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Eye Drop Instillation Success and Hand Function in Adults with Glaucoma: A Pilot Study

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38	Running head: Hand Function and Eye Drop Success in Adults with Glaucoma

39 Abstract

40 Purpose: To investigate hand function and eye drop instillation success in adults with and

- 41 without glaucoma.
- 42
- 43 Design: Cross-sectional pilot study
- 44

Subjects: Adults aged 65+ with glaucoma who use eye drops daily, and adults aged 65+ without
 glaucoma who do not regularly use eye drops.

47

Methods: Hand function was evaluated using the Saehan Hydraulic Pinch Gauge, Jamar Hand
Dynamometer, Grooved Pegboard Test (GPT), Arthritis Hand Function Tests (AHFT), SemmesWeinstein Monofilament Test, and a tactile pattern recognition device. Eye drop instillation
success was analyzed through videography.

52

53 *Main Outcome Measures*: Proportion of successful eye drop instillation trials assessed by: 1)

54 overall success in getting at least one drop in the eye; 2) average number of drops dispensed; 3)

55 bottle tip contact with the eye, eyelashes, or skin. Hand function measures: mean pinch and grip

strength, GPT and AHFT completion times, smallest detected monofilament, tactile pattern
 identification time and accuracy.

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60 Results: 25 participants with glaucoma and 79 participants without glaucoma were

61 included. There was no difference in the proportion of trials where at least one drop was

62 successfully instilled, disregarding bottle tip contact (glaucoma: 0.95, non-glaucoma: 0.91;

63 p=0.88). Participants with glaucoma were more likely to make contact between the bottle tip and

the eye, eyelashes, or skin compared to those without (glaucoma: 0.49, non-glaucoma: 0.28;

p=0.01). Participants with glaucoma dispensed a similar number of drops as those without

66 (glaucoma: 1.37, non-glaucoma: 1.46; p=0.47). Participants with glaucoma had significantly

reduced pinch strength (glaucoma: 4.8 kg, non-glaucoma 6.1 kg, p=0.01), grip strength
(glaucoma: 23.4 kg, non-glaucoma: 27.7 kg, p=0.02), longer completion times for the GPT

(glaucoma: 25.4 kg, non-glaucoma: 27.7 kg, p=0.02), longer completion times for the GP1
 (glaucoma: 113.5 seconds, non-glaucoma: 85.5 seconds, p=0.02) and specific AHFT tasks of

fastening/unfastening buttons (glaucoma: 36.6 seconds, non-glaucoma: 27.7 seconds, p=0.03)

and pinning/unpinning safety pins (glaucoma: 35.4 seconds, non-glaucoma: 27.3 seconds,

p=0.02), and worse tactile acuity on monofilament (p=0.04) compared to participants without

73 glaucoma.

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76 *Conclusions*: Despite hand function deficits, in this exploratory pilot study, adults with glaucoma

77 demonstrated eye drop instillation success comparable to those without glaucoma, though with

higher rates of bottle tip contact with the eye, skin or eyelashes, suggesting an increased risk of

79 potential eye drop bottle contamination. These findings suggest that though regular practice may

80 help mitigate the effects of hand function deficits on the targeted activity of getting the eye drop

81 in the eye, the deficits may make this activity difficult.

82 Introduction

Glaucoma is a leading cause of irreversible blindness worldwide.¹ The global prevalence of 83 glaucoma is expected to increase from 76 million cases in 2020 to approximately 111.8 million 84 by 2040 in individuals aged 40-80 years.² The primary management for glaucoma involves the 85 daily application of medicated eye drops to lower intraocular pressure.³ However, adherence to 86 87 the medication regimen remains a significant challenge, with rates of non-adherence ranging from 30% to 80% among patients.^{4,5} Poor adherence to eve drop medications significantly 88 impairs treatment effectiveness and accelerates disease progression. Notably, adherence rates 89 90 below 80% are associated with an increased risk of severe visual field deficits and progressive visual field loss.^{6,7} 91

Factors contributing to non-adherence include forgetfulness, side effects, medication 92 costs, and difficulties in instilling eye drops, among others.⁸ Even among individuals attempting 93 to adhere to their prescribed eye drop regimen, a significant portion of adults with glaucoma still 94 face challenges in self-administering their medication.^{9,10} Approximately one third of glaucoma 95 patients are unable to instill a single drop into their eye effectively,¹¹ with older patients facing 96 greater difficulties.¹¹⁻¹⁴ Poor instillation technique may be associated with visual impairments¹⁴ 97 and physical limitations,¹⁵ which can interfere with both squeezing the bottle and accurately 98 targeting where the drop lands. 99

Hand function, essential for tasks requiring fine sensorimotor control such as instilling eye drops, is known to decline with age.^{16,17} Given that the average age of onset for glaucoma is 66 years,¹⁸ adults with glaucoma may experience age-related declines in hand function, which could affect their ability to administer eye drops. While previous studies have highlighted the instillation challenges faced by adults with glaucoma,^{11,19} there remains a significant gap in

understanding how age-related sensorimotor deficits impact the ability to self-administer eyedrops.

