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# Incidence of ocular pathology following bariatric surgery for with morbid obesity across a large United States National Database

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**BACKGROUND/OBJECTIVES:** Bariatric surgery, as indicated for treatment of morbid obesity, has been studied in association with short term effects on ocular pathology. However, effects of surgery on postoperative disease incidence is largely unknown. **SUBJECTS/METHODS:** In this retrospective cohort study, the TriNetX United States Collaborative Network national database, was queried for patients with an ICD-10 code for morbid obesity and a procedural code for bariatric surgery. Patients were propensity score matched across baseline demographics at the time of surgery and compared to those presenting with an ICD10 code for morbid obesity surgery, identifying 42,408 patients per cohort. New diagnoses or procedural codes found after the surgical index date for diabetic retinopathy, age-related macular degeneration, glaucoma, low vision, and blindness along with pertinent treatment metrics were monitored.

**RESULTS:** Bariatric surgery was found to be associated with reduced future risk of diabetic retinopathy (RR: 0.283; 95% CI: 0.252–0.319), macular edema (RR: 0.224; 95% CI: 0.170–0.297), vitreous hemorrhage (RR: 0.459; 95% CI: 0.323–0.653), ocular hypertension (RR: 0.387; 95% CI: 0.387–0.487), glaucoma (RR: 0.360; 95% CI: 0.326–0.399), use of ocular pressure lowering medications (RR: 0.565; 95% CI: 0.496–0.644), age-related macular degeneration (RR: 0.628; 95% CI: 0.447–0.882), cataract surgery (RR: 0.524; 95% CI: 0.448–0.612), and low vision and blindness (RR: 0.328; 95% CI: 0.294–0.365) compared to patients not surgically managed. **CONCLUSIONS:** The present analysis comprising a large US cohort of patients suggests that bariatric surgery is associated with a decreased risk of future ocular morbidity and mortality.

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## INTRODUCTION

As obesity co-exists with and exacerbates systemic health conditions, obesity has been associated with increased risk of all-cause mortality [1]. While diet and weight loss efforts are often first line in mitigation of obesity and restoration of normal weight, intensive surgical interventions such as bariatric surgery is an increasingly utilized option for patients who present with a body mass index (BMI) above 40 and are refractive to conservative medical management [2]. Since the advent of bariatric surgery, the efficacy of this procedure on weight loss and systemic health has been profound. Bariatric surgery has been shown in large clinical trials to reduce mean BMI, along with risk factors for cardiovascular disease and metabolic syndrome [3, 4].

Obesity has been implicated as a driver of various ocular disease states, increasing risk for diabetic retinopathy, glaucoma, and age-related macular degeneration [5]. Due to these associations, increasing attention has been focused on effects of bariatric surgery and mitigation of ocular pathology. A two-year analysis of the STAMPEDE trial, a prospective randomized trial examining bariatric surgery compared to intensive medical management for treatment of morbid obesity, showed patients treated with bariatric surgery to experience no worsening or benefit with respect to diabetic retinopathy (DR) outcomes [6]. An subsequent 5-year analysis further found no differences in DR worsening or improving in patients undergoing bariatric surgery compared to medically managed patients [7]. However, a recent systematic review pooling 876 patients undergoing bariatric surgery showed patients who underwent surgery were at decreased odds of DR progression than those who were obese and did not undergo surgery [8].

With respect to other ocular pathologies, small cohorts of patients undergoing bariatric surgery have also shown decreases in intraocular pressure (IOP) short term post-surgery [9, 10]. However long-term effects examining development of glaucoma have yet to be examined. Zhang et al. explored a pooled analysis of studies examining age related macular degeneration (AMD) and body mass index BMI, finding every 1 kg/m<sup>2</sup> increase in BMI incurred a 2% increased risk of carrying a diagnosis of AMD [11]. While these results seem promising, no studies to date have examined odds of developing AMD after intervention with bariatric surgery. Obesity has been found to incur increased risk of cataract formation [12, 13]. Recently, Burkard et al. examined patients undergoing

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bariatric surgery, finding them to be at significantly decreased risk of cataract formation [14]. However, this design did not stratify by types of cataract, and need for future cataract surgery, leaving areas for continued investigation.

