



Updates on Neuro-ophthalmology: Optic nerve

14-SEP-2024, 15:15-15:30, The Islands Meeting
College Of Ophthalmologists of Sri Lanka
axel petzold



Disclosures

NIHR UK, UCSF
Stichting MS Research NL
Novartis, Heidelberg Academy



Content

Optic Nerve Updates:

- Anatomical
- Imaging: AI for OCT & MRI
- Treatment



Optic Nerve Glymphatic System in rodents

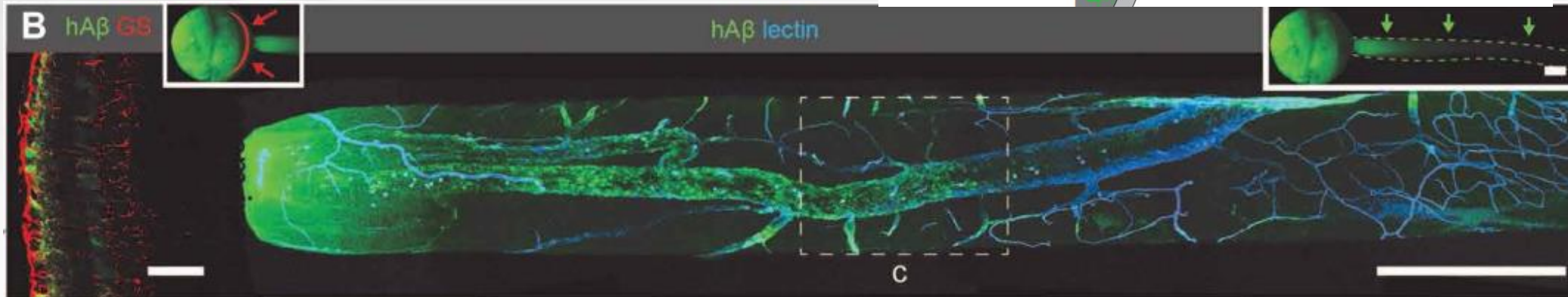
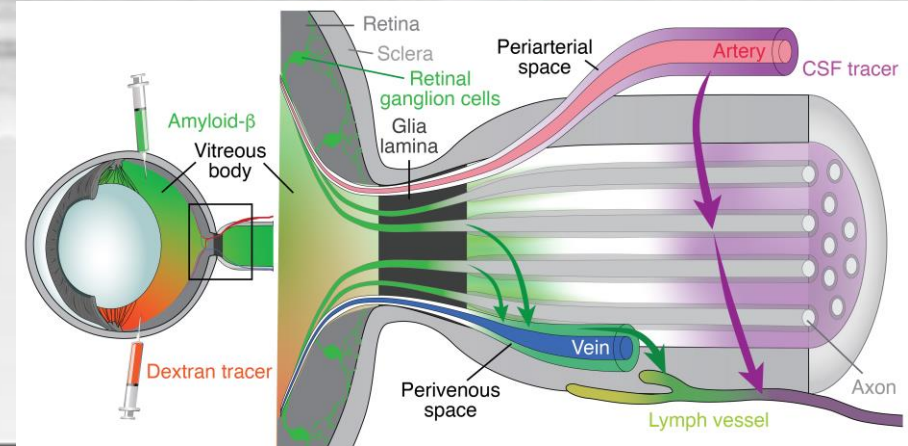
SCIENCE TRANSLATIONAL MEDICINE | RESEARCH ARTICLE

EYE PHYSIOLOGY

An ocular glymphatic clearance system removes β -amyloid from the rodent eye

Xiaowei Wang^{1,2}, Nanhong Lou², Allison Eberhardt², Yujia Yang³, Peter Kusk¹, Qiwu Xu², Benjamin Förster^{4,5}, Sisi Peng², Meng Shi³, Antonio Ladrón-de-Guevara², Christine Delle¹, Björn Sigurdsson¹, Anna L. R. Xavier¹, Ali Ertürk^{4,5}, Richard T. Libby⁶, Lu Chen^{3*}, Alexander S. Thrane^{1,7}, Maiken Nedergaard^{1,2*}

Despite high metabolic activity, the retina and optic nerve head lack traditional lymphatic drainage. We here identified an ocular glymphatic clearance route for fluid and wastes via the proximal optic nerve in rodents. β -amyloid