107 Comparative studies evaluating eye drop instillation success between adults with 108 glaucoma who use chronic daily eye drops and adults without glaucoma who do not use daily 109 eye drops are limited, and are needed in order to understand the hand function learning curve 110 associated with instilling drops. Understanding how hand function declines associated with aging 111 may impact eye drop instillation success could potentially inform how to teach the best 112 techniques to promote successful eye drop instillation when someone is newly diagnosed with 113 glaucoma. A previous study compared eye drop instillation success among visually impaired 114 older adults with either glaucoma or retinal disease, finding that both groups, who regularly used eye drops, faced significant challenges in self-administering their medication.²⁰ To date, only one 115 116 study has specifically compared eye drop instillation success between older adults with glaucoma 117 who use daily eye drops and adults of similar age without glaucoma who do not use eye drops,²¹ 118 which found that the participants with glaucoma had lower rates of successful instillation. Our 119 study seeks to build upon this foundation by comparing hand function and eye drop instillation 120 success between older adults with and without glaucoma. We hypothesize that those with 121 glaucoma will have worse hand function and lower eye drop instillation success than those 122 without glaucoma. We used standardized measures from the motor control literature to assess hand function.²²⁻²⁶ This investigation is exploratory and will generate hypotheses as to how 123 124 sensorimotor deficits associated with aging may impact eye drop instillation success.

125 Methods

126 **Participants**

127 The study sample included 25 adults \geq 65 years of age with any type of glaucoma using medicated drops, and 79 adults \geq 65 years of age without glaucoma who do not regularly use eye 128 129 drops, defined as using eye drops less than once per week. As this was a pilot study, the sample 130 size was based on feasibility, with the primary focus on generating hypotheses for future studies. 131 We excluded individuals with a history of allergic reaction to over-the-counter eye drops and any 132 individuals with glaucoma who did not self-administer their drops. Approval for this study was 133 obtained from the Michigan Medicine Institutional Review Board, and written informed consent was obtained for each participant. The study adhered to all the tenets of the Declaration of 134 135 Helsinki. All participants were recruited from the University of Michigan research participant 136 registry (https://umhealthresearch.org) and via community and university flyer postings. Participants were not recruited from eye clinics. 137

138 Standardized Assessments

139 Participants completed standardized assessments of visual acuity, physical activity, grip and 140 pinch strength, dexterity, and tactile registration. Visual acuity was evaluated using a 141 standardized Snellen Chart, and better-eye scores were converted to logMAR for analysis. 142 Participants were assessed for physical activity using the Physical Activity Scale for the Elderly (PASE).²⁷ Grip and three-point pinch strength were measured three times each in the dominant 143 144 hand using a Jamar Hand Dynamometer and a Saehan Hydraulic Pinch Gauge. Assessments of 145 dexterity included both the Jamar Grooved Pegboard Test (GPT) and the Arthritis Hand Function Test (AHFT). The GPT, one of many pegboard tests, is a commonly used standardized test of 146 manual dexterity²⁸ that is particularly effective in correlating with activities of daily living and 147 quantifying changes in hand function among older adults.²²⁻²⁴ In this test, participants are timed 148

149 while using their dominant hand to manipulate grooved pegs into a board with holes of varied orientations. The AHFT, originally designed for patients with arthritis and related conditions, 150 151 includes simulated functional tasks that are commonly encountered in daily life, such as lacing 152 shoes and buttoning clothing. This test was selected because it requires the dexterous use of both hands, a skill required for eye drop bottle use.^{25,26} In this test, participants were timed while 153 completing bimanual simulated activities of daily living, including buttoning a button board, 154 lacing up a shoe and tving a bow, pinning safety pins to fabric, placing coins into a slot, and 155 cutting a piece of clay with a fork and knife. Tactile registration was assessed in the index and 156 ring finger of the dominant hand using the Semmes-Weinstein Monofilament Test (SWMT).²⁹ In 157 the SWMT, participants were asked to close their eyes while monofilaments of various sizes 158 were applied to their finger pads and indicate when they perceived a touch. The smallest 159 160 detectable monofilament size was recorded for each finger.

161 Experimental Assessments

162 *Tactile Discrimination*

Tactile discrimination was evaluated using a custom-designed pattern recognition device. While seated in front of the device, participants were given five seconds to feel a visually concealed pattern of raised dots with one finger. After five seconds, participants removed their finger from the pattern and used a touchscreen display to select the pattern they felt from a choice of four patterns. Participants completed four trials each with their dominant index and ring finger, for a total of eight trials in the dominant hand. Patterns were randomly selected for each trial from a

- set of twelve unique patterns. Outcome measures included the amount of time taken to identify
- 170 the perceived pattern and accuracy rate of correct identifications.
- 171 Assessment of Eye Drop Instillation

