

Are the European Society of Cardiology guidelines on lipid-lowering treatment implemented in morbidly obese patients qualified for bariatric surgery?

Jan Bylica^{1,2}, Piotr Major³, Tomasz Grodzicki¹, Maria Fornal¹

¹Department of Internal Medicine and Gerontology, Jagiellonian University Medical College, Kraków, Poland

²Doctoral School of Medical and Health Sciences, Jagiellonian University Medical College, Kraków, Poland

³2nd Department of General Surgery, Faculty of Medicine, Jagiellonian University Medical College, Kraków, Poland

Correspondence to:

Jan Bylica, MD,
Department of Internal Medicine
and Gerontology,
University Hospital in Krakow,
Jakubowskiego 2,
30-688 Kraków, Poland,
phone: +48 69 741 89 79,
e-mail:
jan.bylica@doctoral.uj.edu.pl

Copyright by the Author(s), 2023

DOI: 10.33963/KPa2023.0120

Received:

March 203, 2023

Accepted:

May 2, 2023

Early publication date:

May 19, 2023

INTRODUCTION

The primary aims of atherosclerotic cardiovascular disease (ASCVD) prevention are to reduce morbidity and mortality as well as increase life expectancy [1]. An important aspect of this prevention is control of apo-B-containing lipoproteins, which contribute to the induction and progression of ASCVD [1]. Prompt diagnosis and implementation of lipid-lowering treatment at every stage of dyslipidemia result in a better prognosis and lowers the risk of cardiovascular events [1, 2].

Adiposity is another major risk factor of ASCVD. This condition increases the risk of developing cardiovascular disease (CVD) by promoting dyslipidemia, diabetes mellitus (DM), and other disorders [1]. According to recent reports, 12.5% of the world's population is obese [3]. This number is twice as great in Poland [4].

We aimed to assess physicians' adherence to the latest guidelines on primary CVD prevention. To address this problem, we focused on the pharmacological treatment of dyslipidemia in morbidly obese patients before bariatric surgery. Our objective was to assess CVD risk, indications for hypolipidemic agents' introduction, and their efficacy in this group of patients.

METHODS

We enrolled consecutive patients 35 years of age or older, with morbid obesity, who had been admitted to the surgery ward to undergo bariatric surgery and agreed to participate in the study. The recruitment took place between December 2021 and April

2022. The exclusion criteria were secondary CVD prevention, chronic kidney disease, and familial hypercholesterolemia.

Medical records were used to characterize patients.

The study protocol conformed to the ethical guidelines of the 1975 Declaration of Helsinki and was approved by the institutional ethics committee. Informed consent was obtained from each study participant.

IBM SPSS Statistics for Windows, Version 28.0. (Armonk, NY: IBM Corp) software was used for statistical analysis. Continuous data values were presented as mean (standard deviation [SD]) or median (interquartile range [IQR]), and qualitative data as numbers and percentages. The χ^2 test and Fisher's test were used to compare qualitative data, while Student's t-test (for data with normal distribution) and the Mann-Whitney U-test (for variables with other than normal distribution) were used for quantitative data. Continuous variables were first checked for normal distribution with the Shapiro-Wilk test. For all tests, a *P*-value less than 0.05 was considered significant. A detailed description of the study methods is presented in Supplementary material.

RESULTS AND DISCUSSION

We enrolled 57 patients with morbid obesity (*n* = 41; 71.9% women) at a median (IQR) age of 46 (38–51) years. More than one-fourth (*n* = 15; 26.3%) had DM. Detailed basic characteristics are presented in Supplementary material, *Table S1*. The mean (SD) non-high-density lipoprotein cholesterol (non-HDL-C) level

Table 1. Comparison between hypolipidemic treatment groups in terms of basic characteristics and selected parameters