Although the associations between bariatric surgery and development of ocular disease are mixed, studies to date largely reveal protective trends. Furthermore, these associations have yet to be explored in the context of large U.S. cohort. The advent of TriNetx, a large U.S. collaborative national database, allows such studies to be performed. Herein, this study aims to characterize the effect of bariatric surgery on post-operative prevalence of diabetic retinopathy, glaucoma, age-related macular degeneration, blindness, and associated disease specific worsening events using the TriNetX U.S. Collaborative network.

## MATERIALS/SUBJECTS AND METHODS

This study was performed utilizing the TriNetX U.S. Collaborative Network: a large database containing aggregated, standardized, and de-identified medical records data that utilizes International Classification of Diseases (ICD) codes. The TriNetX U.S. Collaborative Network provides access to electronic medical records information on approximately 86 million patients from 54 contributing healthcare organizations. TriNetX, LLC is a Health Insurance Portability and Accountability Act (HIPAA) compliant, certified to the ISO 27001:2013 standard, and maintains an Information and Security Management System to ensure protection of data.

Using the TriNetX database, patients were queried for presence of a procedural code for bariatric surgery (43,774) and international classification of disease-10 (ICD-10) code for morbid obesity (E66.01), identifying 42,408 patients. A control cohort of patients with a diagnosis of morbid obesity (E66.01) and no procedural code for bariatric surgery (43774) was also queried, identifying 972,234 patients. To achieve balance amongst cohorts, propensity score matching was performed using the TriNetX analytic feature (greedy matching, calliper of 0.25 SD). The following criteria were matched upon: age, gender, race, ethnicity, BMI, hypertension (HTN), and hemoglobin A1c.

Once cohorts were obtained, development of pertinent ICD-10 codes were queried for after the date of bariatric surgery for surgical patients or after a diagnosis of morbid obesity in control cohorts as seen in Supplementary Table 1. For surgical patients, the analysis window was any date after bariatric surgery. For non-surgical patients, the analysis window was any time after the diagnosis of morbid obesity. To ensure only incident cases were included in the analysis, patients with an ICD pathology, procedure, or medication code of interest before the surgical or matching date were excluded from the analysis. Risk ratios were calculated to determine the impact of a bariatric surgery on the risk of new ocular diagnoses. Risk ratios were reported, along with confidence intervals.

# RESULTS

# Demographics

In the TriNetX national database, a total of 42,408 patients were identified to carry a diagnosis of morbid obesity and underwent bariatric surgery. A total of 831,791 patients were identified to carry a diagnosis of morbid obesity lacking a procedural code for bariatric surgery. Propensity score matching was used as stated above. Before matching, patients in the bariatric cohort were significantly younger, included a greater percentage of female patients, and were more obese than those who were not treated surgically. After matching, each cohort contained 42,408 patients and patient cohorts were more similar with respect to age, gender, race, BMI, HbA1c, and HTN status although differences were still present (Table 1).

## Ocular outcomes

Of the 42,408 patients who underwent bariatric surgery, 347 of them were given a new diagnosis of any type of diabetic retinopathy after their surgical date. Of the patients who did not undergo surgery, 1232 of them experienced a new diagnosis of diabetic retinopathy in the future as seen in Table 2. Patients who underwent surgery had significantly reduced risk of developing new diabetic retinopathy compared to those who were not (RR: 0.283; 95% CI: 0.252-0.319). When stratified by category of DR, patients who were surgically managed were significantly less likely to experience nonprolifereative DR (RR: 0.324; 95% CI: 0.274–0.382) or proliferative DR (RR: 0.453; 95% CI: 0.339-0.604) compared to those who did not undergo surgery (Table 2). Patients who underwent bariatric surgery were also significantly less likely to experience complications of diabetic retinopathy worsening such as vitreous hemorrhage, intravitreal injection, pars plana vitrectomy, pan retinal photocoagulation, and tractional retinal detachment (p < 0.05 for all, Table 2).

Patients who underwent surgery were significantly less likely to be newly diagnosed with all types of AMD (RR: 0.628; 95% CI: 0.447–0.882) than those who did not have bariatric surgery. When stratified by type of AMD, patients in the bariatric cohort were still significantly less likely to experience nonexudative AMD (RR: 0.671; 95% CI: 0.464–0.971) and exudative AMD (RR: 0.500; 95% CI: 0.269–0.929) (Table 2).