Participants completed three eye drop instillation trials for each eye across three different 172 postural configurations (seated, standing, and supine), starting with the right eye in each position. 173 During each trial, participants were instructed to administer one eye drop into their eye, 174 175 dispensing as many drops as needed to achieve this goal. Participants used a 15 mL bottle of 176 Refresh Tears Lubricant Eye Drops (Allergan, Irvine, CA, USA) for all trials. No instruction was provided for instillation technique; participants self-selected which hand they preferred to hold 177 178 the bottle with, whether they used their other hand for support, and whether they used a mirror during instillation. Trials were recorded using an iPod touch (7th generation) with high-definition 179 180 video (8-megapixel camera with 1080p HD video recording). The camera was positioned close 181 to the side of the participant's eye for clear visualization. The video recordings were transferred 182 to a laptop computer for review. The recordings were carefully reviewed by pausing, zooming in 183 and, when necessary, slowing down the footage frame by frame to accurately assess the number 184 of drops dispensed and determine if the trial was successful. Each trial video was reviewed by a research associate who did not take the video recording, though they were not masked to 185 participant glaucoma status. The following measures were collected and managed in a secure 186 187 research database (UM1TR004404)^{30,31}: location of eye drop instillation, number of drops used 188 to instill one drop, and whether the tip of the eye drop bottle touched the ocular surface, 189 eyelashes, or skin. In rare cases where the solution was dispensed as a stream instead of 190 individual drops, it was counted as four drops if a clear start and end of the stream could be

determined. A single research associate reviewed each video. To assess the reliability of the eye
drop instillation success measures graded from videos (location of eye drop, number of drops
dispensed, bottle tip contact with the ocular surface, skin, or eyelashes), a 10% random sample of
videos were selected stratified by grader, position, and eye, with an oversampling of videos
where more than one eye drop was dispensed. This sample was re-graded by additional research
associates masked to the initial grading. Inter-grader agreement of measures obtained from
videos was good and ranged from 86.9% to 95.6% (Supplemental Table 1).

198 Statistical Analysis

199 Characteristics of the sample (participant demographics, eye drop instillation success,

200 sensorimotor function) were summarized with descriptive statistics, including mean and standard 201 deviation for continuous measures and frequency and percentage for categorical measures. Six 202 measures of eye drop installation success were calculated, aggregated over all 18 trials, including 203 1) proportion of trials with no bottle tip contact with the eye, eyelashes, or skin, 2) proportion of 204 trials with only one drop dispensed, 3) mean number of drops dispensed to instill a single drop, 205 4) proportion of trials with at least one drop successfully placed in the eye, 5) proportion of trials 206 with at least one drop successfully placed in the eye with no bottle tip contact, and 6) proportion 207 of trails with exactly one drop successfully placed in the eye with no bottle tip contact.

208 Participants with and without glaucoma were compared for differences in demographic

209 characteristics, eye drop installation success, and sensorimotor function with 2-sample t-tests (for

- 210 normally distributed continuous measures), 2-sample Wilcoxon tests (for non-normally
- 211 distributed continuous measures), Chi-square tests (for categorical measures with expected
- counts \geq 5), or Fisher's exact tests (for categorical measures with expected counts <5). Linear

regression was used to estimate the differences in eye drop success and sensorimotor function between participants with and without glaucoma, adjusting for age and diabetes. Model results are reported with regression estimates and 95% confidence intervals (CI). Due to the limited variability in monofilament detection in the sample, models adjusted for age and diabetes could not be performed for this outcome. SAS version 9.4 (SAS Institute, Cary NC) was used for all statistical analysis.

219 **Results**

A total of 25 glaucoma participants and 79 participants without glaucoma were

studied. Participants with glaucoma reported using 1 (n=9, 36.0%), 2 (n=11, 44.0%), or 3 (n=5,

222 20.0%) daily eye drop medications to control their disease, for durations that were less than 1

223 year (n=1, 4.0%), 1-5 years (n=10, 40.0%), 6-10 years (n=7, 28.0%), or >10 years (n=7,

224 28.0%). Those without glaucoma reported using over the counter eye drops either weekly (n=6,

225 7.6%), monthly (n=11, 13.9%), rarely (n=31, 39.2%), or never (n=31, 39.2%). Glaucoma

226 participants (n=25) had significantly worse vision than non-glaucoma participants (n=79).

227 Specifically, glaucoma participants had average logMAR visual acuity of 0.45 (Snellen

equivalent 20/57; median 20/30), whereas non-glaucoma participants had average logMAR

visual acuity of 0.12 (Snellen equivalent 20/26; median 20/25), p=0.0007. No significant

230 demographic differences were found between participants with versus without glaucoma for age

231 (mean of 75.3 years [SD=1.8] vs 73.3 years [SD=5.8], p=0.15), sex (45.8% male vs 44.9,

232 p=0.93), race (80.0% White vs 85.5%, p=0.50), ethnicity (0.0% Hispanic vs 1.3%, p=1.00),

education (p=0.42), and income (p=0.81) (Table 2). Further, no significant difference between

groups was found with respect to physical activity (mean PASE score 150.6 [SD=73.7] vs 155.7

235	[SD=73.3], p=0.76). However, a larger percentage of glaucoma participants reported having
236	diabetes than participants without glaucoma (20.8% vs 5.1%, p=0.03). For the reduced sample
237	with tactile measures, no significant demographic differences were observed between those with
238	and without glaucoma (Supplemental Table 3).

