	BMD lumbar spin	e p	
patients		(L1-L4), g/cm ²	Р	
А	without fractures (n=358)	0.992±0.180	>0.05	
	with non-vertebral fractures (n=61)	0.955±0.164		
В	without fractures (n=365)	1.119 ± 0.183	< 0.001	
Б	with non-vertebral fractures (n=77)	1.059 ± 0.163	~0.001	
С	without fractures (n=236)	1.139 ± 0.199	<0.001	
C	with non-vertebral fractures (n=64)	1.052 ± 0.860	< 0.001	
	TABLE II			
	BMD OF FEMORAL NECK IN GROUPS			
Groups of patients	Subgroups of patients	BMD femoral neck, g/cm ²	Р	
patients	without fractures (n=358)	0.814±0.130		
А		0.780 ± 0.101	>0.05	
	with non-vertebral fractures $(n=61)$			
В	without fractures (n=365)	0.879±0.135	< 0.001	
	with non-vertebral fractures (n=77)	0.834 ±0.119		
С	without fractures (n=236)	0.888±0.153	>0.05	
	with non-vertebral fractures (n=64)	0.852±0.133	0.02	
	TABLE III BMD of Total Body in Groups	OF WOMEN		
Groups of	BWD OF TOTAL BODT IN GROUPS	BMD total		
patients	Subgroups of patients	body, g/cm^2	Р	
punonio	without fractures (n=358)	0.884±0.140		
А	with non-vertebral fractures $(n=61)$	0.832±0,119	< 0.001	
	without fractures (n=365)	0.977±0.148		
В	with non-vertebral fractures (n=77)	0.977 ± 0.148 0.928 ± 0.129	< 0.001	
С	without fractures (n=236)	0.991±0.163	< 0.001	
	with non-vertebral fractures (n=64)	0.931±0.140		
	TABLE IV			
В	TABLE IV MD of Ultradistal Radius in Gro	UPS OF WOMEN		
	MD of Ultradistal Radius in Gro	UPS OF WOMEN BMD ultradistal		
B Groups of patients			Р	
Groups of patients	MD of Ultradistal Radius in Gro	BMD ultradista	Р	
Groups of	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients	BMD ultradistal radius, g/cm ²	^l P <0.05	
Groups of patients A	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61)	BMD ultradistal radius, g/cm ² 0.382±0.077	P <0.05	
Groups of patients	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365)	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079	Р	
Groups of patients A	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77)	BMD ultradistai radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083	P <0.05	
Groups of patients A	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26)	BMD ultradistai radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088	P <0.05	
Groups of patients A B	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77)	BMD ultradistai radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083	P <0.05 <0.001	
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Groups of patients A B C	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64)	BMD ultradistai radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842	P <0.05 <0.001	
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Groups of patients A B C	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients	BMD ultradistai radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 OMEN TBS (L1-L4)	P <0.05 <0.001	
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Groups of patients A B C Groups of patients A B C	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=236) with metabolic syndrome (n=73) TABLE VI TRIGLYCERIDES LEVEL IN GROUP Subgroups of patients	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 0MEN TBS (L1-L4) 1.194±0.151 1.152±0.439 1.179±0,150 1.119±0.171 1.156±0.177 1.158±0.156 S OF WOMEN triglycerides level, mmol/l	P <0.05 <0.001 <0.001 P <0.05 <0.001 >0.05	
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Groups of patients A B C Groups of patients A B C Groups of patients	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=236) with non-vertebral fractures (n=77) without fractures (n=236) with metabolic syndrome (n=73) TABLE VI TRIGLYCERIDES LEVEL IN GROUP Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61)	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 0MEN TBS (L1-L4) 1.194±0.151 1.152±0.439 1.179±0,150 1.119±0.171 1.156±0.177 1.158±0.156 S OF WOMEN triglycerides level, mmol/l	P <0.05 <0.001 <0.001 P <0.05 <0.001 >0.05 P	
Groups of patients A B C Groups of patients A B C Groups of patients A	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=236) with non-vertebral fractures (n=77) without fractures (n=236) with metabolic syndrome (n=73) TABLE VI TRIGLYCERIDES LEVEL IN GROUP Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365)	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 0MEN TBS (L1-L4) 1.194±0.151 1.152±0.439 1.179±0.150 1.119±0.171 1.156±0.177 1.158±0.156 s OF WOMEN triglycerides level, mmol/1 1.057±0.393	P <0.05 <0.001 <0.001 P <0.05 <0.001 >0.05 P >0.05	
Groups of patients A B C Groups of patients A B C Groups of patients	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=236) with non-vertebral fractures (n=77) without fractures (n=236) with metabolic syndrome (n=73) TABLE VI TRIGLYCERIDES LEVEL IN GROUP Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61)	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 0MEN TBS (L1-L4) 1.194±0.151 1.152±0.439 1.179±0,150 1.119±0.171 1.156±0.177 1.158±0.156 s OF WOMEN triglycerides level, mmol/1 1.057±0.393 0.996±0.288	P <0.05 <0.001 <0.001 P <0.05 <0.001 >0.05 P	
Groups of patients A B C Groups of patients A B C Groups of patients A B C Groups of patients A B B B B	MD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=236) with non-vertebral fractures (n=77) without fractures (n=236) with metabolic syndrome (n=73) TABLE VI TRIGLYCERIDES LEVEL IN GROUP Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365)	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 0MEN TBS (L1-L4) 1.194±0.151 1.152±0.439 1.179±0,150 1.119±0.171 1.156±0.177 1.158±0.156 s OF WOMEN triglycerides level, mmol/1 1.057±0.393 0.996±0.288 1.035±0.332	P <0.05 <0.001 <0.001 P <0.05 <0.001 >0.05 P >0.05 >0.05	
Groups of patients A B C Groups of patients A B C Groups of patients A	SMD OF ULTRADISTAL RADIUS IN GRO Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=26) with non-vertebral fractures (n=64) TABLE V TBS (L1-L4 IN GROUPS OF W Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=365) with non-vertebral fractures (n=77) without fractures (n=236) with non-vertebral fractures (n=77) without fractures (n=358) with non-vertebral fractures (n=77) without fractures (n=358) with metabolic syndrome (n=73) TABLE VI TRIGLYCERIDES LEVEL IN GROUP Subgroups of patients without fractures (n=358) with non-vertebral fractures (n=61) without fractures (n=358)	BMD ultradistal radius, g/cm ² 0.382±0.077 0.354±0.071 0.442±0.079 0.410±0.083 0.442±0.088 0.407±0.842 0MEN TBS (L1-L4) 1.194±0.151 1.152±0.439 1.179±0,150 1.119±0.171 1.156±0.177 1.158±0.156 s OF WOMEN triglycerides level, mmol/1 1.057±0.393 0.996±0.288 1.035±0.332 1.006±0.273	P <0.05 <0.001 <0.001 P <0.05 <0.001 >0.05 P >0.05	

