# Efficacy test of micro-dosing with MistGo®

Conducted by Copenhagen University Hospital (Rigshospitalet) 2023

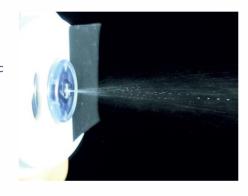


### Background<sup>1</sup>

- Medicating with an eye dropper result in a drop containing 30-50  $\mu$ l which greatly exceeds the physiologic 6-8  $\mu$ L ocular tear film capacity.
- A part of the delivered solution is washed out of the eye, but up to 80% of the dose enters the systemic circulation via transport into surface blood vessels or passage through the nasal lacrimal duct and into the digestive system, leading to an increased risk of systemic adverse events.
- Any excess solution remaining in the eye can lead to local adverse events, incl. hypersensitivity reactions to the drug or excipients.
- From a biopharmaceutical and economic point of view, the ideal would be to instill smaller volumes of eye medication as they can achieve a maximal tear film concentration with far less systemic absorption.
- Studies show that micro-dosing is as safe and efficacious as normal eye-drops.

# MistGo® designed to micro-dose

- MistGo<sup>®</sup> contains a high-precision micropump, consistently metering a dose of 6 μL, dispensing no excess liquid to irritate the skin, enter systemic circulation or risk adverse events.
- The internal geometry of the nozzle is designed to vaporise the drug into a fine mist with a broad impact pattern, coating the entire cornea for optimal absorption (~10-12 mm diameter).



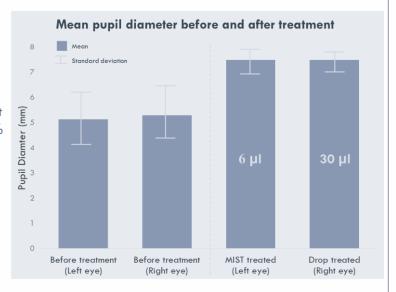
Mist of glaucoma medication latanoprost

## Purpose & Methods

- **Purpose**: to verify whether a micro dosing of 6 μL eye medicine delivered as a mist is comparable to a conventional eyedrop of appr. 30 μL of the same eye medication. Primary endpoint: pupil dilation degree.
- Method: By use of a pupilometer the size of the subject's eye pupils were measured. A 6 µL mist dose of pupil dilation medicine Tropicamide 1% was given to each test subject's left eye by MistGo® test set-up. A drop of the same medicine was given to each test subject's right eye by a conventional eye drop bottle. Min 15 minutes after mist and drop instillation, the size of the subject's eye pupils was measured by use of pupilometer.

#### Results and Conclusions

- Successful dosing using the MistGo® test setup was achieved in 100% of subjects in first attempt.
- Tropicamide treatment gave a significant pupil dilation for both eyes. There is no statistically significant difference in the obtained pupil dilations comparing treatment with a micro dose of 6 μL of Tropicamide 1% delivered as a mist to treatment with a conventional eyedrop of approximately 30 μL Tropicamide 1%.
- It can be concluded that MistGo®
  micro-dosing mist technology provides
  a comparable pupil dilation using
  Tropicamide 1% as when using a
  conventional eyedrop applicator.



<sup>1:</sup> Washington N, Washington C, Wilson CG, "Ocular drug delivery". Physiological Pharmacceutics: Barriers to Drug Absorption, 2nd edition, CRC Press, 2001, pp 249–270.; Mishima S et al, "Determination of tear volume and tear flow". Invest Ophthalmol, 1966, Vol 5(3), pp 264–276.; Scherz W, Doane MG, Dohlman CH, "Tear volume in normal eyes and keratoconjunctivitis sicca". Allbreaft Von Graefes Arch Klin Exp Ophthalmol, 1974, Vol 192(2), pp 141–150.; Guiroz-Mercado H, Ivri E, Gonzalez-Salinas R, Kourtis IC, Gilbert J, Pérez-Vaçuez JF, Blumenkranz M, Jiménez-Román J, Marcellino G. Clinical Evolucition of a Novel Electromechanical Topical Ocular Drug Delivery System: Two Phase 1 Proof of Concept Studies. Clin Ophthalmol. 2020 Jan 20;14 pp 139-147; Van Santvillet L, Ludwig A, "Determinants of eye drop size". Surv Ophthalmol, 2004, Vol 49(2), pp 197-213.; Brown RH, Hotchkiss ML, Davis EB, "Creating Smaller Eyedrops by Reducing Eyedrope Tip Dimension, 1985, Vol 99(4), pp 460-464.; Collins JF, Dartt DA, Dana R. Mist delivery of eye medication to the anterior to the anterior to the anterior of the control of t