Participants showed differing success with eye drop instillation measures (Table 4). 239 240 Bottle tip contact with the eye, eyelashes, or skin was the largest hurdle to successful eye drop 241 instillation with an average rate of non-contact of 0.51 for glaucoma participants (SD=0.55, 242 median=0.67), which was significantly worse than the average rate of 0.72 for participants 243 without glaucoma (SD=0.36, median=0.94; p=0.01; Figure 1A). The rate of dispensing only one 244 drop from the bottle was similar between glaucoma (mean=0.76, SD=0.17, median=0.78) and 245 non-glaucoma groups (mean=0.72, SD=0.23, median=0.72; p=0.59; Figure 1B). There was no 246 difference in the number of drops dispensed to successfully instill a single drop in the eye among 247 those with and without glaucoma (glaucoma: mean=1.37, SD=0.31, median=1.28; non-248 glaucoma: mean=1.46, SD=0.41; median=1.39; p=0.47; Figure 1C). The average proportion of 249 trials that glaucoma participants successfully placed at least one drop in their eye was high at 250 0.95 (SD=0.07, median=0.94), which was similar to the average proportion for participants 251 without glaucoma of 0.91 (SD=0.17, median=1.00; p=0.88; Figure 1D). However, when 252 accounting for bottle tip contact, these rates decreased and were significantly different between 253 groups such that those with glaucoma had an average rate of instilling at least one drop in the eye 254 without contacting the bottle tip of 0.49 (SD=0.42, median=0.56) compared to 0.68 for those 255 without glaucoma (SD=0.36, median=0.83; p=0.03; Figure 1E). These rates reduced futher when 256 also requiring only a single drop be instilled, although no significant difference was observed 257 between those with and without glaucoma (glaucoma: mean=0.39, SD=0.36, median=0.44; non-

258	glaucoma: mean=0.53, SD=0.34, median=0.56; p=0.06; Figure 1F) After adjusting for age and
259	diabetes status, though the directionality of the estimated differences in instillation success
260	between participants with and without glaucoma remained the same, the results were no longer
261	statistically significant (Table 5).

Several differences in sensorimotor function were observed between participants with 262 263 and without glaucoma (Table 4). Participants with versus without glaucoma showed significantly 264 reduced pinch force (mean of 4.8 kg [SD=4.8] vs 6.1 kg [SD=2.1], p=0.01, Figure 2A) and grip 265 strength (mean of 23.4 kg [SD=10.7] vs 27.7 kg [SD=10.6], p=0.02, Figure 2B). Additionally, 266 participants with glaucoma took significantly longer than those without glaucoma to complete the GPT (mean of 113.5 seconds [SD=52.3] vs 85.5 seconds [SD=23.4], p=0.02, Figure 2C) and 267 268 individual activities from the AHFT including fastening/unfastening 4 buttons (mean of 36.6 seconds [SD=19.0] vs 27.7 seconds [SD=8.2], p=0.03), and pinning/unpinning 2 safety pins 269 270 (mean of 35.4 seconds vs 27.3 seconds [SD=7.4], p=0.02, Figure 2D). No significant differences 271 were observed between participants with and without glaucoma for time to complete the tasks of 272 lacing a shoe and tying a bow, picking up and manipulating 4 coins into a slot, and cutting putty 273 into 4 pieces with a knife (all p>0.05, Table 4). A significantly smaller percentage of participants 274 with glaucoma were able to detect the smallest monofilaments compared to those without 275 glaucoma (2.83 monofilament: 8.0% vs 20.3%; 3.61 monofilament: 67.0% vs 77.2%; p=0.04; 276 Figure 2E). Lastly, there was also a marginally significant difference in time to complete tactile 277 testing (Figure 2F) where those with glaucoma took longer than those without glaucoma (mean 278 of 5.1 seconds [SD=2.3] vs 4.2 seconds [SD=1.9], p=0.10), but there was no difference in tactile 279 accuracy (mean of 43.7% accurate [SD=24.1] vs 52.1% [SD=20.9], p=0.20, Table 4).

