Patients' Perspectives on Alternative Drug Delivery Systems for Glaucoma Drop Medications

Name of Ref Clinician: Dr. _____ Date of Informed Consent: ____/ ___/ Investigator Code: _____

COUNTRY

Part 1: General patient information concerning glaucoma

1. Do you take glaucoma eye drops?

 \Box_1 Yes \Box_{\circ} No

2. Do you experience any side effects from your glaucoma eye drops?

 \Box_1 Yes \Box_0 No

- 3. If yes, which ones?
- 4. How strong are your side effects on a range from 0 (barely any side effects) to 10 (massive side effects)?

5. How long does it take you to reach our glaucoma clinic?

_____ hours

6. How do you get to our clinic?

 \Box_1 by car

 \square_2 by public transport

 \square_3 someone drives me

 \square_4 etc

 \square_5 Other, please specify: _____

Part 2: Demographics

- 1. Are you a subsidized or a private patient?
- \Box_1 Subsidized \Box_2 Private
 - 2. What is your age?

□ ₁ 20-30	□ ₂ 31-40
□ ₃ 41-50	□₄ 51-60
□ ₅ 61-70	□ ₆ 71-80
□ ₇ 81-90	□ ₈ >91

3. What is your ethnicity?

	White	(Europear	n, American))
-1		(Earopoar	, <i>i</i> anonoan <i>j</i>	

 \Box_3 Black

 \square_2 Asian \square_4 Hispanic

 $\square_5 I$ don't want to give an answer.

 \square_6 Other, please specify: _____

Country of origin: _____

4. Which religion do you belon

 \Box_1 none \Box_2 Christian

 \Box_3 Islam \Box_4 Judaism

- \Box_5 Buddhism \Box_6 Hinduism
- □₇ Other, please specify: _____
 - 5. What is your highest level of education?
- \Box_1 No formal education \Box_2 Primary
- \Box_3 Secondary \Box_4 A level

 \Box_5 Polytechnic / Diploma \Box_6 Graduate

 \Box_7 Post Graduate

- 6. What is your occupation? \square_1 Professional, Executive & Managerial \square_3 Production, Technical, Mechanical \square ⁵ Clerical, Administrative \Box_7 No occupation \square_8 Others, Please specify:
- 7. What is your monthly income (net income)? □₁less than 1,000 € \Box_2 1,000 \in to less than 2,000 \in \Box_3 2,000 \in to less than 5,000 \in □₄5,000 € to less than 10,000 € □₅10,000 € or more \square_6 I do not know. $\square_7 I$ don't want to give an answer. \square_8 No income..
 - 8. Do you live alone?
- □ _°No \Box_1 Yes
 - 9. Do you administer your glaucoma eye-drops for yourself?
- \Box_1 Yes \square_2 No. \square_3 Sometimes
 - 10. If no or sometimes, who helps you drop your glaucoma eye-drops? (Tick all that applicable)

(a) Husband/Wife/Partner	\Box ₁ Yes	🗆 。No
(a) Son/Daughter	\Box_1 Yes	🗆 。No
(b) Other relatives/friends/neighbours	\Box_1 Yes	$\Box_{\circ} No$
(c) Others, please specify:	\Box_1 Yes	🗆 。No

- \square_2 Self-employed
- □ ₄ Housewife

\square_{6} Retired

11. How many oral medications are you on?
0
1
2
3
4

 \Box More than 4

8. Are you taking any other medications?

 \Box_0 Sprays (number)

 \Box_1 Injections (number)

 \square_2 Others, please specify: _____

How I feel about my medicine

1. My health depends on my eye-drops.

□,I agree

□,I disagree

2. My life would be impossible without my eye-drops.

 $\Box_{A} I \text{ agree} \qquad \qquad \Box_{A} I \text{ disagree}$

3. My eye-drops protect me from becoming worse.

□,I agree

4. Having to take eye-drops worries me

□,I agree

□,I disagree

 \Box_{I} I disagree

5. I do not understand how my eye drops work.

□,I agree	\Box_{a} I disagree
6. My eye-drops disrupt my life	
□,I agree	□,I disagree
7. I sometimes worry about becoming too c	lependent on my eye-drops
□,I agree	□,I disagree
8. Most medicines are addictive	
□,I agree	□,I disagree
9. Medicine do more harm than good	
□,I agree	□.I disagree
10. Most medicines are addictive	
□,I agree	□,I disagree
11. Natural remedies are safer than medicin	ne
□,I agree	□,I disagree
12. Doctors place too much trust on medici	ne
□,I agree	□,I disagree
13. If doctors had more time with patients the	hey would prescribe fewer medicines
□,I agree	□,I disagree
14.Medicine help many people to live better	
□,I agree	□,I disagree
15. Medicine help many people to live long	jer
□,I agree	□,I disagree