While less cases of retinal vascular occlusion were observed in the surgically treated cohort, these differences were not found to be significant (RR: 0.746; 95% CI: 0.523–1.064). When stratifying by retinal vein or artery occlusion, no associations were uncovered (p > 0.05, Table 2).

Patients who underwent surgery were significantly less likely to present with new onset ocular hypertension (RR: 0.387; 95% CI: 0.308–0.487) or be diagnosed as a glaucoma suspect (RR: 0.406; 95% CI: 0.358–0.461) than those did not undergo surgery. Regarding all glaucoma associated diagnoses, patients undergoing surgery were significantly less likely to develop associated disease states (RR: 0.365; 95% CI: 0.330–0.403) as seen in Table 3. Concerning clinical management, patients undergoing surgery were significantly less likely to require pressure lowering medications (RR: 0.565; 95% CI: 0.496–0.644), or undergo future glaucomatous procedures or surgeries (RR: 0.461; 95% CI: 0.340–0.626). Finally, surgical patients were less likely to be diagnosed with new OSA (RR: 0.267; 95% CI: 0.250–0.285).

Patients treated surgically significantly less likely to receive new diagnoses of cataract formation including age-related cataract, nuclear sclerotic cataract, cortical cataract, posterior subcapsular cataract, and diabetic cataract (p < 0.001 for all, Table 4). Surgically managed patients were also at significantly decreased risk of future cataract surgery than those not managed with surgery (RR: 0.524; 95% CI: 0.448–0.612).

Concerning low vision and blindness, patients undergoing bariatric surgery were at significantly reduced likelihood of development compared to those who did not receive surgery (RR: 0.328; 95% CI: 0.294–0.365). When stratified by patients with only low vision and only blindness, those who underwent bariatric surgery were still significantly less likely to experience low vision (RR: 0.337; 95% CI: 0.302–0.377) and blindness (RR: 0.355; 95% CI: 0.274–0.460) (Table 5).

# DISCUSSION

The present study examining associations between bariatric surgery and common ocular pathology across a large national database, suggests bariatric surgery and associated systemic improvements may serve a protective role in development of ocular morbidity and mortality. Patients whose morbid obesity was managed surgery were significantly less likely to experience diabetic retinopathy, vitreous hemorrhage, intravitreal injection,

Variable	Bariatric surgery [SD] or (%) (N = 42,408)	Obese [95% SD] or (%) (N = 42,408)	p value
Age (years)	43.2 +/- 12.4	43.6 +/- 13.9	<0.001
Gender			
Female	34,273 (82.3%)	34,323 (80.9%)	0.662
Male	8131 (19.2%)	8073 (19.0%)	0.612
Race			
White	27,307 (64.4%)	27,961 (65.9%)	<0.001
Black	11,237 (26.5%)	10,639 (25.1%)	<0.001
Ethnicity			
Hispanic	3772 (8.9%)	3300 (7.8%)	<0.001
BMI			
BMI	42.9 +/- 5.4	40.8 +/- 5.5	<0.001
Labs			
HbA1c	$6.03 \pm 1.24$	6.2 ± 1.6	<0.001
LDL cholesterol	106 ± 32.8	106 ± 33.1	0.076
Systemic diagnoses			
Essential HTN	23,378 (55.1%)	23,542 (55.5%)	0.257
Type 1 DM	1230 (2.9%)	1126 (2.7%)	0.029
Type II DM	12,055 (28.4%)	11,674 (27.5%)	0.003
Long term use of insulin	2176 (5.1%)	2074 (4.8%)	0.108
Hypercholesterolemia	5670 (13.4%)	5504 (12.9%)	0.092
CAD	2623 (6.2%)	2403 (5.1%)	0.005
OSA	21,005 (49.5%)	20,778 (48.9%)	0.119
Ocular diagnoses			
Type I DM with DR	90 (0.2%)	69 (0.2%)	0.095
Type II DM with DR	644 (1.5%)	450 (1.1%)	<0.001
non exudative AMD	30 (0.1%)	30 (0.1%)	1
exudative AMD	10 (0.1%)	15 (0.1%)	0.317
Glaucoma suspect	661 (1.4%)	467 (0.9%)	<0.001
OHTN	195 (0.5%)	153 (0.4%)	0.024
Glaucoma	926 (2.2%)	738 (1.7%)	<0.001
Blindness and low vision	651 (1.5%)	537 (1.3%)	<0.001

Table 1. Demographic information of Bariatric surgery and Obese cohorts.

