

Lifestyle Counseling for Medication Adherence in Glaucoma

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Background: Medication adherence in glaucoma patients remains sub-par despite proven benefits of regular administration. The objective was to analyze medication adherence before and after lifestyle counseling in patients with ocular hypertension (OHT) or primary open-angle glaucoma (POAG) (mild, moderate, severe).

Methods: Prospective cohort study from May to July 2018 at a single academic center. From 391 consecutive records, 247 were excluded based on exclusion criteria with 28 patients not meeting inclusion criteria resulting in the final sample of 116 patients (33 had OHT, 83 had POAG – 28 mild, 39 moderate, 16 severe). Scripted lifestyle counseling focusing on diet, exercise, vitamin intake, stress management, and medication adherence was administered by a team of trained medical students. Primary outcome measure was self-reported medication adherence, defined as not missing an eye drop administration in the past month. A 2–3 week follow-up with scripted telephone survey assessing medication adherence, diet, and exercise was collected.

Results: At baseline, in 116 patients, 59.5% were adherent to their medication with a breakdown of 42.4% OHT, 64.3% mild, 66.7% moderate, and 68.7% severe and an increasing trend in medication adherence was found across increasing disease severity ($p=0.055$). Of the 76 (65.5%) patients reached for follow-up, 17 (22.4%) became adherent following lifestyle counseling ($p=0.02$) increasing overall adherence to 78.9% from 62.5%.

Conclusion: In our study, comprehensive lifestyle counseling succeeded in increasing medication adherence in patients with OHT and POAG.

Keywords: glaucoma, lifestyle counseling, medication adherence

Introduction

Despite effective medical and surgical therapy, glaucoma is still a leading cause of irreversible blindness worldwide.¹ This trend is exacerbated by non-adherence to medication regimens requiring daily eye drop administration to lower intraocular pressure (IOP) and prevent disease progression. Research has shown that patients who administer 80% or more of their prescribed eye drops have an 86% reduction in their odds of having moderate or severe visual field defects.² A recent study by Newman-Casey and colleagues confirmed a dose-dependent relationship between medication non-adherence and visual field loss over a period of 8 years with patients who reported not missing doses having similar annual visual field loss as those with stable glaucoma.³

Despite the benefits of medication adherence, various studies have found that almost half of patients are non-adherent with their medications.^{4–7} Reported reasons include forgetfulness,⁸ cost, lack of knowledge surrounding the glaucoma disease

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process or purpose of medical therapy,^{9,10} and provider's lack of communication regarding proper medication administration and adherence during clinic visits.¹¹ The chronic and asymptomatic nature of vision loss coupled with preventive rather than curative properties of glaucoma medication make it difficult for patients to adhere to a medication regimen they cannot see the direct benefits of.

Most studies analyzing the effects of educational interventions on medication adherence have designed interventions focused on eye drop administration technique and patient provider communication by showing patients instructional videos for eye drop administration,¹² videotaping and transcribing clinic visits,^{13,14} and assigning trained personnel to counsel patients.¹⁵ Interventions built on patient education and motivational interviewing are hypothesized to have improved glaucoma medication adherence by increasing patient confidence in following medication regimens^{14,16–20}. Carpenter and colleagues found that provider education regarding medication adherence was associated with improved IOP at an eight-month follow-up.¹⁴ Despite the evidence that these interventions are successful in improving medication adherence, there remains a high proportion of medication non-adherence in glaucoma patients indicating that educating patients on proper administration and regimen may not be the only limiting factor.

Fewer studies have focused on or incorporated lifestyle counseling in glaucoma despite evidence of benefit in similar "silent" diseases, such as diabetes and hypertension. Interventions focused on long-term behavioral control, such as diet, exercise, and weight loss, have shown glucose and blood pressure levels under control for as long as two years.^{21–27} While there exist few epidemiological studies on lifestyle risk factors for POAG, some research has shown evidence that certain lifestyle activities such as exercise, diet, vitamin intake, and smoking affect glaucoma disease onset and progression.²⁸

Given these findings, our study aimed to assess the effects of a more holistic, lifestyle-centered counseling on medication adherence. We hypothesized that lifestyle-based sessions would improve adherence to medication regimens by motivating patients to overall live healthier lifestyles.