16. In most cases the benefits of medicine outweigh the risks

□,I agree

 \Box_{a} I disagree

The Brief Illness Perception Questionnaire regarding your glaucoma disease

The Brief Illness Perception Questionnaire

For the following questions, please circle the number that best corresponds to your views:

10 severely affects my life
severely
and the my me
10 forever
10 extreme amount of control
10 extremely helpful
10 many severe symptoms
10 extremely concerned
10 understand very clearly
e you angry, scare
10 extremely affected

Patient Adherence Questionnaire

How frequently do you take your eyedrops?	□ ₀ 100%
	□ ₁ 90 - 99 %
	□ ₁ 51 - 89 %
	□ ₁ 0 - 50 %

Taking the prescribed glaucoma medication every day can be very demanding. It is very common to forget taking eye drops from time to time. The following questions are supposed to help to give us an insight on how often patients forget to take their prescribed medication. Please answer them truthfully. Your answers will remain anonymous and your doctor will not get access to your answers.

1. Do you sometimes forget to take your glaucoma eye drops?	□₁Yes □₀No
2. Over the past two weeks, were there any days when you did not take your glaucoma eye drops?	□₁Yes □₀No
3. Have you ever cut back or stopped taking your glaucoma eye drops without telling your doctor because you felt worse when you took it?	□₁Yes □₀No
4. When you travel or leave home, do you sometimes forget to bring along your glaucoma eye drops?	□₁Yes □₀No
5. Did you take your glaucoma eye drops yesterday?	□₁Yes □₀No
6. When you feel like your glaucoma is under control, do you sometimes stop taking your eye drops?	□₁Yes □₀No

7. Taking medication everyday is a real inconvenience for some people. Do you ever feel hassled about sticking to your glaucoma eye drops?	□₁Yes □₀No
8. How often do you have difficulty remembering to take all your glaucoma eye drops?	□ ₀ Never □ ₁ Very rarely □ ₂ A little of the time □ ₃ Sometimes □ ₄ All the time

Modified 8 item Medication Adherence Scale – Morisky et al., 2008

Part 6: Ocular Drug Delivery (ODD) Questionnaire

The treatment of glaucoma can be very challenging due to the many side effects, especially of eye drops, that may occur and reduce the patient's quality of life. A treatment option using invasive surgery can cause complications and therefore involves a certain risk.

Glaucoma treatment offers some promising minimally invasive approaches that could improve the treatment outcome and patients' quality of life. This involves finding alternative ways of applying the medications as an alternative to drops but without major glaucoma surgery being performed. Please imagine you could choose between the following options:

1) Contact lens

A contact lens similar to what people use to replace spectacles can be used to release glaucoma medication. A contact lens is placed onto the surface of the cornea (the clear window on the front of the eye) so it does not involve entering the eye in any way. Certain types of contact lenses can be inserted once, either by yourself or your eye specialist, and remain for weeks/months on the surface of the eye whilst releasing glaucoma medications slowly. The contact lens would have to be replaced at intervals (months) to replenish the medication supply - this can also be done by yourself, carer/partner or eye specialist. This option does not require a visit to the operating room and can theoretically be self-administered at home.



Source: https://www.aao.org/eye-health/diseases/contact_lenses (last accessed 23/04/22)

Have you ever heard of this treatment modality?

 \Box_1 Yes

 $\Box_{o} No$

How helpful do you feel this method of treatment would be for the treatment of your glaucoma?

1 2 3 4 5 6 7 8 9 10 Not at all very helpful

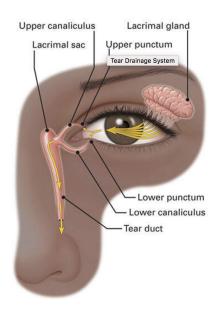
Would you consider undergoing this type of treatment?