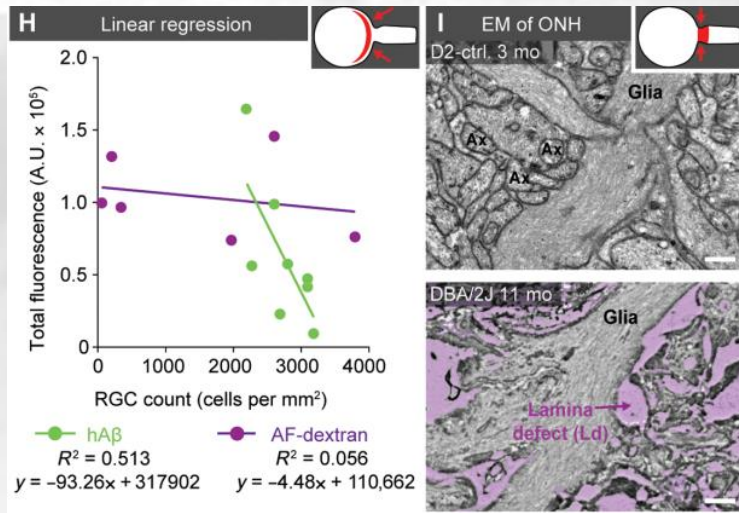


Wang, X. et al. Sci Translat Med 2020

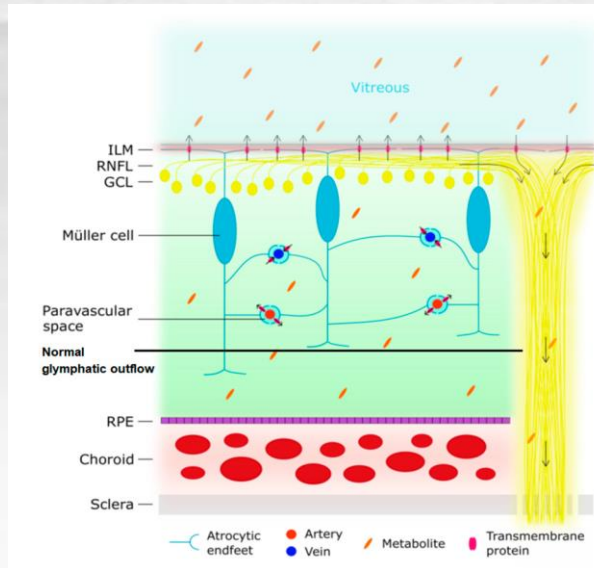
Delle, C et al. Mol Sci 2024



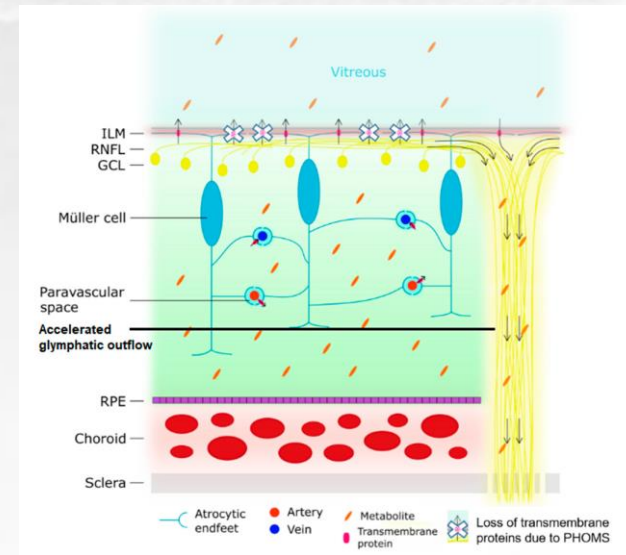
Optic Nerve Glymphatic System in humans



CTRL



Optic atrophy



Wang, X. et al. Sci Translat Med 2020

Kacar, S. et al. Brain Sciences 2023;14:36



Pioneering Sri Lankan Astronaut

NASA includes Sri Lankan in crew for simulated Mars journey

April 17, 2024



NASA has selected a new crew of four volunteers, including a Sri Lankan, to participate in a simulated mission to Mars within a habitat at the agency's Johnson Space Center in Houston.

Jason Lee, Stephanie Navarro, Shareef Al Romaithi, and Piyumi

Wijesekara will step into the agency's Human Exploration Research Analog, or HERA, on Friday, May 10.

Life Sciences in Space Research 42 (2024) 40–46



Contents lists available at [ScienceDirect](https://www.sciencedirect.com)

Life Sciences in Space Research

journal homepage: www.elsevier.com/locate/lssr



Review article

Imaging in spaceflight associated neuro-ocular syndrome (SANS): Current technology and future directions in modalities

Benjamin Soares^{a,*}, Joshua Ong^b, Ethan Waisberg^c, Prithul Sarker^d, Nasif Zaman^d, Alireza Tavakkoli^d, Andrew G. Lee^{d,e,f,g,h,i,j,k,l,*}



Literature update: A new Journal
The optic nerve is in the 1st edition



Content

- Optic Nerve News:
 - Anatomical update
 - Imaging update: OCT & AI
 - Treatment update



OCT Diagnostic Criteria Update

Panel 1: Diagnostic criteria for optic neuritis

Clinical criteria

- A: Monocular, subacute loss of vision associated with orbital pain worsening on eye movements, reduced contrast and colour vision, and relative afferent pupillary deficit
- B: Painless with all other features of (A).
- C: Binocular loss of vision with all features of (A) or (B).