280	After adjusting for age and diabetes status, differences between participants with and
281	without glaucoma on sensorimotor function measurements remained significant or remained with
282	a trend in the hypothesized direction (Table 5). Participants with glaucoma were estimated to
283	have weaker pinch force than participants without glaucoma by 0.9 kg (95% CI: -1.9 to 0.1,
284	p=0.07), took 22.7 seconds longer complete the GPT (95% CI: 8.2 to 37.3, p=0.003), and took
285	1.0 second longer to complete tactile testing (95% CI: -0.1 to 2.1, p=0.08). Participants with
286	glaucoma also took longer to complete the AHFT, including an estimated additional 6.6 seconds
287	to finish fastening/unfastening 4 buttons (95% CI: 1.2 to 12.1, p=0.02), an additional 5.6 seconds
288	to finish lacing a shoe and tying a bow (95% CI: -0.7 to 11.9, p=0.08), and an additional 7.4
289	seconds to finish pinning/unpinning 2 safety pins (95% CI: 2.3 to 12.6, $p=0.005$)

290 Discussion

291 In this study, we explored specific hand function testing and eye drop instillation success among 292 adults with glaucoma who regularly use medicated eye drops compared to adults without 293 glaucoma who do not use eye drops regularly. Though we had hypothesized that people with 294 glaucoma would have worse hand function and lower success instilling eye drops compared to 295 people without glaucoma, we did not find this in our study. We found that although participants 296 with glaucoma had reduced hand function including decreased grip and pinch strength, decreased 297 manual dexterity, and decreased tactile acuity compared to adults who do not use eye drops 298 regularly, they achieved similar success in eye drop instillation as the non-glaucoma participants, 299 though with a higher incidence of bottle tip contact with the eye, skin or eyelashes, suggesting an 300 increased risk of potential bottle contamination.

301 These findings suggest that the individuals with glaucoma may have developed adaptive302 strategies to compensate for their hand function deficits, potentially through the routine and

303 repetitive task of daily eve drop use. Among glaucoma participants in our study, 96% had been using drops for over a year, and 56% for over six years. It may be possible that regular practice 304 305 of the specific task of eye drop instillation over time enables refinement of skills compared to 306 other activities involving precise control of hand movements. While the transfer of motor skill learning from one arm to the other has been demonstrated,³² it has also been demonstrated that 307 the generalizability of more complex movements involving the coordination of different joints is 308 poor^{33,34} and independent of the intensity of training.³⁵ This literature suggests that learning to do 309 310 a very specific task well, like instilling eye drops, may not generalize to being able to do other 311 tasks requiring good hand function well. These observations support earlier work demonstrating 312 that transfer of motor skill learning is most effective when tasks share similar joint configurations³⁶ rather than end-point control of body segments, as in what is necessary to hold 313 314 the eye drop bottle above the eye and dispense a drop. Thus, in the present study, the ability to 315 successfully instill an eye drop in the eye despite poor hand function may be an example of non-316 transferable task-specific training.

317 The question remains, however, as to why glaucoma patients had poor hand function on 318 standardized measures of dexterity. Both the grooved pegboard test and the arthritis hand 319 function test require visual monitoring of hand movements. Although research on hand function 320 in the glaucoma population is limited, prior research suggests that adults with glaucoma may experience impaired eye-hand coordination compared to adults without glaucoma.^{37,38} Of 321 particular interest, Zwierko et al.³⁷ found that adults with moderate to advanced glaucoma 322 323 exhibited eve-hand coordination deficits that could not be entirely attributed to visual field 324 defects. They suggested that other factors, such as accelerated age-related changes in visuomotor 325 processing and reduced physical activity, may be contributing to these findings. While previous

326 research suggests that adults with glaucoma are less physically active than their non-glaucoma counterparts,³⁹ our study found no significant differences in overall physical activity levels 327 328 between the two groups, though our measure of physical activity was through self-report. The 329 participants with glaucoma did have significantly worse visual acuity than the participants 330 without glaucoma. Poor visual acuity in adults with glaucoma has been associated with restricted participation in activities of daily living,⁴⁰ many of which require skilled hand use. A reduction in 331 332 hand use over time could contribute to diminished hand strength and dexterity, potentially leading to difficulties in tasks requiring fine sensorimotor control, such as manipulating an eye 333 334 drop bottle. It is thus reasonable to hypothesize that the impaired hand function observed in the 335 glaucoma group may reflect a decline in overall hand use due to impairments in visual acuity. This impaired hand function may be why we see an increase in bottle tip contact among those 336 337 with glaucoma alongside the increased success in getting a drop in the eye.

Previous studies have highlighted the challenges faced by adults with glaucoma in self-338 administering eve drops. A review of 15 observational studies indicates that up to 61% of 339 340 glaucoma patients struggle to dispense the correct dosage of one drop, as many as 37% miss the eye entirely, and up to 80% contaminate their eye drop bottles.⁹ A study by Naito et al.²¹ 341 342 compared eye drop instillation success, defined as the deposition of one drop onto the ocular 343 surface on the first attempt without any bottle tip contact with the eye, skin or eyelashes, between 344 volunteers with and without glaucoma. They found that the adults with glaucoma had lower 345 success rates (38.5%) compared to adults without glaucoma (56.5%). Using a similar definition 346 of success, we observed a marginally significant difference between groups in the proportion of 347 trials with instillation of only one drop into the eye and no contact between the bottle tip and eye 348 or ocular adnexae (glaucoma: 0.39, non-glaucoma: 0.53; p=0.06). When success was defined less

349	stringently as the instillation of one or more drops into the eye and no eye drop bottle tip contact,
350	the glaucoma group had significantly fewer successful trials (glaucoma: 0.49, non-glaucoma:
351	0.68; p=0.03). Under our broadest definition of success, getting at least one drop into the eye
352	irrespective of eye drop bottle tip contact, both groups had similar success (glaucoma: 0.95, non-
353	glaucoma: 0.91; p=0.88). Potential contamination of the bottle tip through contact with the eye or
354	ocular adnexae was more prevalent among the glaucoma participants, with a smaller proportion
355	of their trials having no contact (glaucoma: 0.51, non-glaucoma: 0.72; p=0.013). While regular
356	practice of instilling eye drops daily may enable glaucoma participants to administer drops
357	successfully, their ability to perform the finer movements necessary to prevent bottle tip contact
358	may be impaired by their reduced hand function.