 \Box_1 Yes

 $\Box_{\circ} No$

2) Punctum Plugs

Punctal plugs are tiny devices that are placed in the eye's tear ducts (called puncta). Puncta are the tiny openings that drain tears from your eyes. About the size of a grain of rice, the plug is usually used to stop fluid draining from the eye to treat dry eye conditions but the plug can also be used to slowly release glaucoma medications. Plugs trialled for glaucoma are made of a long-lasting medical plastic (such as silicone or acrylic) and are designed to stay in the eye for many months. They can be removed and replaced by your ophthalmologist if needed. Another type of semi-permanent punctal plug is placed in a deeper part of the tear duct called the canaliculus. These plugs cannot be seen at all in the eye. This option does not require a visit to the operating room but would need an eye specialist to insert/remove and/or ascertain position.



Source: https://www.aao.org/eye-health/diseases/punctal-plugs (last accessed 23/04/22)

Have you ever heard of this treatment modality?

 \Box_1 Yes

 $\Box_{\circ} No$

How helpful do you feel this method of treatment would be for the treatment of your glaucoma?

1 2 3 4 5 6 7 8 9 10 Not at all very helpful

Would you consider undergoing this type of treatment?

 \Box_1 Yes

 $\Box_{\circ} No$

3) Subconjunctival implants

A subconjunctival implant can either be placed under the eyelids or injected beneath the conjunctiva (the transparent cling film layer which covers the sclera (the white of the eye). These implants slowly release glaucoma medications and are covered either by the eyelid or conjunctiva itself, making them less exposed to the external environment when compared to a contact lens or punctal plug. This option does not require a visit to the operating room but would need an eye specialist to insert/remove and/or ascertain position. The injection would involve a very small needle but this is only inserted under the superficial surface of the eye (conjunctiva) and *not into the eye*.



Have you ever heard of this treatment modality?

 \Box_1 Yes

 $\square_{\circ} \mathsf{No}$

How helpful do you feel this method of treatment would be for the treatment of your glaucoma?

1 2 3 4 5 6 7 8 9 10 Not at all very helpful

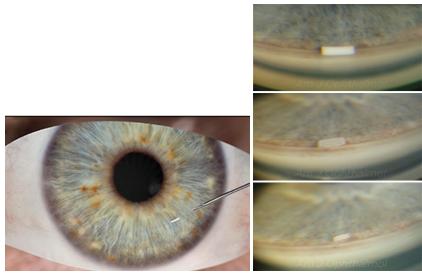
Would you consider undergoing this type of treatment?

 \Box_1 Yes

 $\square_{\circ} No$

4) Anterior Chamber (Intracameral) Administration

Slow release glaucoma medications can be administered into the anterior chamber of the eye (the space between the cornea at the front and iris at the back) via a very small biodegradable, preservative-free implant. Once placed, the drug is slowly released for about 3 to 4 months and the implant slowly dissolves in about 12 to 15 months. The added advantage of this route of administration is that side-effects related to the surface of the eye and skin are significantly reduced compared to drops and even other surface administrations such as mentioned above. On the other hand, the implant is placed into the eye itself, meaning it is slightly more invasive, and it needs to be inserted by an eye specialist in a clean clinic room environment or the operating theatre. Further implantations may/will be needed as the original one loses effect.



Source: AAO.org

Have you ever heard of this treatment modality?

 \Box_1 Yes

 \Box_{\circ} No

How helpful do you feel this method of treatment would be for the treatment of your glaucoma?

1 2 3 4 5 6 7 8 9 10 Not at all very helpful

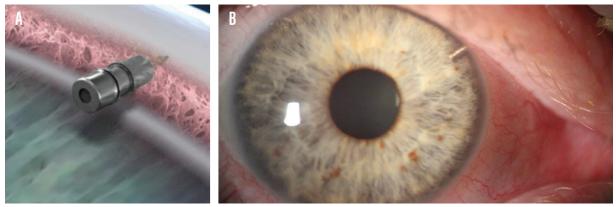
Would you consider undergoing this type of treatment?