Paraclinical criteria

- OCT: Corresponding optic disc swelling acutely or an inter-eye difference in the mGCIPL of >4% or >4 μm or in the pRNFL of >5% or >5 μm within 3 months after onset.**
- MRI: Contrast enhancement of the symptomatic optic nerve and sheaths acutely or an intrinsic signal (looking brighter) increase within 3 months.
- Biomarker: AQP4, MOG, or CRMP5 antibody seropositive, or intrathecal CSF IgG (oligoclonal bands).

Application of the clinical and paraclinical criteria

Definite optic neuritis

- (A) and one paraclinical test
- (B) and two paraclinical tests of different modality
- (C) and two different paraclinical tests of which one is MRI

Possible optic neuritis

- (A), (B), or (C) if seen acutely but in absence of paraclinical tests, with fundus examination typical for optic neuritis and consistent with the natural history during follow-up
- Positive paraclinical test or tests, with a medical history suggestive of optic neuritis

ICON study group TLN 2022

Multicenter Study > Neurol Neuroimmunol Neuroinflamm. 2024 Nov;11(6):e200291.

doi: 10.1212/NXI.0000000000200291. Epub 2024 Sep 4.

Diagnostic Value of Inter-Eye Difference Metrics on OCT for Myelin Oligodendrocyte Glycoprotein Antibody-Associated Optic Neuritis

Giulio Volpe¹, Neringa Jurkute¹, Gabriela Girafa¹, Hanna G Zimmermann¹, Seyedamirhosein Motamedi¹, Charlotte Bereuter¹, Lekha Pandit¹, Anitha D'Cunha¹, Michael R Yeaman¹, Terry J Smith¹, Lawrence J Cook¹, Alexander U Brandt¹, Friedemann Paul¹, Axel Petzold¹, Frederike C Oertel¹

> J Neurol Neurosurg Psychiatry. 2023 Jul;94(7):560-566. doi: 10.1136/jnnp-2022-330608. Epub 2023 Feb 21.

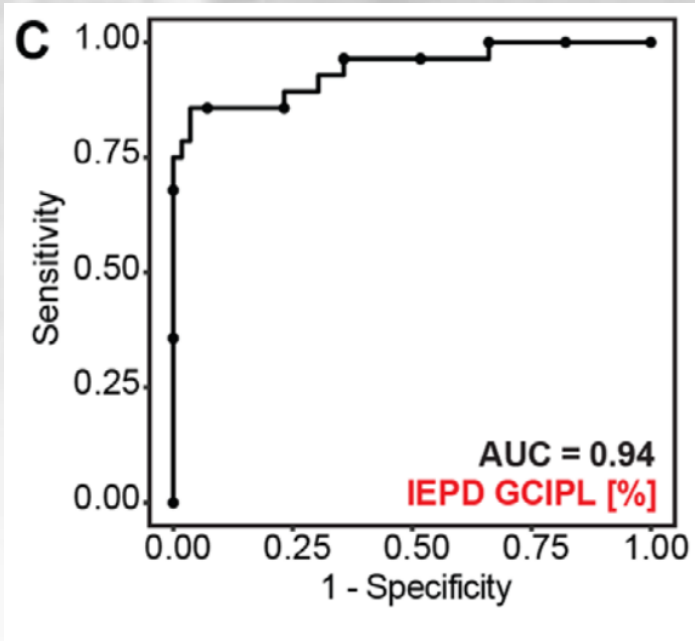
Diagnostic value of intereye difference metrics for optic neuritis in aquaporin-4 antibody seropositive neuromyelitis optica spectrum disorders

Frederike Cosima Oertel^{1 2 3}, Hanna G Zimmermann^{1 2 4}, Seyedamirhosein Motamedi^{1 2}, Claudia Chien^{1 2}, Orhan Aktas⁵, Philipp Albrecht⁵, Marius Ringelstein^{5 6}, Anitha Dcunha⁷, Lekha Pandit⁷, Elena H Martinez-Lapiscina⁸, Bernardo Sanchez-Dalmau⁸, Pablo Villoslada^{8 9}, Jacqueline Palace¹⁰, Adriana Roca-Fernández¹⁰, Maria Isabel Leite¹⁰, Srilakshmi M Sharma¹¹, Letizia Leocani¹², Marco Pisa¹², Marta Radaelli¹², Marco Aurélio Lana-Peixoto¹³, Mariana Andrade Fontenelle¹³, Joachim Havla¹⁴, Fereshteh Ashtari¹⁵, Rahele Kafieh¹⁶, Alireza Dehghani¹⁷, Mohsen Pourazizi¹⁷, Romain Marignier¹⁸, Alvaro Cobo-Calvo^{18 19}, Nasrin Asgari^{20 21}, Anu Jacob^{22 23}, Saif Huda²², Yang Mao-Draayer²⁴, Ari J Green³, Rachel Kenney²⁵, Michael R Yeaman^{26 27}, Terry J Smith^{28 29}, Lawrence Cook³⁰, Alexander U Brandt^{1 2 31}, Friedemann Paul^{1 2 32 33}, Axel Petzold^{1 34 35}



