ORIGINAL CONTRIBUTIONS





Prevalence of and Risk Factors Associated with Alcohol Overconsumption at 2 Years After Bariatric Surgery

Lara Siikaluoma¹ · Erik Stenberg² · Mustafa Raoof²

Received: 6 December 2021 / Revised: 13 April 2022 / Accepted: 13 April 2022 / Published online: 25 April 2022 © The Author(s) 2022

Abstract

Introduction Alcohol overconsumption remains one of the adverse effects associated with bariatric surgery. Many previous studies have used subjective methods to evaluate the prevalence of alcohol overconsumption. In 2018, Örebro University Hospital started to use phosphatidylethanol 16:0/18:1 (PEth) as a screening tool pre- and postbariatric surgery. Research exploring alcohol use after bariatric surgery assessed with PEth is scarce.

Aim The aim of this study is to evaluate the prevalence of alcohol overconsumption in bariatric surgery patients measured 2 years postoperatively with PEth and to identify possible risk factors associated with alcohol overconsumption.

Methods This was a register-based retrospective, observational cohort study with PEth results collected from medical records at Örebro University Hospital. Patients who underwent bariatric surgery between January 2016 and June 2019 and who were registered in the Scandinavian Obesity Surgery Registry (SOReg) were included.

Results PEth results from 410 bariatric surgery patients were identified. PEth values significantly increased from baseline to the postoperative follow-up (from 3.0% before surgery to 8.3% at the 2-year follow-up). In a univariate logistic regression analysis, the associated risk factors were found to be male sex (odds ratio, OR 2.14), older age (OR 1.06), and hypertension (OR 3.32).

Conclusion The prevalence of alcohol overconsumption measured with PEth 2 years after bariatric surgery was 8.3% and was associated with male sex, older age, and hypertension. More studies are needed to validate the results of this study because it is not known whether PEth values are affected by bariatric surgery.

Keywords Bariatric surgery · Weight loss surgery · Alcohol overconsumption · Alcohol use disorder · Phosphatidylethanol

Key Points

Alcohol overconsumption at 2 years after sleeve gastrectomy or Roux-en-Y gastric bypass was 8.3%.

Men, patients of older age, and those with hypertension at baseline were at higher risk for alcohol overconsumption after surgery.

More studies are needed to evaluate how PEth values are affected by gastric surgery.

Mustafa Raoof mustafa.raoof@regionorebrolan.se

> Lara Siikaluoma larasiikaluoma@hotmail.com

Erik Stenberg erik.stenberg@regionorebrolan.se

¹ School of Medical Sciences, Örebro University, Örebro, Sweden

² Department Surgery, Faculty of Medicine and Health, Örebro University, Örebro, Sweden

Introduction

Significant changes in alcohol metabolism have been observed after Roux-en-Y gastric bypass (RYGB), such as a higher peak concentration of alcohol, faster absorption, and a longer time to become sober [1, 2]. Alterations in alcohol metabolism may result in an increased risk for alcohol use disorders (AUDs) after surgery, a risk that persists over time [3, 4]. However, it has been suggested that AUD after bariatric surgery may not represent new-onset AUD but rather a relapse after full sustained remission around the time of bariatric surgery [5]. Alternatively, there are theories suggesting that people with obesity have changed one addiction for another; in this case, it is alcohol to food, and after bariatric surgery, they change back from food to alcohol [5-7]. Phosphatidylethanol (PEth), an abnormal phospholipid, can be used as an alcohol marker to detect and monitor alcohol abuse; moreover, it can be used to identify early harmful alcohol use with a theoretical diagnostic specificity of 100% because PEth is formed only when alcohol is present [8]. PEth can be detected for up to 30 days, but the results can be affected by how often alcohol is consumed and the blood alcohol concentration [9]. There are different homologs of PEth, but the most common homolog is 16:0/18:1, representing 46% of total PEths [10, 11]. In a prospective multicenter cohort study evaluating the Alcohol Use Disorders Identification Test (AUDIT), the prevalence of AUD was 7.3% 1 year after bariatric surgery, increasing in prevalence to 9.6% 2 years after surgery [12]. A meta-analysis and systematic review by Azam et al. found that there was no significant difference in the prevalence of AUD 1 and 2 years postbariatric surgery, but the prevalence increased significantly 3 years after the RYGB procedure [13]. All of these studies were based on self-reported alcohol use. Studies evaluating alcohol use with PEth are limited. In a prospective cohort study, Walther et al. used a reference group of healthy blood donors and two separate groups of patients undergoing RYGB to assess alcohol intake with PEth at preoperative baseline and at 1 and 2 years postoperatively. The study found that the PEth levels of the bariatric surgery groups were lower than those of the reference group at baseline and postoperatively; notably, in both bariatric surgery groups, the PEth levels increased significantly postoperatively compared to baseline but did not reach the PEth levels of the control group. However, this study included only PEth $\geq 0.05 \ \mu mol/L$ without classifying it further as $0.05-0.30 \ \mu mol/L \ or > 0.30 \ \mu mol/L;$ in addition, the prevalence of alcohol overconsumption and risk factors were not examined [14]. The aim of this study was to evaluate the prevalence of alcohol overconsumption at 1 and 2 years after bariatric surgery and to identify possible risk factors associated with alcohol overconsumption.

