

# Rehabilitation of Unresolved Corneal Problems

Descemetorhexis and ROCK inhibitors

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## **Historical Background of DSO**

- Descemetorhexis without endothelial keratoplasty (DSO) emerged in the early 2010s as a response to the lack of tissue donor and the risks associated with donor grafts and chronic immunosuppression.
- Macsai et al. (2015) published the first clinical series demonstrating successful corneal clearance without grafting in Fuchs dystrophy.

• Source: Macsai MS et al. Cornea. 2015;34(8):873-880.

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## **Rationale for Development**

- The recognition that peripheral corneal endothelium retains the ability to migrate and repopulate damaged areas led to the concept of removing only the diseased central zone without replacing it.
- Source: Borkar DS et al. Ophthalmology. 2016;123(2):247–252.
- Peripheral endothelial cells exhibit limited mitotic activity and enhanced migratory capacity. R
- ROCK inhibitors such as ripasudil potentiate this regenerative response by modulating cytoskeletal tension and promoting adhesion and proliferation.

• Source: Okumura N et al. Invest Ophthalmol Vis Sci. 2012;53(10):6066–6072.

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## **Minimum Endothelial Cell Density for DSO**

 A minimum peripheral ECD of 1000–1200 cells/mm<sup>2</sup> is recommended to ensure sufficient regenerative capacity post-DSO. Specular microscopy and endothelial maps help assess cell distribution preoperatively.

• Source: Wacker K et al. Am J Ophthalmol. 2019;205:109–122.

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## When to Choose DSO vs. DMEK

DSO is indicated in Fuchs dystrophy with central guttae, clear periphery, and ECD >1000 cells/mm<sup>2</sup>.
DMEK is preferred in diffuse guttae, low ECD, or failed prior surgery. DSO offers no rejection risk, lower cost, and avoids long-term steroid use.

• Source: Wacker K et al. Am J Ophthalmol. 2019;205:109–122. Macsai MS. Eye Contact Lens. 2020;46(2):S46–S49.

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## **Surgical Technique of DSO**

 Under topical or peribulbar anesthesia, a 4–6 mm central descemetorhexis is created using blunt dissection with a Sinskey hook or reverse stripper. No graft is implanted. Viscoelastic may be used. The anterior chamber is reformed and no endothelial tissue is replaced.

Source: Borkar DS et al. Ophthalmology. 2016;123(2):247–252.



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## **Descemetorhexis Size and Centering**

 Standard diameters range from 4 to 6 mm. Larger diameters (>6 mm) are associated with delayed recovery and greater risk of failure. Centering should align with the visual axis and zone of confluent guttae, typically guided by retroillumination and AS-OCT.

• Source: Macsai MS, Shiloach M. Cornea. 2015;34(8):873-880.

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## **Mechanism of Action of Ripasudil**

 Ripasudil inhibits Rho-kinase, relaxing actin-myosin tension in endothelial cells. This enhances cellular adhesion, migration and mitosis. It also modulates the extracellular matrix and suppresses inflammatory pathways.

• Source: Koizumi N et al. Sci Rep. 2016;6:28102.

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## **Ripasudil Posology and Duration**

- Ripasudil 0.4% is typically prescribed as 1 drop 4 to 6 times daily for 3 months postoperatively.
- Treatment duration may be extended based on repopulation rate assessed via specular microscopy or AS-OCT.
- It is safe for long-term use.
- Ripasudil should be tapered or discontinued if signs of central edema persist beyond 3 months or if no improvement in visual acuity is observed. Consider DMEK rescue if corneal thickness remains >650 μm or BCVA <20/40 at 12 weeks.</li>

• Source: Borkar DS et al. Ophthalmology. 2016;123(2):247–252.

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## **Visual Outcomes After DSO with Ripasudil**

 BCVA improvement to ≥20/25 achieved in 80–85% of successful DSO cases by 3 months. Time to visual recovery shorter with ripasudil (median 6–8 weeks). CCT normalizes progressively.