	Did not require lipid-lowering treatment (n = 30)	Required lipid-lowering treatment (n = 27)	P-value
Age, years, median (IQR)	45.50 (39.00–51.00)	48.00 (37.00–52.00)	0.88
Female sex, n (%)	25 (83.3)	16 (59.3)	0.04
BMI, kg/m ² , median (IQR)	40.41 (37.13–46.65)	42.52 (38.57–47.40)	0.24
Maximal noted body weight, kg, median (IQR)	126.50 (108.00–141.00)	130.00 (120.00–164.00)	0.23
Waist circumference, cm, median (IQR)	111.00 (108.00–128.00)	125.50 (114.00–132.00)	0.01
Hip circumference, cm, median (IQR)	132.00 (121.00–145.00)	133.50 (122.00–139.00)	0.64
Waist/hip ratio, mean (SD)	0.89 (0.02)	0.93 (0.02)	0.09
Active smoker, n (%)	0 (0)	10 (37.0%)	<0.001 ^a
SBP, mm Hg, mean (SD)	132.81 (2.18)	137.26 (2.73)	0.20
DBP, mm Hg, mean (SD)	82.98 (1.61)	84.19 (1.37)	0.57
CVD risk, %, median (IQR)	2 (1–2) ^b	4 (3–5.25) ^c	<0.001
ALT, U/l, median (IQR)	41.00 (28.00–59.00)	56.00 (46.00–77.00)	0.02
Total cholesterol mmol/l, mean (SD)	4.18 (0.16)	4.55 (0.14)	0.09
HDL-C mmol/l, median (IQR)	1.15 (1.07–1.37)	1.11 (1.01–1.21)	0.24
Non-HDL-C, mmol/l, mean (SD)	2.94 (0.15)	3.41 (0.14)	0.02
LDL-C, mmol/l, mean (SD)	2.49 (0.14)	2.85 (0.11)	0.05
TG, mmol/l, median (IQR)	1.27 (0.96–1.59)	1.31 (0.92–1.56)	0.94
Glucose, mmol/l, median (IQR)	5.53 (5.01–6.29)	5.76 (5.18–6.64)	0.37
HbA1c, %, median (IQR)	5.60 (5.40–5.90)	5.90 (5.60–6.40)	0.02
CK, U/l, median (IQR)	211.50 (152.00–301.00)	203.00 (151.00–260.00)	0.81
Creatinine, μmol/l, mean (SD)	71.82 (1.99)	72.14 (3.13)	0.93
eGFR, ml/min/1.73 m ² , mean (SD)	88.24 (3.25)	90.85 (3.50)	0.59

Abbreviations: ALT, alanine aminotransferase; BMI, body mass index; CK, creatine kinase; CVD, cardiovascular disease; DBP, diastolic blood pressure; GFR, glomerular filtration rate; HbA1c, glycated hemoglobin; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; Non-HDL-C, non-high-density lipoprotein cholesterol; SBP, systolic blood pressure; TG, triglycerides

^aMore than 20% of expected counts were <5. ^bCVD risk calculated for 27 individuals. ^cCVD risk calculated for 17 individuals

was 3.16 (0.11) mmol/l, low-density lipoprotein cholesterol (LDL-C) 2.66 (0.09) mmol/l, median (IQR) serum concentration of triglycerides (TG) was 1.28 (0.94–1.56) mmol/l, total cholesterol 4.3 (3.8–4.9) mmol/l. Ten patients (17.5%) were smokers or have quitted during the last stage of preparation for surgery (≤3 months before surgery). Half of the patients had a high CV risk (n = 29; 50.9%), followed by low-moderate (n = 26; 45.6%), and very high (n = 2; 3.5%). More than one-fifth (n = 12; 21.1%) of study participants had hypercholesterolemia diagnosed before enrollment. Of these, ten subjects (17.5%) had some sort of pharmacological treatment. Most of them were statins-only users (n = 6; 60% of patients on hypolipidemic treatment), two (20%) were on fibrates, one (10%) had statin and fibrate, and one (10%) took statin and ezetimibe. However, only one of those patients had the LDL-C level within normal limits, while all others could have their treatment intensified. Two patients (3.5%, with dyslipidemia diagnosed earlier) had no treatment introduced at the time it was required.

Dyslipidemia was newly diagnosed in other 15 patients. Altogether, almost half of the study population (n = 27; 47.4%) had hypercholesterolemia but was on no or insufficient treatment. This number included 5 (8.8%) participants with hypertriglyceridemia. None of the previously mentioned patients met the exclusion/discontinuation criterion for statin treatment, which is creatine kinase >4 times the upper limit of normal, and only 4 (14.8% of those in need of lipid-lowering therapy) could take these

drugs because of their alanine aminotransferase levels, which were >3 times the upper limit of normal [2].

As shown in **Table 1**, the patients who required dyslipidemia treatment had higher levels of alanine aminotransferase, non-HDL-C, and LDL-C. Moreover, the group that needed hypolipidemic treatment consisted mostly of men.

Although there is a large number of previously published articles on lipid profile and pharmacological lipid-lowering treatment in morbidly obese patients before bariatric surgery, they do not provide data on undiagnosed lipid metabolism disorders or untreated patients [7–9]. Our study is the first to provide this information.