Statistically significant values are in bold.

SD standard deviation, BMI body mass index, HTN Hypertension, HbA1c hemoglobin A1c, LDL low density lipoprotein, DM diabetes mellitus, CAD coronary artery disease, OSA obstructive sleep apnea, DR diabetic retinopathy, AMD age related macular degeneration, OHTN ocular hypertension.

vitrectomy, ocular hypertension, glaucoma, glaucomatous surgeries, use of ocular pressure lowering medications, age-related macular degeneration, cataract formation, cataract surgery, low vision, and blindness compared to patients who were not treated surgically.

This study largely supports and adds clarity to what has been reported in the literature regarding patients undergoing bariatric surgery and ocular pathology. With respect to diabetic retinopathy, this study supports findings of Akerblom et al. who found bariatric surgery to reduce future development of diabetic retinopathy in 5321 Swedish patients with diabetes. However, Akerblom et al also reported no difference in diabetic retinopathy complications such as development of proliferative DR, need for PRP, or IVI, which the present study finds to be reduced, possibly due to larger sample sizes leading to an appropriately powered analysis [15]. The present study also examines additional complications of DR not examined by Akerblom et al. such as development VH, finding reduced frequency in those whose obesity was managed surgically. The present data runs contrary to analyses of the STAMPEDE trial, which found no difference in DR incidence comparing patients who underwent bariatric surgery to intensive medical management. While these differences may be due to larger sample size in the present study, it is also plausible these differences are due to variations in the comparator group. In the STAMPEDE trial, patients undergoing surgery were compared to those undergoing intensive medical management of systemic disease while the present study compared outcomes to real world morbidly obese patients who were not managed surgically. Therefore, it is plausible the control cohort in the present study may have reduced access to care, decreased long term follow up, were poor candidates for surgery, or other differences unable to be controlled for in the study design. However, if this were the driving factor, it would be expected for this population to have reduced ophthalmic visits and diagnoses relative to the surgical cohort, potentially diminishing true effect of the surgical intervention. The diabetic cohorts as a whole suggest that treatment with bariatric surgery may be associated with reduced future development of diabetic retinopathy in morbidly obese patients along with sight-threatening complications of DR such as development of VH, TRD, DME, and need for treatment with IVI, and PRP.

Table 2. Risk of post operative retinal pathology and complications after surgery.

Cohorts	Patients in cohort	Patients with outcome	Risk ratio (CI)	P value
All forms of di	abetic retinopat	hy		
Bariatric surgery	41,661	347	0.283 (0.252,	<0.001
Obese	41,880	1232	0.319)	
Nonproliferativ	ve diabetic retine	opathy		
Bariatric surgery	42,156	183	0.324 (0.274,	<0.001
Obese	42,221	566	0.382)	
Mild nonprolif	erative diabetic	retinopathy		
Bariatric surgery	42,197	160	0.334 (0.279,	<0.001
Obese	42,247	480	0.399)	
Moderate non	proliferative diab	petic retinopathy		
Bariatric surgery	42,341	54	0.342 (0.251,	<0.001
Obese	42,365	158	0.466)	
Severe nonpro	liferative diabeti	ic retinopathy		
Bariatric surgery	42,376	25	0.373 (0.236,	<0.001
Obese	42,383	67	0.591)	
Proliferative di	iabetic retinopat	hy		
Bariatric surgery	42,295	67	0.453 (0.339,	<0.001
Obese	42,316	148	0.604)	
Diabetic retine	pathy with mac	ular edema		
Bariatric surgery	42,046	60	0.224 (0.17,	<0.001
Obese	42,150	268	0.297)	
Vitreous Hemo	orrhage			
Bariatric surgery	42,351	45	0.459 (0.323,	<0.001
Obese	42,348	98	0.653)	
Intravitreal inje	ection			
Bariatric surgery	42,339	83	0.522 (0.4,	<0.001
Obese	42,334	159	0.68)	
Pars Plana Vitr	rectomy			
Bariatric surgery	42,381	23	0.46 (0.281,	0.001
Obese	42,394	50	0.754)	
Pan Retinal Ph	otocoagulation			
Bariatric surgery	42,367	21	0.467 (0.278,	0.003
Obese	42,378	45	0.783)	
Tractional Reti	nal Detachment			
Bariatric surgery	42,391	10	0.476 (0.224,	0.048
Obese	42,388	21	1.011)	
All forms of A	MD			
Bariatric surgery	42,370	54	0.628 (0.447,	0.006
Obese	42,366	86	0.882)	