TABLE I BMD of Lumbar Spine (L1-L4) in Groups of Women

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(c)

Fig. 1 Correlation between waist circumference and BMD of (A) lumbar spine (L1-L4), (B) femoral neck, (C) TBS

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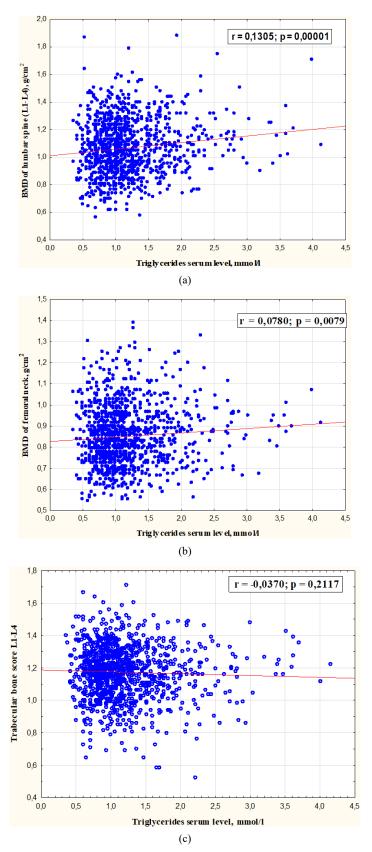