 \Box_1 Yes

 \Box_{\circ} No

5) Trabecular Meshwork Microstents

Using a minimally invasive approach an implant/stent can be placed into the anterior chamber of the eye (the space between the cornea at the front and iris at the back) which reduces the intraocular pressure through a mechanical bypass process. These stents have been around for quite a few years now and have been shown to be very effective in reducing the number of glaucoma drops needed by patients. In many cases the stent insertion is combined with cataract surgery (which can also has a pressure lowering effect) but stents can also be used in eyes which have already had cataract surgery or even in eyes with the natural lens still in place (i.e. *not* combined with cataract surgery). Stents are now also being developed with the ability to slowly release glaucoma medications into the anterior chamber of the eye. Although still minimally invasive, stent insertion is more invasive than the above-mentioned options (as it is placed into the eye itself and not onto the surface or superficial layers). It also needs to be performed by an eye surgeon in the operating theatre and may need to be replaced or more stents added later as the effect of the original stent wears off.



Source:https://glaucomatoday.com/articles/2020-nov-dec/idose-an-intraocular-drug-eluting-delivery-device-for-glaucoma

Have you ever heard of this treatment modality?

 \Box_1 Yes

 $\Box_{\circ} No$

How helpful do you feel this method of treatment would be for the treatment of your glaucoma?

1 2 3 4 5 6 7 8 9 10 Not at all very helpful

Would you consider undergoing this type of treatment?

 \Box_1 Yes

 \Box_{\circ} No

If you had the choice to choose between these different therapy options, would you rather

- A: Keep your eye drops
- B: Use one or more of the aforementioned therapy options.

Would it make a difference to the answer before if the procedure had to be performed in the operating room?

A: Yes

B: No

Please consider that there are these alternative treatment methods for your glaucoma that could replace the current prescribed eye drops. How important are the following aspects of a treatment modality in your opinion?

A. HOW IMPORTANT IS EACH OF THE FOLLOWING TO YOU?

1 2 3 4 5 6 7 8 9 10 Not at all very important

Factors	Priority
Monetary cost to me	1 2 3 4 5 6 7 8 9 10 Not at all very important
Monetary cost to hospital / insurance	1 2 3 4 5 6 7 8 9 10 Not at all very important
Effectiveness	1 2 3 4 5 6 7 8 9 10 Not at all very important
Reduction in side effects	1 2 3 4 5 6 7 8 9 10 Not at all very important
Less frequent follow-ups	1 2 3 4 5 6 7 8 9 10 Not at all very important
Reversibility of Treatment *	1 2 3 4 5 6 7 8 9 10 Not at all very important
Biodegradability **	1 2 3 4 5 6 7 8 9 10 Not at all very important
Ability to apply it on your own	1 2 3 4 5 6 7 8 9 10 Not at all very important
Long duration	12345678910 Not at all very important

<u>* Possibility to take it out without additional risk</u>
 <u>** It will disappear from your system eventually</u>

Which of these are the 2 most important points for you? Please tick the two most important ones.

- □ Cost
- Effectiveness
- □ Reduction in side effects
- □ Less frequent follow-ups
- □ Reversibilty of Treatment
- Biodegradability
- □ Ability to apply it on your own
- □ Long duration

What impact would the cost of the treatment TO YOU have on your decision if the treatment was more expensive than your actual treatment?

A: I don't care how much it costs.

B: I would not choose this option, but rather go for another treatment.

C: It depends on the difference in cost.

If you ticked C: How much would you be willing to pay:

Have a second to think and ask our interviewer about any doubts

IN THE CASE ALL OF THESE are equally effective, which one would you choose? (only tick 1)

- \Box_1 Contact Lens
- \square_2 Punctal Plugs
- \Box_3 Subconjunctival
- \Box_4 Intracameral
- \Box_5 Stent

IN THE CASE WHERE EFFICACY AND INVASIVENESS BOTH INCREASED FROM CONTACT LENS THROUGH TO STENTS (I.E. CL LEAST INVASIVE/EFFECTIVE, STENT MOST INVASIVE/EFFECTIVE, which one would you choose? (only tick 1)

- \Box_1 Contact Lens
- \square_2 Punctal Plugs
- \Box_3 Subconjunctival
- \Box_4 Intracameral
- \Box_5 Stent

IN THE CASE WHERE DURATION AND INVASIVENESS BOTH INCREASED FROM CONTACT LENS THROUGH TO STENTS (I.E. CL LEAST INVASIVE/SHORTEST DURATION/MORE FOLLOW-UPS, STENT MOST INVASIVE/LONGEST DURATION/LESS FOLLOW-UPSwhich one would you choose? (only tick 1)

- \Box_1 Contact Lens
- \square_2 Punctal Plugs
- \Box_3 Subconjunctival
- \Box_4 Intracameral
- \Box_5 Stent