Material and Methods

Study Design

This was a retrospective observational cohort study based on prospectively collected data registered in the Scandinavian Obesity Surgery Registry (SOReg). The SOReg, which has been shown to have high data quality and is constantly validated, covers 98% of all bariatric procedures performed in Sweden since 2009 with records entered in the database preoperatively, at surgery and at outpatient visits at 6 weeks and 1, 2, and 5 years after surgery [15]. Data from the SOReg includes information on patient demographics, postoperative complications, associated medical problems, and weight measurements. Associated medical problems were specified as specific conditions (diabetes, depression, dyslipidemia, hypertension, or sleep apnea) treated with pharmacological treatment or nocturnal continuous positive airway pressure in the case of sleep apnea. Previous AUD was defined as a previous diagnosis from a specialized unit or in the case of previous use of medication for alcohol disorder (Anatomical Therapeutic Chemical code: N07BB). PEth results were collected retrospectively from the electronic patient records at Örebro University Hospital and registered within the SOReg database. All available baseline and postoperative PEth results were included.

Study Population

All patients who were eligible for and underwent sleeve gastrectomy (SG) or RYGB at Örebro University Hospital and a County Hospital between January 2016 and June 2019 with a registration in the SOReg were included in this study. To be eligible for bariatric surgery after fulfilling the BMI requirements (BMI \geq 35 kg/m² with or without associated medical problems), patients need to be free from drug or alcohol abuse for at least 2 years before surgery if they have any history of abuse [16]. Bariatric surgery was performed by the same team; the standard RYGB procedure followed the H. Lönroth method [17, 18], and the SG procedure followed the New York summit consensus [19].

PEth Measurements

PEth was introduced as a routine measurement in this patient cohort in 2018. The results were registered within the clinical registry. Alcohol overconsumption was defined as PEth > 0.30 μ mol/L; moderate alcohol consumption, PEth 0.05–0.30 μ mol/L; and low or no alcohol consumption, PEth < 0.05 μ mol/L, as suggested by Equalis, an expert group for pharmaceuticals and toxicology, to standardize the PEth results in Sweden [20]. PEth results were obtained from blood samples analyzed in accordance with regional laboratory guidelines.

Statistical Methods

The prevalence of alcohol consumption and patients' baseline demographics were analyzed using descriptive statistical methods. Differences between the prevalence of alcohol overconsumption (PEth > 0.30 μ mol/L) and at least moderate consumption (PEth > 0.05 μ mol/L) from baseline to 1 and 2 years were assessed for statistical significance with the χ 2 test. Associated baseline factors for alcohol overconsumption were evaluated using an unadjusted logistic regression analysis, with odds ratios (ORs) and 95% confidence intervals (CIs) as measures of association. *P* values < 0.05 were considered indicative of statistical significance. All analyses were carried out using IBM SPSS Statistics MAC version 26.0 (Armonk, NY, USA).

Results

Collected Data

During the study period, 621 patients were included in the SOReg and followed for 2 years after surgery. Since PEth was introduced as a routine measurement in this patient cohort in 2018, samples were not available for all patients at all points in time. Of the 621 patients included, 236 (84.9%; of those with potential registration) had a registered PEth result at baseline, 381 (61.4%; of those with potential registration) had a result at 1 year, and 410 (66.0%; of those with potential registration) had a result at 2 years. In total, 114 patients (18.4%) had PEth results registered at all three time points (baseline and 1 and 2 years, postoperatively).