Fig. 2 Correlation between triglycerides serum level and BMD of (A) lumbar spine (L1-L4), (B) femoral neck, (C) TBS

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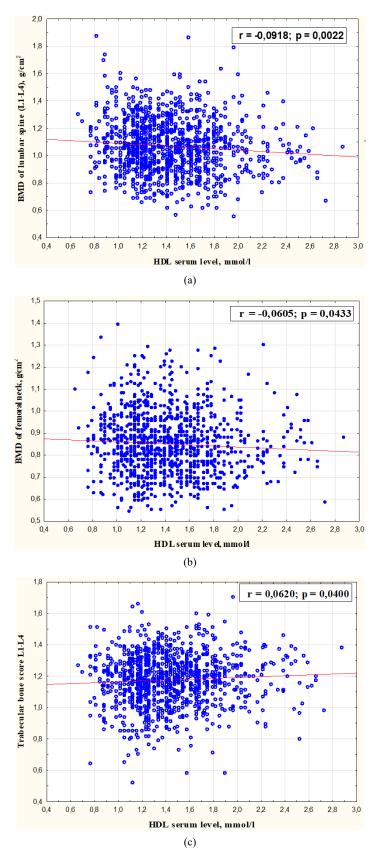


Fig. 3 Correlation between HDL serum level and BMD of (A) lumbar spine (L1-L4), (B) femoral neck, (C) TBS

HDL LEVEL IN GROUPS OF WOMEN				
Groups of patients	Subgroups of patients	HDL level, mmol/l	Р	
А	without fractures (n=358)	$1.523 {\pm} 0.037$	>0.05	
	with non-vertebral fractures (n=61)	1.579 ± 0.327		
В	without fractures (n=365)	1.501 ± 0.319	>0.05	
	with non-vertebral fractures (n=77)	1.546 ± 0.272		
С	without fractures (n=236)	1.148 ± 0.229	< 0.001	
	with non-vertebral fractures (n=64)	1.252 ± 0.325		

TABLE VII

In analysis of metabolic syndrome components, the waist circumference component was positively associated with BMD of lumbar spine and femur (Fig. 1). The study reveals significant positive correlation between serum triglycerides level and both investigated BMD sites (Fig. 2). A number of investigators have suggested relationship in accordance with our own findings [2]. It was found a significant positive correlation between HDL serum level and TBS and inversely association with BMD of lumbar spine and femur (Fig. 3).

We calculated the percentage of non-vertebral fractures in anamnesis (Fig. 4).

Low-trauma non-vertebral fractures occurred in 14.6% female with increased body weight, 17.4% of women with obesity and 21.3% of patients with metabolic syndrome.

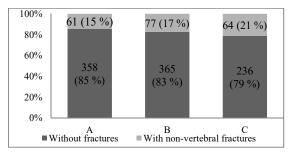


Fig. 4 Frequency of low-trauma vertebral fractures in women with increase bode weight (A), obesity (B) and with metabolic syndrome (C)

Significant differences were not found in the frequency of non-vertebral fractures in the groups of women with obesity and increased body weight or metabolic syndrome (X2=1.312, p>0.05 and X2=1.780, p>0.05, respectively), but it was significant in the groups of pre-obese female and patients with metabolic syndrome (X2=5.590, p<0.05). The similar results were found by other investigators [9].

IV. CONCLUSION

Menopausal women with obesity and metabolic syndrome have a significantly higher BMD at all measured sites compared to females with pre-obesity. TBS is significantly lower in women with non-vertebral fractures and increased body weight or obesity. A significant positive correlation is established between waist circumference, triglycerides level and BMD of lumbar spine and femoral neck. Correlation between HDL level and BMD at all levels is significant and negative. At the same time, it is positively associated with

TBS indexes. There is no significant difference in frequency of low-trauma non-vertebral fractures in the groups of preobese and obese women. At the same time, the incidence of osteoporotic non-vertebral fractures is significantly higher in female with metabolic syndrome in compared to other patients. Metabolic syndrome may not protect from any type of fractures, but future investigations are necessary.

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