In our work, the percentage of morbidly obese patients with hypercholesterolemia before bariatric surgery was comparable to other research, although the percentage of our participants with hypertriglyceridemia was significantly lower compared to 24% and 53% reported in the literature [5, 8, 9]. This discrepancy might partially result from the diagnostic criterion we used when setting the threshold value of TG: 2.3 mmol/l [1].

The average levels of LDL-C and TG presented in our article (2.66 and 1.28 mmol/l, respectively) are lower than the values reported in a similar study from Austria (3.12 and 1.78 mmol/l) [6], Singapore (3.05 and 1.66) [7], and the US (2.88 and 1.74 mmol/l) [10]. Conversely, serum levels of non-HDL-C in the Austrian study (3.9), and total cholesterol shown in the Singaporean investigation (4.9 mmol/l) were lower than those quoted in our article (3.16 and 4.3 mmol/l,

respectively). Furthermore, the perioperative statin user ratio observed in our study was much lower in comparison to the results reported in American publications (10.5% vs. 24.5%, and 34%) [10, 12].

Our investigations, showing insufficiencies in the pharmacological treatment of dyslipidemia in morbidly obese patients before bariatric surgery, have some limitations. One factor that might have influenced the outcomes was the COVID-19 pandemic, which reduced healthcare accessibility. Another reason may be the lack of patient adherence to prescribed medications. In any case, our results should encourage all physicians involved in treating morbidly obese patients to actively screen them for dyslipidemia and regularly evaluate effectiveness of lipid-lowering treatment. There are three main reasons why the above-mentioned checks should be performed particularly in individuals qualified for bariatric surgery. First, the process of preparing for surgery should last 6–12 months, so there is plenty of time to introduce effective treatment before, hypothetically, more effective intervention [13]. Second, quite a considerable number, reaching 40% of post-bariatric patients, do not demonstrate dyslipidemia remission. What is more, its relapse rate may come up to 24% [14]. Lastly, patients on preoperative statins may have a higher DM rate and hypertriglyceridemia remission [11,14].

The main limitation of our research is the fact that the study was conducted in one bariatric center, and the enrolled group was relatively small. Moreover, we did not have data on target organ damage in DM patients, which might have impaired their CV risk assessment. Nevertheless, if we had possessed this information, we could only have assigned such patients to the higher CV risk group, which would support the outcomes.

In conclusion, our study is the first to show that morbidly obese patients before bariatric surgery may be underdiagnosed and undertreated for dyslipidemia. These findings, if confirmed by further research involving a larger number of clinical centers, would indicate a need for revisiting clinical practices applied to patients qualified for bariatric surgery.

Supplementary material

Supplementary material is available at https://journals.viamedica.pl/kardiologia_polska.

Article information

Conflict of interest: None declared.

Funding: This work was supported by the Jagiellonian University Medical College grant no. N41/DBS/000621.

Open access: This article is available in open access under Creative Commons Attribution-Non-Commercial-No Derivatives 4.0 International (CC BY-NC-ND 4.0) license, which allows downloading and sharing articles with others as long as they credit the authors and the publisher, but without permission to change them in any way or use them commercially. For commercial use, please contact the journal office at kardiologiapolska@ptkardio.pl.