Both groups demonstrated comparable success in instilling at least one drop into their 359 360 eye. Participants with glaucoma used an average of 1.37 drops, and non-glaucoma participants used an average of 1.46 drops to successfully instill a single drop into their eye, and this 361 362 difference was not statistically significant (p=0.47). Although the excess usage beyond a single 363 drop for both groups may appear minor, it translates into notable medication waste over time, particularly for patients requiring multiple daily instillations. This can result in a shortage before 364 the next refill is available that will be covered by insurance, leading people to pay out of pocket 365 366 for the medication or have a gap in eye drop usage. Further, this study used a 15mL artificial tear 367 bottle, that was likely more pliable and easier to squeeze than many of the bottles of glaucoma 368 medications, which likely underestimated difficulties with eye drop instillation. Several 369 researchers have demonstrated that the force required to squeeze the different glaucoma medication bottles is highly variable, based on bottle shape, pliability of the bottle material, 370 orientation of the bottle, and the number of drops left in the bottle.^{41,42} Drew and Wolffsohn 371

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372 demonstrated that average pinch grip strength among people with glaucoma was not always sufficiently forceful to dispense a drop from the glaucoma medication bottle, leading them to 373 374 conclude that many glaucoma patients likely struggle with the force required to dispense an eve 375 drop from several of the bottle designs.⁴¹

Considering that hand function continues to decline with advancing age,⁴³ there may be a 376 377 threshold at which the combined effects of hand function deficits and vision loss from glaucoma significantly impair the ability to administer eye drops, thereby affecting the overall 378 effectiveness of the treatment. This highlights the need for developing targeted interventions to 379 380 address instillation challenges, particularly for individuals newly diagnosed with glaucoma, those 381 aged 65 and older who are more likely to experience motor control deficits, and anyone with decreased hand function. Furthermore, as hand function continues to decline with age, it brings 382 383 to light the potential that targeted interventions to improve hand function overall might impact not only the ability to instill eye drops, but also the ability to engage with more activities of daily 384 living. Additionally, identifying older patients with hand function deficits who may struggle with

386 eye drop instillation could help guide clinicians toward considering alternative treatment options 387 such as selective laser trabeculoplasty (SLT) or surgical interventions, which may reduce the 388 drop burden for patients who have difficulty utilizing topical medications.

385

389 The strengths of this study include its targeted focus on older adults and the prospective 390 assessment of hand function using standardized methods. Our study has limitations. Given its 391 exploratory nature, no sample size calculations were conducted, and the study was not powered to detect differences across all outcomes. Because this is a pilot study, the results should be seen 392 393 as hypothesis generating rather than hypothesis testing. Eye drop instillation success was 394 assessed using artificial tear bottles, which typically require less squeezing force than glaucoma

395 medication bottles. The artificial tear bottles utilized in this study had a volume of 15 mL, in contrast to the average 5 mL volume of typical glaucoma medication bottles, which could result 396 397 in the artificial tear bottles being more pliable and underestimating the difficulty people may 398 have instilling glaucoma medication. We did not assess whether non-glaucoma participants faced 399 limitations that would prevent them from self-administering eye drops, an exclusion criterion for 400 the glaucoma participants. Consequently, the glaucoma group may consist of individuals more adept at administering their own drops, while the non-glaucoma group could include those who 401 require assistance. There were few instances where participants dispensed a stream of eye drops 402 403 and the number of drops were not countable, and so we counted these instances as four drops, 404 this involved 2.4% of the 1979 trials. Contrast sensitivity was not assessed, which could have 405 provided further insight into the visual factors affecting hand function. The accessibility of this 406 study was limited to individuals able to drive to the research location, potentially introducing a healthy volunteer bias into our findings, though this would be present across both groups. 407 Additionally, participants were recruited from a restricted geographic area characterized by high 408 409 income and education levels, which does not reflect the broader population. Future studies 410 should address whether these differences exist in more diverse settings.