Methods

Study Design

In this prospective cohort study, we interviewed 391 consecutive patients and selected 116 eligible patients at the

University of Texas Southwestern (UTSW) Medical Center Eye clinic from May to July 2018, from which we obtained baseline data. During the single face-to-face counseling session, patients were counselled on lifestyle habits (diet, exercise, vitamin supplements, stress) and medication adherence. They were sent home with printed educational materials and interviewed at 2–3 weeks follow-up via phone call regarding changes made to lifestyle and medication adherence. Information about the purpose of the study and verbal consent was obtained from patients prior to counseling. A team of 3 trained medical students who followed a standardized script provided counseling in clinic. The same medical student research team conducted similarly scripted phone interviews to follow-up with patients after counseling and obtain post-intervention data. The UTSW Institutional Review Board approved our protocol and verbal consent process, and we followed the tenets of the United States Health Insurance Portability and Accountability Act of 1996 (HIPAA) and the Declaration of Helsinki for research involving human subjects.

Lifestyle Counseling Intervention

Prior to their office visit, eligible patients were given lifestyle counseling. The session consisted of first obtaining baseline data unavailable in the EMR (education, employment status, family history of glaucoma, stress level, medication adherence). Then, a comprehensive lifestyle counseling was conducted by going over a pamphlet with pictures of recommended fruits and vegetables in correct portion sizes, stress control strategies, exercise frequency and duration, OTC vitamins, and tips for medication adherence. These counseling categories were selected based on data that these variables may affect glaucoma disease onset and progression.²⁸ The script for the intervention is shown in Table 1.

Patient Selection

Eligibility criteria included a diagnosis of OHT or mild/moderate/severe POAG and prescription of at least one glaucoma medication. Patients with OHT were included if they had no visual field (VF) defects and at least one of the following: IOP \geq 21 mmHg, suspicious optic discs, anti-glaucoma medication prescription. Patients were diagnosed with POAG if they had suspicious optic discs, open iridocorneal angles confirmed by gonioscopy, and VF defects characteristic of glaucoma. Suspicious optic discs were defined as noticeable cupping, neuroretinal rim

Table 1 Lifestyle Counseling Intervention

Variable	Counseling Script
Medication Adherence	Continued encouragement of daily drop use Use drops before 7 AM if possible Use drops the same time everyday Wait at least 5 minutes between drops Match the timing of drops with meals, with use by 8–9 PM
Diet	Do you eat a balanced diet daily? Do you eat processed foods daily? Do you eat fast food daily? How many servings (1/2 cup) of fruit/vegetables do you eat per day?
Vitamin	What vitamins do you take?
Exercise	Do you exercise at least 30 min/day for ≥ 5 days a week?
Stress	Suggested stress reducing measures: Exercise Yoga Meditation, deep breathing Improving sleep patterns Plant based diet (reducing processed, animal-based products)

thinning or notching, or a retinal nerve fiber layer (RNFL) defect characteristic of glaucoma. Patients with POAG were separated into mild, moderate, or severe, based on their VF mean deviation (MD) of less than -6 dB and -12 dB and greater than -12 dB, respectively, using the modified Hodapp-Parrish-Anderson classification.²⁹ Exclusion criteria included age less than 18 years, secondary glaucoma, acute angle closure glaucoma, recent glaucoma or cataract surgery, significantly restricted mobility, and memory/cognitive impairment (Figure 1).

Data Collection

Data collected included age, race, sex, education, employment, body mass index (BMI), family history of glaucoma, type of glaucoma, number of prescribed glaucoma medications, and information on missed drops. Patients were also questioned on reasons for missing drops such as forgetfulness, cost, side effects, and physical difficulties.

The validated perceived stress scale was used to measure patient stress scores.³⁰ Patients were asked 10 questions that addressed how often they felt a negative emotion in the past month, such as anxiety, inability to control important things in their life, and anger because of things

that had happened outside of their control. Possible responses were never (0), almost never (1), sometimes (2), fairly often (3), and very often (4). Answers were summed into a perceived stress score from 0 to 40.

The 2–3-week follow-up phone interview asked about changes made to the patients' diet, vitamin intake, exercise, stress level, and medication adherence in response to the previous lifestyle counseling session. Responses were categorized into more than before, same as before, and less than before. This interview was primarily for data collection and was not a second intervention.