Preoperative and Postoperative Characteristics of the Study Cohort

Table 1 shows the preoperative characteristics of the study cohort at baseline. The mean age was 40 years, 73% were females, and the mean preoperative BMI was 42. At 2 years after surgery, remission of several metabolic associated *medical problems* was seen, but more patients were current smokers; in addition, a higher prevalence of clinical depression was noted (Table 2).

| Table 1 | Preoperative | (baseline) | demographics | and | characteristics | of |
|----------|--------------|------------|--------------|-----|-----------------|----|
| the stud | y cohort | | | | | |

| Variable | Preopera- tive cohort, n = 621 |
|--------------------------------------|--------------------------------------|
| Age (y), mean \pm SD | 40.2 ± 12.4 |
| Preoperative BMI, mean \pm SD | 42.0 ± 5.5 |
| Sex, <i>n</i> (%) | |
| Female | 455 (73.3) |
| Male | 166 (26.7) |
| Associated metabolic problems, n (%) | |
| Diabetes | 80 (12.9) |
| Depression | 52 (8.4) |
| Dyslipidemia | 42 (6.8) |
| Hypertension | 153 (24.6) |
| Sleep apnea | 83 (13.4) |
| Previous AUD ¹ , n (%) | 29 (4.7%) |
| Current smoker, n (%) | 85 (14.0%) |

BMI body mass index. SD standard deviation. AUD alcohol use disorder

¹Missing data for 1 patient (0.2%)

Characteristics of the Study Cohort at the 2-Year Follow-Up

At baseline before bariatric surgery, the prevalence of alcohol overconsumption (PEth > $0.30 \ \mu$ mol) was 3.0%. The prevalence of alcohol overconsumption at the 1-year followup was 6.8% and at 2 years 8.3%. The number of patients with low or no alcohol consumption declined with time after surgery (Table 3).

Patients with high alcohol overconsumption at 2 years were more likely to be older and male. They were more likely to require pharmacological treatment for diabetes and hypertension 2 years after surgery (Table 2). RYGB was the predominant surgical method in all groups, with no difference in alcohol overconsumption between the surgical methods. Increased age, male sex, and hypertension at baseline were all associated with a higher risk for AUD at 2 years after surgery (Table 4).

Discussion

Alcohol overconsumption increased over time, with a prevalence of alcohol overconsumption of 8.3% at 2 years postoperatively. Male sex, older age, hypertension, and previous AUD were associated with alcohol overconsumption 2 years after bariatric surgery.

Walter et al. reported a prevalence of alcohol consumption (PEth > 0.05 μ mol) of 26% 2 years after surgery among RYGB patients, which is in agreement with the alcohol consumption in our study (26.1%). In their study, a group of healthy blood donors was used as the reference, with PEth results > 0.05 μ mol found in 44% of the controls, indicating that there could be higher alcohol consumption in the general population than among bariatric surgery patients [14].

Many previous studies have used the Alcohol Use Disorders Identification Test (AUDIT/AUDIT-C) questionnaires [3, 4, 12, 21], reporting an increased prevalence of AUD from the first to second postoperative year, reaching 9.6% at 2 years after surgery [12]. While comparisons between studies using self-estimates and PEth values may not be directly comparable, PEth results have been shown to overlap with AUDIT-C results [22–24]. The rate of AUD in our study reflects those reported in previous studies that have defined alcohol overconsumption with more subjective methods.

In our study, the risk of AUD increased within the first postoperative year and increased further over time. Similar findings have been reported previously in studies using AUDIT-based definitions as well [4, 12]. Interestingly, one study showed that the prevalence of new-onset alcohol over-consumption was 7% 1 year after bariatric surgery and 6% after 2 years, whereas more than 50% of baseline high-risk drinking resolved after surgery [21]. In the context of our

Table 2Demographicsand characteristics of thestudy cohort at 2 yearspostoperatively, dividedinto groups according toalcohol overconsumption(PEth > 0.30 μ mol/L) or not