Our study highlights the critical need to understand the factors that contribute to successful eye drop instillation, with a particular focus on hand function deficits that may impact one's ability to effectively administer eye drops. Such information could be utilized to develop personalized and timely educational interventions aimed at refining eye drop instillation techniques particulary among those newly diagnosed with glaucoma. However, with projected workforce shortages among ophthalmologists,⁴⁴ combined with the aging US population and subsequent increasing prevalence of glaucoma,² ophthalmologists may be constrained in their

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ability to provide such instruction. Novel strategies are needed to address this gap in glaucoma
care. By standardizing assessment and coaching techniques, perhaps ophthalmic technicians
could play a crucial role in filling this gap. Additionally, through standardized assessment, those
with difficulties instilling eye drops could be identified and potentially referred to occupational
therapy for additional coaching and strengthening and/or the physician could identify alternative
approaches for intraocular pressure management.

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552 Figure captions

553

554	Figure 1. Boxplots displaying the distribution of eye drop installation success measures, stratified
555	by glaucoma status, including A) proportion of trials with no eye drop bottle tip contact with the
556	eye, eyelashes, or skin, B) proportion of trials with only one drop dispensed, C) mean number of
557	drops dispensed to instill a single drop, D) proportion of trials with at least one drop successfully
558	placed in the eye, E) proportion of trials with at least one drop successfully placed in the eye
559	with no bottle tip contact, and F) proportion of trails with exactly one drop successfully placed in
560	the eye with no bottle tip contact.
561	Figure 2. Plots (boxplots and bar chart) displaying the distribution of motor function measures,
562	stratified by glaucoma status, including A) Pinch force, B) Grip strength, C) Time to complete
563	the grooved pegboard test, D) Time to complete the arthritis hand function pinning/unpinning 4
564	safety pins test, E) Smallest monofilament detected with the index finger of the dominant hand,
565	and F) Median time to complete tactile testing.
566	

Table 2. Sample descriptives stratified by glaucoma status

	Glaucoma (n=25)		No Glaucoma		
Continuous Variable	Mean (SD)	Median	Mean (SD)	Median	P-value ^a
Age (years)	75.3 (6.2)	74.0	73.3 (5.8)	72.0	0.1503
PASE Score	150.6 (73.7)	147.0	155.7 (73.3)	151.0	0.7623
Categorical Variable	Frequency (Pe	ercent)	Frequency (Pe	ercent)	P-value ^b
Sex - Male	11 (45.8)	35 (44.9)	0.9340
Race					
White	20 (80.0)	65 (85.5)	
Black	4 (16.0))	6 (7.9)		0.5043
Asian	1 (4.0)		5 (6.6)		
Ethnicity - Hispanic/Latino	0 (0.0)		1 (1.3)		1.0000
Education					
High School/GED	3 (12.0))	2 (2.5)		
Vocational/Technical	0 (0.0)		1 (1.3)		
Some College/Associate Degree	5 (20.0))	15 (19.0)	0.4201
College Graduate	7 (28.0))	25 (31.7)	
Graduate/Professional Degree	10 (40.0		36 (45.6)	
Income					
<\$20k	1 (4.6)		3 (4.4)		
\$20k-\$39k	3 (13.6)		14 (20.3)	
\$40k-\$59k	3 (13.6)		8 (11.6))	
\$60k-\$79k	2 (9.1)		8 (11.6))	
\$80k-\$99k	1 (4.6)		8 (11.6)	0.8059	
\$100k-\$119k	5 (22.7)		11 (15.9		
\$120k-\$139k	1 (4.6)		6 (8.7)		
\$140k-\$160k	1 (4.6)		5 (7.3)		
>\$160k	5 (22.7)		6 (8.7)		
Handed					
Right-handed	22 (88.0)	70 (88.6)	
Left-handed	2 (8.0)		5 (6.3)		0.8674
Both	1 (4.0)		4 (5.1)		
Comorbidities					
Cardiac	8 (33.3)		16 (20.8)	0.2071
Vascular	16 (64.0)	37 (48.1)	0.1655
Pulmonary	3 (12.0)		5 (6.4)		0.3981
Neurologic	0 (0.0)		1 (1.3)		1.0000
Endocrine/Diabetes	5 (20.8)		4 (5.1)		0.0314
Renal	3 (12.5)		6 (7.7)		0.4360
Hepatic	0 (0.0)		1 (1.3)		1.0000
Gastrointestinal	0 (0.)		5 (5.1)		0.5704
Mental	1 (4.2)		6 (7.9)	,	1.0000
Osteoarthritis	16 (66.7)	41 (55.4)	0.3311
Immunocompromised	3 (12.0)		8 (10.3)		0.7257
Blood Disorder	1 (4.0)		1 (1.3)		0.4283

PASE, Physical Activity Score for Elderly; GED, General Education Diploma; ^a2-sample t-test, ^bChisquare or Fisher's exact test; Percentages are reported on the non-missing sample