Variables

Medication adherence was defined as whether patients had ever missed their drops in the past month and dichotomized into a yes/no variable. Age was left a continuous variable, while all other variables were coded as categorical variables. Sex was dichotomized into male and female. Given sample size restrictions, certain variables had to be stratified into larger groups, such as race (white, black, other), education (high school or less, college or more), employment (unemployed, employed), and stress level (mild stress, moderate stress and high stress). For stress level, a perceived stress score from 0 to 13 was defined as mild stress, 14–26 as moderate stress, and 27–40 as high stress. No patient fell within the high-stress category (Table 2).

Statistical Analysis

Study data were collected and managed using REDCap electronic data capture tools hosted at UTSW³¹ and de-identified according to IRB protocol. From an exported Excel data file, one randomized eye per patient was selected for analysis using the Microsoft Excel³² (Microsoft Corporation in Redmond, Washington) random number function. R software (R foundation in Vienna, Austria) was used for the following statistical analyses.³³ First, a contingency table was calculated for Fisher's Exact Test to analyze disease group differences in medication adherence at baseline. A Cochran Armitage Trend Test was then performed using this contingency table to analyze the trend in adherence across glaucoma disease groups. A second contingency table was tabulated to analyze differences in medication adherence between baseline and follow-up. Finally, predictive factors (age, race, sex, education, employment, BMI, family history of glaucoma, stress level, disease category, and number of prescribed medications) of medication adherence were