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Table 2. conti	nued				
Cohorts	Patients in cohort	Patients with outcome	Risk ratio (Cl)	P value	
Non exudative	AMD				
Bariatric surgery	42,378	47	0.671 (0.464,	0.033	
Obese	42,378	70	0.971)		
Exudative AM	)				
Bariatric surgery	42,399	15	0.5 (0.269,	0.025	
Obese	42,393	30	0.929)		
Retinal vascula	r occlusion				
Bariatric surgery	42,362	53	0.746 (0.523,	0.105	
Obese	42,331	71	1.064)		
Retinal vein oo	clusion				
Bariatric surgery	42,386	32	1 (0.613, 1.632)	0.999	
Obese	42,376	32			
Retinal artery occlusion					
Bariatric surgery	42,396	16	0.762 (0.398,	0.411	
Obese	42,386	21	1.46)		
Statistically significant values are in hold					

Statistically significant values are in bold.

AMD Age related macular degeneration.

The present study is also one of the first to examine bariatric surgery and associations with development of glaucomatous disease. To date, a retrospective study of 22 bariatric surgery patients by Shimonov et al. showed that BMI reduction was associated with significant and continued decline in IOP beyond 1 year after surgery. Specifically it was found that average BMI decreased from 41.9  $\pm$  7.3 to 25.5  $\pm$  5.7 kg/m<sup>2</sup> at 1-year follow-up from bariatric surgery, which corresponded to a mean IOP decrease of 21% [16]. Additionally, small clinical cohorts examined by Burgansky-Eliash et al. have also shown bariatric surgery to reduce intraocular pressure [10]. Mechanistically, obesity associated orbital adipose tissue may elevate episcleral venous pressure, decreasing aqueous humor outflow resulting in IOP elevation [17]. Strong associations also exist between glaucoma and OSA, as carrying a diagnosis of OSA significantly increases the likelihood of being diagnosed with glaucoma [18]. These associations are thought to be a result of increased hypoxia in OSA, leading to optic nerve hypoxic damage and glaucomatous phenotype. Supporting these associations, patients who elected surgery were found to have lower incidences of being diagnosed with glaucomatous diseases such as ocular hypertension, glaucoma suspect, glaucoma, primary open angle glaucoma, and glaucomatous optic atrophy. Interestingly patients who underwent bariatric surgery were found to have significantly decreased incidence of OSA, in line with the above proposed mechanistic links between the two. In terms of clinical outcomes, surgically managed patients were less likely to use pressure lowering drops or undergo glaucoma surgeries in the future, highlighting relevant metrics for future analyses of clinical trials. The present study did not investigate secondary glaucoma mechanisms, such as neovascular glaucoma, which may also be affected by the many metabolic syndromes seen in morbidly obese patients.

While obesity has been identified as a risk factor for likelihood of being diagnosed with AMD, bariatric surgery has not been explored with respect to future development of disease. Although 
 Table 3. Risk of post-operative glaucomatous diseases and complications in cohorts.