• Source: Wacker K et al. Am J Ophthalmol. 2019;205:109–122.

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## **Complications Associated with Ripasudil**

- Most common adverse events are transient hyperemia and mild irritation.
- Discontinue if epithelial defects persist beyond 2 weeks or if there is anterior chamber reaction.
- No significant intraocular toxicity reported.
- Source: Okumura N et al. Cornea. 2018;37(6):730–735.

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## **Evidence: DSO With vs. Without Ripasudil**

- Studies show DSO with ripasudil significantly accelerates endothelial repopulation (mean 6–8 weeks vs. 12–16 weeks without).
- Final visual outcomes are similar, but ripasudil reduces time to clearance and improves ECD in central cornea.

• Source: Macsai MS. Eye Contact Lens. 2020;46(2):S46–S49.

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## **Long-Term Durability of DSO**

- Studies with >24 months of follow-up report stable corneal clarity and endothelial cell density after successful DSO. No progressive endothelial failure observed in compliant patients with proper selection criteria.
- DMEK grafts show long-term survival in 90–95% at 5 years, whereas DSO success depends on initial ECD and response to therapy. With adjunctive ripasudil, 2-year data suggest comparable durability in mild to moderate cases.

• Source: Macsai MS. Eye Contact Lens. 2020;46(2):S46–S49.7–252.

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## **Economic Impact: DSO vs. Transplant**

• DSO eliminates the need for donor tissue, storage, transportation, surgical prep, and immunosuppressive therapy. One study estimated a 40–60% cost reduction per case vs. DMEK/DSAEK when successful.

• Source: Saad A et al. Cornea. 2021;40(2):163–169.

• Although ripasudil represents an added pharmaceutical cost (~€70–90/month), the overall procedure remains more cost-effective by avoiding graft and OR time. Insurance coverage varies by region.

• Source: Koizumi N et al. Invest Ophthalmol Vis Sci. 2020;61(4):29.

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## #1 70y/o Preop BCVA 0,45



### #1 70y/o Preop BCVA 0,45



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**1** day after the surgery





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## **#1 ripasudil stopped BCVA 0,85**



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# #2 70y/o Preop BCVA 0,45



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## #2 Ripasudil stopped BCVA 0,65

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# #3 62y/o Preop BCVA 0,05



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## #3 Ripasudil stopped BCVA 0,5 (AMD)

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#4 65 y/o BCVA 0,45



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## 1 month

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## 3 month

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## **#4 Ripasudil stopped BCVA 0,75**

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# **#5** Trauma on a 56 y/o BCVA 0,2

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# #5 Trauma (n a 56 y/o BCVA 0,3

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## **Conclusions DSO**

Ideal indications:

- Fuchs endothelial dystrophy with central guttae, clear peripheral cornea, and peripheral ECD ≥ 1000– 1200 cells/mm<sup>2</sup>.
- 2. Good visual potential and absence of stromal edema.

#### Descemetorhexis diameter:

3. A size of **5–6 mm** is optimal. Larger diameters (>6 mm) increase the risk of delayed or incomplete healing. **Surgical centration**:

4. Should align with the visual axis and fully encompass the area of confluent guttae

#### Absolute contraindications:

5. Advanced stromal edema, ECD <1000 cells/mm<sup>2</sup>, glaucoma, prior keratoplasty, or diffuse guttae.

#### Durability:

6. Clinical studies up to 24 months show **stable outcomes** with no progressive decompensation in wellselected patients.

Be very gentle..

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## **Conclusions DSO+Ripasudil**

#### Proven benefit:

•Reduces mean corneal clearance time from **12–16 weeks to 6–8 weeks**.

#### Adverse effects:

•Mostly mild hyperemia and transient discomfort. No significant intraocular toxicity reported. When to suspend treatment:

•If no clinical improvement in corneal clarity or visual acuity after **3 months**.

•Persistent corneal thickness >650  $\mu$ m or BCVA <20/40 at 12 weeks suggests failure.

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## Thank you

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