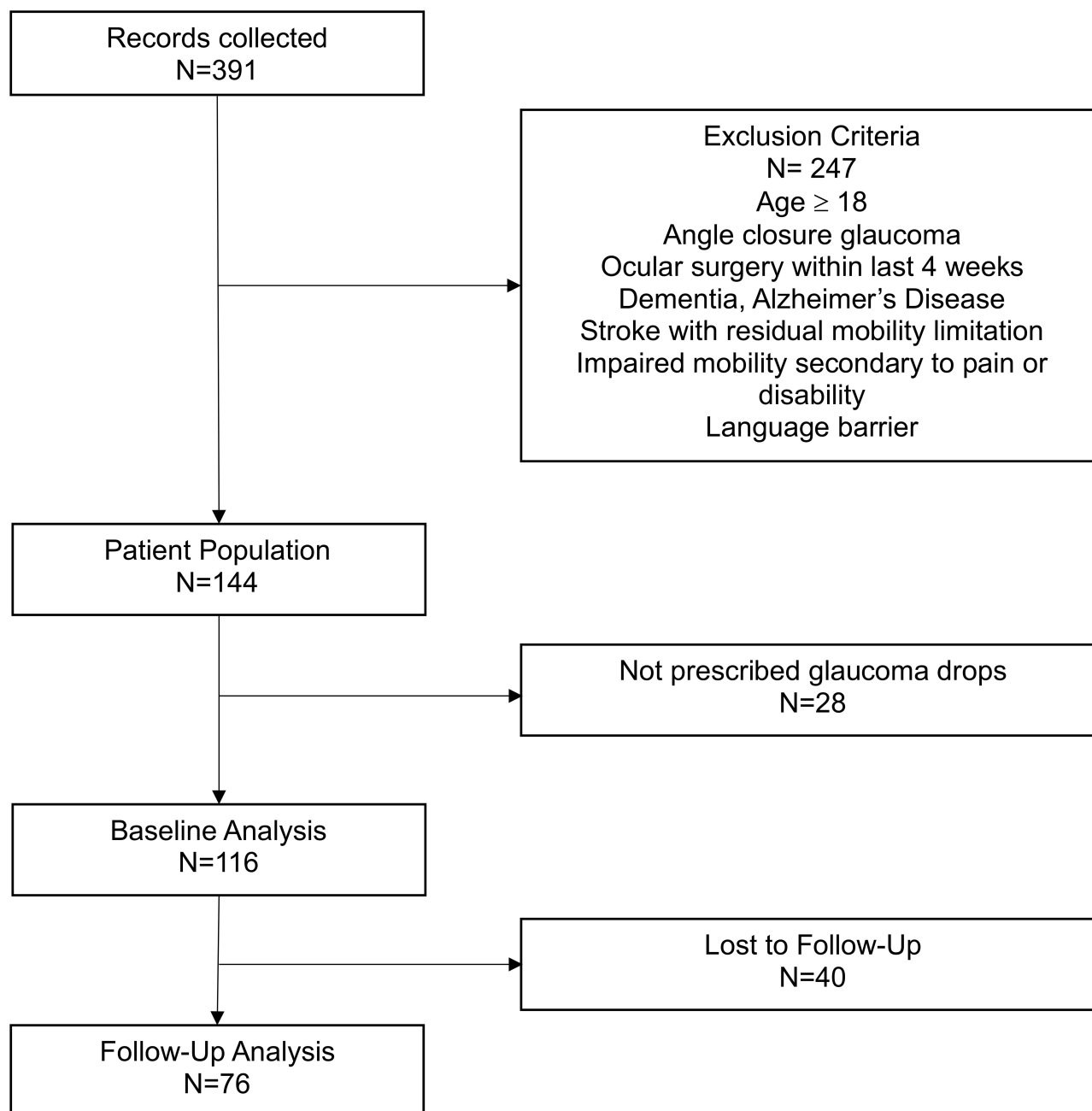


Figure 1 Study flow diagram.

analyzed using multivariate logistic regression. An alpha value of 0.05 was used as the cutoff for statistical significance.

Results

Patient Demographics

Of 116 patients, there were 33 patients with OHT and 83 patients with POAG (28 mild, 39 moderate, and 16 severe). At baseline, 40.5% of patients were found to be non-adherent with their medication regimen. Mean age

was 67.6 years \pm 11.2, 43.1% were white, 48.3% male, 70.7% college educated, and 50.9% employed. Mean BMI was 28.3 \pm 6.4, 62.1% had a family history of glaucoma, 70.7% were found to have mild stress, and 61.2% had 2 or more prescribed glaucoma medications. There were no statistically significant differences in age, race, sex, education, employment, mean BMI, family history of glaucoma, stress level, disease category, and number of prescribed medications between non-adherent and adherent patients (Table 3).

Table 2 Data Collection

Variable	Question	Analysis
Age	Retrieved from EMR	Mean (yr)
Race	Retrieved from EMR	White, Black, Other
Sex	Retrieved from EMR	Male Female
Education	Highest level of education?	High School or Less College or More
Employment Status	Employed/Unemployed/Retired/Disability/Homemaker	Unemployed Employed
Body Mass Index (BMI)	Retrieved from EMR	Mean (kg/m ²)
Family History of Glaucoma	Any family histories were noted	No Yes
Stress Level	Perceived Stress Scale (ref)	Mild Moderate Severe
Disease Category	Retrieved from EMR, one eye randomized	OHT Mild POAG Moderate POAG Severe POAG
Number of Prescribed Medications	Retrieved from EMR	1 2 3 or more
Medication Adherence	Does patient miss drops?	Yes – nonadherent No – Adherent

Table 3 Characteristics of Patient Population

Variables	Nonadherent	Adherent	p
	N=47 (40.5%)	N=69 (59.5%)	
Age, yr mean	68 ± 11.7	67 ± 10.9	0.77
Race			0.46
White	17 (36.2)	33 (47.8)	
Black	19 (40.4)	23 (33.3)	
Other	11 (23.4)	13 (18.8)	
Sex			1.00
Male	23 (48.9)	33 (48.5)	
Female	24 (51.1)	35 (51.5)	
Education			0.91
High School or Less	13 (27.7)	21 (30.4)	
College or More	34 (72.3)	48 (69.6)	
Employment			0.55
Unemployed	21 (44.7)	36 (52.2)	
Employed	26 (55.3)	33 (47.8)	
BMI, kg/m ² mean	28 ± 5.7	29 ± 6.9	0.31
Family Hx of Glaucoma			0.51
No	20 (42.6)	24 (34.8)	
Yes	27 (57.4)	45 (65.2)	
Stress Level			0.69
Mild	36 (78.3)	46 (73.0)	
Moderate	10 (21.7)	17 (27.0)	
Disease category			0.13
OHT	19 (40.4)	14 (20.3)	
Mild POAG	10 (21.3)	18 (26.1)	
Moderate POAG	13 (27.7)	26 (37.7)	
Severe POAG	5 (10.6)	11 (15.9)	
Number of Prescribed Medications			0.95
1	18 (38.3)	27 (39.1)	
2	17 (36.2)	23 (33.3)	
3 or more	12 (25.5)	19 (27.5)	

Abbreviations: BMI, body mass index; POAG, primary open angle glaucoma; OHT, ocular hypertension.