Table 3Prevalence of alcoholconsumption levels among the621 surgical patients who hadPEth measurements taken atbaseline, 1 year and/or 2 years

| Variable | Entire cohort | Alcohol overcon- sumption | No alcohol over- consumption | P value |
|----------------------------------|-----------------|------------------------------|---------------------------------|-----------|
| Age, mean \pm SD | 43.0 ± 12.4 | 50.8 ± 9.9 | 42.3 ± 12.3 | < 0.001 |
| Preoperative BMI, mean \pm SD | 41.7 ± 5.2 | 41.2 ± 5.3 | 41.8 ± 5.2 | 0.541 |
| Change in BMI, mean \pm SD | -13.6 ± 4.4 | -12.9 ± 4.3 | -13.6 ± 4.4 | 0.336 |
| %EBMIL, mean \pm SD | 84.1 ± 24.1 | 84.2 ± 28.6 | 84.0 ± 23.6 | 0.981 |
| Sex, <i>n</i> (%) | | | | |
| Female | 307 (74.9%) | 20 (58.8%) | 287 (76.3%) | Reference |
| Male | 103 (25.1%) | 14 (41.2%) | 89 (23.7%) | 0.027 |
| Surgical method, n (%) | | | | |
| Gastric bypass | 289 (70.5%) | 26 (76.5%) | 263 (70.0%) | Reference |
| Sleeve gastrectomy | 121 (29.5%) | 8 (23.5%) | 113 (30.0%) | 0.426 |
| Associated medical problems, n (| %) | | | |
| Current smoker ² | 68 (20.2%) | 7 (23.3%) | 61 (19.9%) | 0.659 |
| Diabetes ³ | 20 (5.4%) | 5 (15.2%) | 15 (4.4%) | 0.015 |
| Depression ³ | 50 (13.4%) | 5 (15.2%) | 45 (13.3%) | 0.763 |
| Dyslipidemia ³ | 27 (7.3%) | 5 (15.2%) | 22 (6.5%) | 0.076 |
| Hypertension ³ | 67 (18.0%) | 14 (42.4%) | 53 (15.6%) | < 0.001 |
| Sleep apnea ³ | 13 (3.5%) | 2 (6.1%) | 11 (3.2%) | 0.408 |

BMI body mass index. SD standard deviation; EBMIL excess BMI loss

¹Alcohol overconsumption was defined as PEth > 0.30 µmol/L

²P values based on unadjusted logistic regression analysis

³Missing data from 38 patients. Depression defined as if the patient was receiving pharmacological treatment against depression

⁴Missing data from 74 patients

| PEth (µmol/L) ¹ | Baseline, $n = 236$ | 1 year, <i>n</i> =381 | P^2 1 year vs. baseline | 2 years, <i>n</i> =410 | P^2 2 years vs. base- line |
|----------------------------|---------------------|-----------------------|---------------------------|------------------------|------------------------------------|
| <0.05, <i>n</i> (%) | 193 (81.8) | 285 (74.8) | Reference | 303 (73.9) | Reference |
| 0.05–0.30, <i>n</i> (%) | 36 (15.3) | 70 (18.4) | | 73 (17.8) | |
| >0.30, n (%) | $7(3.0)^3$ | 26 (6.8) | 0.038 | 34 (8.3) | 0.007 |

PEth phosphatidylethanol

¹PEth < 0.05 μ mol/L was defined as low or no alcohol consumption; PEth 0.05–0.30 μ mol/L, as moderate alcohol consumption; and PEth > 0.30 μ mol/L, as alcohol overconsumption

 ^{2}P values based on the χ^{2} test

³At that time period, we had not established reference values for the use of PEth values in the evaluation of this as a contraindication for bariatric surgery. Thus, 7 patients whom we would now consider to have contraindications for surgery were included (based on best available information at that time)

study, this could indicate that many of the patients with AUD at follow-up had developed new-onset alcohol overconsumption after surgery.

Another possible explanation for the increase in alcohol overconsumption after bariatric surgery could be a relapse in alcohol use rather than the development of a new-onset alcohol problem. Patients with a lifetime history of AUD who undergo bariatric surgery are more likely to develop an AUD postoperatively, although patients who underwent bariatric surgery in the USA have been reported to have a lifetime prevalence of AUD comparable to that of the general population [5]. In line with our findings, male sex has been documented as a risk factor for postsurgery AUD [3, 12]. Moreover, previous studies have reported that smoking and younger age are associated with an increased risk for AUD after bariatric surgery [3, 12]. In contrast, in our study, smoking did not influence the risk of alcohol overconsumption 2 years postbariatric surgery. Furthermore, older age was associated with a higher risk for overconsumption.