	Glaucoma (n=25)			No Glaucoma (n=79)					
Outcome	n	Mean	SD	Median	n	Mean	SD	Median	p-value ^a
Eyedrop Installation									
Proportion of trials with no contamination	25	0.51	0.44	0.67	79	0.72	0.36	0.94	0.0128
Proportion of trials with only 1 drop dispensed	25	0.76	0.17	0.78	79	0.72	0.23	0.72	0.5899
Average number of drops used to instill one over all trials	25	1.37	0.31	1.28	79	1.46	0.41	1.39	0.4686
Proportion of trials with ≥1 drop in the eye	25	0.95	0.07	0.94	79	0.91	0.17	1.00	0.8829
Proportion of trials with ≥1 drop in the eye & no contamination	25	0.49	0.42	0.56	79	0.68	0.36	0.83	0.0301
Proportion of trials with only 1 drop in the eye & no contamination	25	0.39	0.36	0.44	79	0.53	0.34	0.56	0.0631
Sensorimotor Function				, ,					
Median Pinch Force - Dominant Hand (kg)	25	4.8	2.1	5.0	79	6.1	2.1	6.0	0.0098
Median Grip Strength - Dominant Hand (kg)	25	23.4	10.7	20.0	79	27.7	10.6	24.0	0.0185
Time to complete Grooved Pegboard - Dominant Hand (sec)	23	113.4	52.3	93.0	79	85.8	23.4	80.0	0.0166
AHF Fasten/Unfasten 4 buttons (sec)	25	36.6	19.0	30.0	79	27.7	8.2	26.0	0.0324
AHF Lacing shoe and tying bow (sec)	24	49.6	18.1	44.0	79	42.2	12.0	39.0	0.0718
AHF Pinning/Unpinning 2 safety pins (sec)	23	35.4	17.5	32.0	79	27.3	7.4	26.0	0.0214
AHF Picking up/Manipulating 4 coins into a slot (sec)	24	14.4	5.3	12.0	79	12.6	4.6	12.0	0.1510
AHF Cutting putty into 4 pieces with knife (sec)	23	13.3	4.5	12.0	77	12.6	3.5	12.0	0.6924
AHF Sum all times (sec)	23	143.2	49.2	127.0	77	121.3	23.3	120.0	0.0647
Median time to complete tactile testing (sec)	20	5.1	2.3	4.4	41	4.2	1.9	3.9	0.0986
Percent tactile accuracy	20	43.7	24.1	40.2	41	52.1	20.9	50.0	0.1972
Э — Э — Э — Э — Э — Э — Э — Э — Э — Э —	Freq	uency (C	olumn F	Percent)	Freq	uency (Co	olumn P	ercent)	p-value ^b
Smallest Monofilament detected - Index finger, Dominant Hand									
'2.83		0 (0.0)			5 (6.3)		
'3.61		22 (88.0)			70 (88.6)		0 1761
'4.31		2 (8.0)			4 (5.1)		0.1701
'4.56		1 (4.0)			0 (0.0)		
Smallest Monofilament detected - Ring finger, Dominant Hand									
'2.83		2 (8.0)			16 (20.3)		
'3.61		19 (76.0)			61 (77.2)		0.0360
'4.31		3 (1	2.0)			2 (2.5)		0.0000
'4.56		1 (4.0)			0 (0.0)		

	Age and Diabetes Adjusted Estimate				
Outcome	Estimate	95% CI	P-value		
Eyedrop Installation					
Proportion of trials with no contamination	-0.13	-0.31, 0.05	0.1506		
Proportion of trials with only 1 drop dispensed	0.05	-0.05, 0.16	0.3257		
Average number of drops used to instill one over all trials	-0.17	-0.30, 0.08	0.2454		
Proportion of trials with ≥1 drop in the eye	0.07	-0.01, 0.14	0.0747		
Proportion of trials with ≥1 drop in the eye & no contamination	-0.11	-0.28, 0.07	0.2228		
Proportion of trials with only 1 drop in the eye & no					
contamination	-0.07	-0.23, 0.09	0.4098		
Sensorimotor Function					
Median Pinch Force - Dominant Hand (kg)	-0.9	-1.9, 0.1	0.0661		
Median Grip Strength - Dominant Hand (kg)	-3.7	-8.9, 1.4	0.1527		
Time to complete Grooved Pegboard - Dominant Hand (sec)	22.7	8.2, 37.3	0.0026		
AHF Fasten/Unfasten 4 buttons (sec)	6.6	1.2, 12.1	0.0182		
AHF Lacing shoe and tying bow (sec)	5.6	-0.7, 11.9	0.0787		
AHF Pinning/Unpinning 2 safety pins (sec)	7.4	2.3, 12.6	0.0049		
AHF Picking up/Manipulating 4 coins into a slot (sec)	1.2	-0.9, 3.4	0.2649		
AHF Cutting putty into 4 pieces with knife (sec)	0.6	-1.1, 2.3	0.4931		
AHF Sum all times (sec)	17.8	3.2, 32.4	0.0177		
Median time to complete tactile testing (sec)	1.0	-0.1, 2.1	0.0760		
Percent tactile accuracy	-8.5	-20.7, 3.7	0.1671		

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Despite reduced hand function, older adults with glaucoma achieved similar eye drop instillation success to those without glaucoma, but they had a higher incidence of bottle tip contact with the ocular surface, skin or eyelashes, suggesting an increased risk of potential contamination. Hand function deficits that occur as people age can pose challenges to eye drop instillation.