REFERENCES

1. Visseren FLJ, Mach F, Smulders YM, et al. 2021 ESC Guidelines on cardiovascular disease prevention in clinical practice. *Eur Heart J*. 2021; 42(34): 3227–3337, doi: [10.1093/eurheartj/ehab484](https://doi.org/10.1093/eurheartj/ehab484), indexed in Pubmed: [34458905](https://pubmed.ncbi.nlm.nih.gov/34458905/).
2. Mach F, Baigent C, Catapano AL, et al. 2019 ESC/EAS Guidelines for the management of dyslipidaemias: lipid modification to reduce cardiovascular risk. *Eur Heart J*. 2020; 41(1): 111–188, doi: [10.1093/eurheartj/ehz455](https://doi.org/10.1093/eurheartj/ehz455), indexed in Pubmed: [31504418](https://pubmed.ncbi.nlm.nih.gov/31504418/).
3. Chooi YuC, Ding C, Magkos F. The epidemiology of obesity. *Metabolism*. 2019; 92: 6–10, doi: [10.1016/j.metabol.2018.09.005](https://doi.org/10.1016/j.metabol.2018.09.005), indexed in Pubmed: [30253139](https://pubmed.ncbi.nlm.nih.gov/30253139/).
4. Stepaniak U, Micek A, Waśkiewicz A, et al. Prevalence of general and abdominal obesity and overweight among adults in Poland. Results of the WOBASZ II study (2013–2014) and comparison with the WOBASZ study (2003–2005). *Pol Arch Med Wewn*. 2016; 126(9): 662–671, doi: [10.20452/pamw.3499](https://doi.org/10.20452/pamw.3499), indexed in Pubmed: [27535012](https://pubmed.ncbi.nlm.nih.gov/27535012/).
5. Buchwald H, Avidor Y, Braunwald E, et al. Bariatric surgery. *JAMA*. 2004; 292(14): 1724, doi: [10.1001/jama.292.14.1724](https://doi.org/10.1001/jama.292.14.1724).
6. Kruschitz R, Wakolbinger M, Schindler K, et al. Effect of one-anastomosis gastric bypass on cardiovascular risk factors in patients with vitamin D deficiency and morbid obesity: A secondary analysis. *Nutr Metab Cardiovasc Dis*. 2020; 30(12): 2379–2388, doi: [10.1016/j.numecd.2020.08.011](https://doi.org/10.1016/j.numecd.2020.08.011), indexed in Pubmed: [32981799](https://pubmed.ncbi.nlm.nih.gov/32981799/).
7. Panday VB, Shabbir A, Kuntjoro I, et al. Long-term effects of bariatric surgery on cardiovascular risk factors in Singapore. *Singapore Med J*. 2021; 62(9): 472–475, doi: [10.11622/smedj.2020047](https://doi.org/10.11622/smedj.2020047), indexed in Pubmed: [32241069](https://pubmed.ncbi.nlm.nih.gov/32241069/).
8. Milone M, Lupoli R, Maietta P, et al. Lipid profile changes in patients undergoing bariatric surgery: a comparative study between sleeve gastrectomy and mini-gastric bypass. *Int J Surg*. 2015; 14: 28–32, doi: [10.1016/j.ijso.2014.12.025](https://doi.org/10.1016/j.ijso.2014.12.025), indexed in Pubmed: [25576760](https://pubmed.ncbi.nlm.nih.gov/25576760/).
9. Buchwald H, Estok R, Fahrbach K, et al. Trends in mortality in bariatric surgery: a systematic review and meta-analysis. *Surgery*. 2007; 142(4): 621–32; discussion 632, doi: [10.1016/j.surg.2007.07.018](https://doi.org/10.1016/j.surg.2007.07.018), indexed in Pubmed: [17950357](https://pubmed.ncbi.nlm.nih.gov/17950357/).
10. Coleman KJ, Basu A, Barton LJ, et al. Remission and relapse of dyslipidemia after vertical sleeve gastrectomy vs roux-en-y gastric bypass in a racially and ethnically diverse population. *JAMA Netw Open*. 2022; 5(9): e2233843, doi: [10.1001/jamanetworkopen.2022.33843](https://doi.org/10.1001/jamanetworkopen.2022.33843), indexed in Pubmed: [36169953](https://pubmed.ncbi.nlm.nih.gov/36169953/).
11. Perna M, Baker M, Byrne TK, et al. Statins and the bariatric patient: characterization and perioperative effects of statin therapy in the gastric bypass patient. *Am Surg*. 2011; 77(1): 44–47, indexed in Pubmed: [21396304](https://pubmed.ncbi.nlm.nih.gov/21396304/).
12. Kennedy AL, Nelson T, Pettine S, et al. Medication use following bariatric surgery: factors associated with early discontinuation. *Obes Surg*. 2014; 24(5): 696–704, doi: [10.1007/s11695-013-1131-8](https://doi.org/10.1007/s11695-013-1131-8), indexed in Pubmed: [24234778](https://pubmed.ncbi.nlm.nih.gov/24234778/).
13. Carter J, Chang J, Birriel TJ, et al. ASMBS position statement on preoperative patient optimization before metabolic and bariatric surgery. *Surg Obes Relat Dis*. 2021; 17(12): 1956–1976, doi: [10.1016/j.soard.2021.08.024](https://doi.org/10.1016/j.soard.2021.08.024), indexed in Pubmed: [34629296](https://pubmed.ncbi.nlm.nih.gov/34629296/).
14. Taylor BA, Ng J, Stone A, et al. Effects of statin therapy on weight loss and diabetes in bariatric patients. *Surg Obes Relat Dis*. 2017; 13(4): 674–680, doi: [10.1016/j.soard.2016.11.018](https://doi.org/10.1016/j.soard.2016.11.018), indexed in Pubmed: [28159561](https://pubmed.ncbi.nlm.nih.gov/28159561/).