| Variable | OR (95% CI) | P value ¹ | | |
|---------------------------------|-------------------|----------------------|--|--|
| Age, mean ± SD | 1.06 (1.03–1.09) | < 0.001 | | |
| Preoperative BMI, mean \pm SD | 0.98 (0.92-1.06) | 0.644 | | |
| Sex, $n (\%)^1$ | | | | |
| Female | Reference | Reference | | |
| Male | 2.14 (1.05-4.39) | 0.037 | | |
| Surgical method, $n (\%)^1$ | | | | |
| Gastric bypass | Reference | Reference | | |
| Sleeve gastrectomy | 0.69 (0.30-1.56) | 0.369 | | |
| Current smoker ² | 1.20 (0.44–3.24) | 0.726 | | |
| Associated medical problems | | | | |
| Diabetes | 2.07 (0.89-4.82) | 0.092 | | |
| Depression | NA | 1.000 | | |
| Dyslipidemia | 2.15 (0.77-5.98) | 0.144 | | |
| Hypertension | 3.32 (1.64-6.73) | 0.001 | | |
| Sleep apnea | 1.74 (0.72–4.22) | 0.408 | | |
| Previous AUD | 3.57 (1.10–11.54) | 0.034 | | |

Table 4 Baseline factors associated with an increased risk for alcohol overconsumption (PEth > 0.30 μ mol/L)

BMI body mass index. *SD* standard deviation; *EBMIL* excess BMI loss. *AUD* alcohol use disorder

¹Based on unadjusted logistic regression analysis

²Missing data from 16 patients

An unexpected finding in our study was the lack of any significant association between depression and postoperative alcohol overconsumption. In fact, none of the 56 patients who were treated for clinical depression at baseline developed AUD. This finding is supported by King et al., who also investigated preoperative mental health, binge eating, and psychiatric treatment [12].

Last, we did not see any difference in alcohol overconsumption between the RYGB and SG groups, which is also in agreement with another study and in accordance with the fact that SG also significantly alters alcohol metabolism [25, 26].

Patients who developed AUD more often had metabolic *associated medical problems* (in particular in the form of hypertension) at baseline. While it remains unlikely that hypertension per se should be associated with the development of AUD, patients with hypertension tend to be older and may also share additional factors associated with both hypertension and AUD. In addition, patients who did develop AUD had higher rates of metabolic associated *medical problems* (including hypertension) at 2 years after surgery. This finding is consistent with previous reports of a linear increase in the risk of hypertension when alcohol consumption increases [27]. While bariatric surgery improves hypertension, diabetes, and other metabolic *associated medical problems*, alcohol overconsumption has been reported to increase the risk of hypertension

threefold [28], which is in line with the findings of the present study.

Despite the strengths of a well-defined cohort, the inclusion of RYGB and SG, and the administration of standardized counseling, follow-up, and supplementation, there are potential limitations that may affect the interpretation of the results in this study. First, because of the retrospective nature of this study, there were missing data. Since screening with PEth started in 2018 and our study cohort included patients who underwent surgery before that, there were missing data, especially at baseline and 2 years after surgery, affecting the reliability of the results. One could expect that there was no selection bias because patients with the same eligibility criteria underwent surgery before and after the beginning of PEth screening. During the study period, the evaluation of contraindications for bariatric surgery relied on self-reported alcohol use and review of hospital charts which explains why seven patients with a PEth value ≥ 0.30 still underwent surgery.

In addition, this is a small patient sample, which affects the generalizability of the results. Finally, to our knowledge, whether the production of PEth in response to alcohol intake differs when gastrointestinal anatomy is surgically altered remains unknown.

Conclusion

The prevalence of alcohol overconsumption measured with PEth 2 years after bariatric surgery was 8.3% and was associated with male sex, older age, AUD, and hypertension.

Funding Open access funding provided by Örebro University.

Declarations

Ethics Approval The study was approved by the Swedish Ethical Review Authority on July 28th, 2020 (registration number 2021–02896) and follows the ethical standards of the 1964 Helsinki Declaration and its later amendments.

Consent to Participate Informed consent was obtained from all individual participants when entering the Scandinavian Obesity Surgery Registry.

Conflict of Interest The authors declare no competing interests.

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