Cohorts	Patiens in cohort	Patients with outcome	Risk ratio (CI)	P value
	Oc	ular hypertensic	on	
Bariatric surgery	42,211	101	0.387 (0.308,	<0.001
Obese	42,252	261	0.487)	
	G	laucoma suspec	t	
Bariatric surgery	41,733	335	0.406 (0.358,	<0.001
Obese	41,926	829	0.461)	
	Ocular hypert	ension + Glauco	oma suspects	
Bariatric surgery	41,733	335	0.406 (0.358,	<0.001
Obese	41,926	829	0.461)	
		Glaucoma		
Bariatric surgery	41,445	501	0.36 (0.326,	<0.001
Obese	41,626	1396	0.399)	
	Primary	Open Angle Gla	aucoma	
Bariatric surgery	42,292	67	0.466 (0.349,	<0.001
Obese	42,328	144	0.622)	
	Glaucomatou	is procedures ar	nd surgeries	
Bariatric surgery	42,300	60	0.461 (0.34,	<0.001
Obese	42,292	130	0.626)	
	Use of pre	ssure lowering e	eye drops	
Bariatric surgery	41,848	351	0.565 (0.496,	<0.001
Obese	41,918	622	0.644)	
	Obst	ructive sleep ap	nea	
Bariatric surgery	19,184	991	0.267 (0.25,	<0.001
Obese	20,703	4008	0.285)	
	Glauco	matous optic at	rophy	
Bariatric surgery	42,363	33	0.611 (0.396,	0.024
Obese	42,371	54	0.942)	
Glauc	oma Diagnose Glaucoma + G	s (Glaucoma + P ilaucomatous O	Primary Open Ang ptic Atrophy)	gle
Bariatric surgery	41,429	513	0.365 (0.33,	<0.001
Obese	41,605	1413	0.403)	
		Mild glaucoma		
Bariatric surgery	42,355	65	0.747 (0.542,	0.075
Obese	42,373	87	1.03)	
	Mo	oderate glaucom	าล	
Bariatric surgery	42,368	31	0.62 (0.396,	0.035
Obese	42,376	50	0.97)	
	S	evere glaucoma	I	
Bariatric surgery	42,381	19	0.704 (0.391,	0.238
Obese	42,380	27	1.265)	

Eye

 Table 4.
 Risk of post-operative cataract formation and cataract surgery.

C	Cohorts	Patients in cohort	Patients with outcome	Risk ratio (CI)	P value
		Age r	elated cataract		
E	Bariatric surgery	41,378	928	0.479 (0.444,	<0.001
	Obese	41,124	1925	0.518)	
		Nuclear	sclerotic cataract		
E	Bariatric surgery	41,565	758	0.483 (0.444,	<0.001
	Obese	41,400	1562	0.527)	
		Cor	tical cataract		
E	Bariatric surgery	42,296	113	0.451 (0.362,	<0.001
	Obese	42,234	250	0.563)	
		Posterior s	ubcapsular catara	act	
E	Bariatric surgery	42,329	62	0.397 (0.296, 0.533)	<0.001
	Obese	42,311	156		
		Diak	petic cataract		
E	Bariatric surgery	41,999	173	0.326 (0.274,	<0.001
	Obese	42,129	533	0.386)	
		Cata	aract surgery		
E	Bariatric surgery	42,228	239	0.524 (0.448, 0.612)	<0.001
	Obese	42,196	456		

Statistically significant values are in bold.

 Table 5.
 Risk of post-operative low vision or blindness.

	Cohorts	Patients in cohort	Patients with outcome	Risk ratio (CI)	P value
		Blindnes	s and Low Visior	า	
	Bariatric surgery	41,714	435	0.328 (0.294,	<0.001
	Obese	41,772	1329	0.365)	
Low vision					
	Bariatric surgery	41,810	410	0.337 (0.302, 0.377)	<0.001
	Obese	41,827	1217		
			Blindness		
	Bariatric surgery	42,251	78	0.355 (0.274, 0.46)	<0.001
	Obese	42,314	220		

Statistically significant values are in bold.

literature that explores the effect of bariatric surgery on ocular anatomy exists, there is a paucity of literature that explores AMD and bariatric surgery specifically. A prospective controlled study by Ozcelik-Kose et al. showed that total choroidal area and the choroidal vascularity index (CVI) exhibited significant increases when comparing the values 6 months before surgery and 6 months after surgery [19]. Similarly, in a prospective study of 40 patients, ElShazly et al. found significant increases in macular thickness and macular vascular density of the deep capillary plexus when compared 3 months before and 3 months after bariatric surgery [20]. Taken together these early studies may suggest systemic effects of bariatric surgery may include improvement in retinal microvascular circulation. Supporting these early associations, patients who underwent bariatric surgery were significantly less likely to develop nonexudative and exudative AMD. To more thoroughly examine these associations and impact on clinical treatment course, larger cohorts of surgical patients with long term follow up would be necessary.