Baseline

The Fisher’s Exact Test revealed no significant differences in medication adherence between glaucoma disease categories (Table 4). However, the Cochran Armitage Trend test did reveal that there was a slightly significant trend of increasing medication adherence as disease status progressed (p=0.055).

The multivariate logistic regression with dichotomous outcome of medication adherence showed that compared to patients with OHT, patients with moderate POAG were

4.14 times more likely (95% CI 1.33–14.24) to be adherent to their medication regimen, while patients with severe POAG were 6.42 times more likely (95% CI 1.28–37.67). These findings, although significant, did have wide confidence intervals. Other factors, such as age, race, sex, education, employment, BMI, family history, stress level, and number of prescribed medications, did not influence the odds of medication adherence (Table 5).

Table 4 Fisher's Exact Test for Independence Adherence by Disease Category

Contingency Table					
Variable		OHT	Mild POAG	Moderate POAG	Severe POAG
		N=33 (%)	N=28 (%)	N=39 (%)	N=16 (%)
Adherence	Yes	14 (42.4)	18 (64.3)	26 (66.7)	11 (68.7)
	No	19 (57.6)	10 (35.7)	13 (33.3)	5 (31.3)
Fisher's Exact Test		p-value = 0.142			
Cochran Armitage Trend Test		z-statistic = -1.916 p-value = 0.055			

Abbreviations: OHT, ocular hypertension; POAG, primary open angle glaucoma.

Follow-Up

Of the 116 patients, 76 (65.5%) were reached by phone call at the 2–3 weeks post intervention. At baseline 62.2% of the followed population reported adherence while after intervention 78.9% reported medication adherence, an increase of 16.7%. In the population, we lost to follow-up (40 patients), 52.5% reported medication adherence and demographics were similar to followed patients. A contingency table calculating the number adherent at baseline and at second encounter was created, and McNemar's Test was performed (Table 6). Among 76 patients, 17 (22.4%) patients who were initially not adherent at baseline became adherent to their medication regimens following lifestyle counseling ($p=0.017$). Furthermore, of those whom follow-up was obtained, 29% reported improved diet while 68% remained the same and 24% reported improved exercise while 67% remained the same. In our sample, we observed that almost one-third of our patients improved their diet and exercise habits following lifestyle counseling.

Discussion

In this prospective study, we found that comprehensive lifestyle counseling inclusive of diet, exercise, vitamin intake, stress level, and medication adherence was successful in improving glaucoma medication adherence at a 2–3 week follow up (62.5% adherent at baseline and

Table 5 Logistic Regression Results

	Adherence		
	OR	95% CI	
N=107*			
Age, yr	0.97	(0.92,	1.01)
Race, Other [†]			
White	2.10	(0.69,	6.54)
Black	1.05	(0.32,	3.46)
Sex, Male [†]			
Female	0.96	(0.38,	2.41)
Education, High School or Less [†]			
College or more	0.79	(0.28,	2.15)
Employment, Unemployed [†]			
Employed	0.68	(0.25,	1.82)
BMI, kg/m ²	1.05	(0.98,	1.13)
Family Hx, No Hx of Glaucoma [†]			
Hx of Glaucoma	1.43	(0.59,	3.50)
Stress Level, Mild Stress [†]			
Moderate Stress	1.53	(0.56,	4.35)
Disease Category, OHT [†]			
Mild POAG	2.38	(0.76,	7.86)
Moderate POAG	4.14	(1.33,	14.24)
Severe POAG	6.42	(1.28,	37.67)
Number of Prescribed medications, I [†]			
2 medications	0.66	(0.23,	1.82)
3 or more medications	0.53	(0.15,	1.75)

Notes: *n=9 with missing covariate values were dropped from analysis. [†]Reference category

Abbreviations: OR, adjusted odds ratio; OHT, ocular hypertension; POAG, primary open angle glaucoma; BMI, body mass index; Hx, history.