Our study supports findings of Burkard et al., showing bariatric surgery to be associated with decreased risk of cataract diagnoses. However, the present study further adds novelty with the utilization of a large cohort of US patients, while also examining type of cataract developed and future need for cataract surgery [14]. Obesity has been identified as a significant risk factor for cataract formation, thought to be mediated through increases in reactive oxygen species and high circulating leptin in these patients, clouding the lens over time [21]. Diabetes is also a significant risk factor for cataract development, especially in those under 65 years of age, thought to be a result of increased osmotic and oxidative stress [22]. Because our cohorts are similar with respect to age and BMI at baseline, it is reasonable to assume that weight loss and systemic effects from bariatric surgery such as improved glycaemic control may have reduced likelihood of cataract development in our cohorts [4]. Furthermore, our findings show a reduction in incidence of all types of cataract including nuclear sclerotic, cortical, posterior subcapsular, and diabetic which have all been associated with obesity or diabetes [5, 22].

Supporting reductions seen in various ocular pathologies, patients in the bariatric cohort were found to be at significantly decreased risk of experiencing low vision or blindness post operatively. This finding largely aligns with data reported in the present study, as patients with bariatric surgery were less likely to develop DR, glaucomatous pathology, and AMD, all of which in isolation may contribute to low vision or blindness. While low vision and blindness is multifactorial in etiology and not all causes are investigated in the present study design, this finding is imperative to highlight that bariatric surgery not only results in reduced disease burden, but also reduces in visual decline in the process.

The findings of the present study, that morbidly obese patients who undergo bariatric surgery experience decreases in ocular morbidity and mortality, may be largely explained by improvements in systemic health as a result of the surgical intervention. Bariatric surgery has been shown long term to reduce BMI, improve glycaemic control, and reduce cardiovascular disease risk factors. Since DR is a microvascular complication of poor systemic diabetic health, it is reasonable to assume that the reduction in DR incidence may be explained by overall glycaemic improvement in this cohort. Reductions in incidence of glaucoma may be attributed to decreases in IOP or OSA over time in hand resulting in less disease development. Although the mechanistic link between obesity and AMD remains less clear, it is plausible to assume reductions in obesity, a studied risk factor for patients with AMD, is contributing to decreases in disease incidence observed. Future long term high powered clinical studies examining bariatric surgery and development of ocular pathology may add credence to the associations presented.

Strengths of the present study lie in the utilization of a national database to investigate bariatric surgery and its effect on the development of DR, glaucoma, AMD, and low vision and blindness in the future. To date, this is the largest longitudinal cohort to date examining these associations, allowing for analysis of a variety of pertinent ocular outcomes with respect to the above disease states. However, the present study is not without its limitations. Length of follow up is unable to be ascertained from the present data set, making it difficult to associate decreases in disease state seen with set chronologic time points. Even after utilization of propensity score matching, small baseline

differences still existed in the two populations. However, these small differences may have been significant due to the large cohort sizes used, overpowering the measure of association. Because the data obtained through TriNetX is presented in aggregated form, it is difficult to assess patient-level data such as individual visual acuity, disease progression, and treatment course. While the present study reports univariate analysis as allowed by the TrinetX platform, subsequent analyses with incorporation of multivariate analysis would be of interest to further explore these associations. Additionally, it is also possible that patients receiving surgery may have been of higher socioeconomic status than those who did not pursue surgery. Unfortunately, this level of patient data was unable to be extracted as only ICD10 associated data was pulled. Finally, as TriNetX relies on ICD-10 and procedural codes to categorize variables, the conclusions drawn from such data is contingent on proper physician coding, leaving room for bias if disease states are coded inconsistently across health care organizations.

This study provides novel insight examining bariatric surgery and ocular pathology across a large national cohort in the US, uncovering a protective association between the surgery and ocular disease. The associations herein suggest a need for further investigation of bariatric surgery and obesity reduction on development of ocular morbidity and mortality in large long term clinical settings.

# SUMMARY

What was known before

- Bariatric surgery have been studied in association with ocular pathology, showing mixed but promising results.
- However, these studies do not often report postoperative disease development and do not investigate multiple disease pathologies.

What this study adds

 This study is the first to examine the effect of bariatric surgery on risk of future ocular disease development across a large population, showing evidence that bariatric surgery may reduce development of multiple ocular diseases.

#### DATA AVAILABILITY

The data that support the findings of this study are available from the authors but restrictions apply to the availability of these data, which were requisitioned from the TriNetX network, and so are not publicly available. Data are, however, available from the authors upon reasonable request.

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#### **AUTHOR CONTRIBUTIONS**

All authors meet the ICMJE criteria for authorship.

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