78.9% adherent at follow-up). Almost one-quarter of the patients reached for follow-up who were previously non-adherent became adherent following lifestyle counseling. The significant trend between more progressive glaucoma disease status and medication adherence and an increased odds of adherence for patients with moderate and severe POAG compared to patients with OHT may indicate the benefits of counseling on medication adherence. It is possible that these patients may be followed more closely in clinic and thus receive more tailored counseling from the physician or that these patients who have decreased vision status secondary to their glaucoma progression are more cognizant of adverse outcomes associated with a lack of medication adherence. Furthermore, counseling that incorporated diet, exercise, and stress control strategies may

Table 6 McNemar's Test for Follow-Up Adherence

Contingency Table		Adherence at Second Encounter N=76 (%)	
Variable		Yes	No
Adherence at First Encounter N=76 (%)	Yes	43 (56.6)	5 (6.6)
	No	17 (22.4)	11 (14.5)
Became Noncompliant		5 (6.6%)	
Became Compliant		17 (22.4%)	
Odds Ratio		0.29 (0.085, 0.831)	
p-value		0.017	

have helped patients to become motivated to pursue more healthier lifestyles overall. Being able to see and feel the effects of a healthier lifestyle combined with the providers' attention to their overall health status may be a powerful motivation for patients to adhere to their medication regimen.

Contrary to what is reported in the literature, our study did not find a significant difference in medication adherence based on age,^{8,34} race,³⁵ and education.³⁶ In our sample, we did not find these predictive factors of non-adherence. Moreover, we did not find high stress levels to be a predictor of low adherence, although previous studies have found depression and living alone to be significant risk factors.³⁷ It is possible that our small sample size masked differences. Additionally, our population was drawn from a single academic center and the patients of one faculty physician which may also have affected our findings.

Recent studies have found several lifestyle modifiable factors limit glaucoma disease progression, such as smoking cessation, exercise, limit of caffeine intake, control of diabetes, consumption of antioxidants, balanced diet of fruits and vegetables, and maintenance of a healthy weight.^{28,38–40} Given the multifactorial causes of glaucoma, a comprehensive approach that focuses on other aspects of glaucoma disease management including eating a healthy diet, exercising, managing stress, and

supplementing with vitamins may not only improve medication adherence but also ultimately lead to the clinically important outcome of improved IOP.

This study was not without limitations. Self-reported responses to medication adherence were used as our primary outcome, a measure that has been documented previously as being often overestimated.⁴¹ Patients were aware they were part of a research study analyzing the effects of lifestyle counseling on glaucoma. This most likely contributed to the Hawthorne effect,⁴² artificially increasing adherence to medication regimens. Our sample size was limited due to an extensive exclusion criteria and time restrictions for primary data collection, impacting the generalizability of our results. Additionally, we were not able to assess IOP at follow-up as it was conducted over the phone and not in clinic. Furthermore, due to our research team being primarily composed of medical students with limited schedules, we were not able to assess the effects of a more continuous intervention with repeated lifestyle counseling sessions and long-term follow-up. Pursuing a more long-term follow-up rather than 2–3 weeks would have elucidated more information regarding the endurance of medication adherence improvement.¹¹

Despite these limitations, our study proved that lifestyle counseling sessions significantly improved glaucoma medication adherence and opens the door for a more holistic approach towards patient counseling. Many ophthalmologists and glaucoma specialists are already counseling their patients regarding lifestyle choices. This study is the first study to provide statistical evidence that lifestyle counseling increased medication adherence. Current epidemiological studies have attempted to stratify the risk for glaucoma onset by analyzing the effects of certain lifestyle risk factors such as smoking, diet, and exercise on the IOP of healthy subjects.²⁸ More research is needed on whether lifestyle changes in conjunction with medication adherence can improve or prevent progression of glaucoma in patients who are already suffering from the condition.

Lifestyle counseling sessions inclusive of diet, exercise, stress, vitamin intake, and medication improved medication adherence in glaucoma patients. This study was limited by self-reported outcome measures and sample size constraints. Future studies should analyze whether lifestyle counseling improves medication adherence in a larger multi-center study population as well as long-term clinical outcomes, such as continued medication adherence and improved IOP.

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Disclosure

The authors report no conflicts of interest in this work.

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