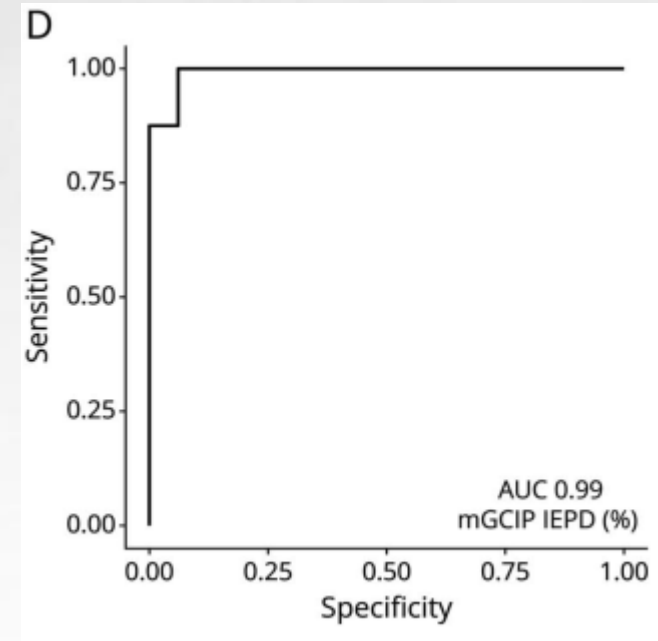
Excellent diagnostic accuracy of mGCIPL >4%

NMO-ON



Oertel, F. et al. JNNP 2023

MOG-ON



Volpe, G. et al. Neurol Neuroimmunol Neuroinflamm 2024



AI Update

Article

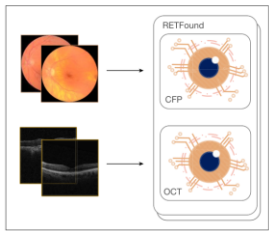
A foundation model for generalizable disease detection from retinal images

<https://doi.org/10.1038/s41586-023-06555-x>

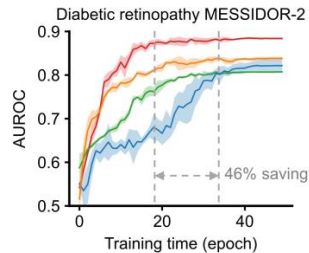
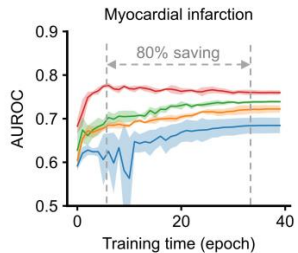
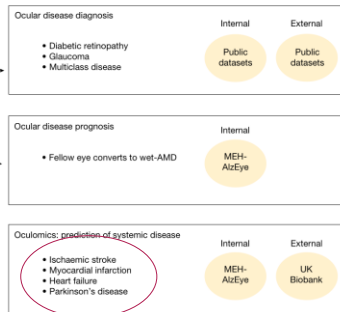
Received: 5 December 2022

Yukun Zhou^{1,2,3,4}, Mark A. Chia^{1,4}, Siegfried K. Wagner^{1,4}, Murat S. Ayhan^{1,4},
Dominic J. Williams^{1,4}, Robbert R. Struyven^{1,4}, Taming Liu^{1,4}, Mou Cheng Xu^{1,4},
Mateo G. Lozano^{1,4}, Peter Woodward-Court^{1,4}, Yuka Kihara^{1,4}, UK Biobank Eye & Vision

Stage 1: Self-supervision on retinal images



Stage 2: Supervised fine-tuning for clinical tasks



— RETFound — SSL-Retinal — SSL-ImageNet — SL-ImageNet

nature medicine



Article

<https://doi.org/10.1038/s41591-024-03139-8>

Integrated image-based deep learning and language models for primary diabetes care

Received: 26 November 2023

Accepted: 18 June 2024

Published online: 19 July 2024

Check for updates

A list of authors and their affiliations appears at the end of the paper

Primary diabetes care and diabetic retinopathy (DR) screening persist as major public health challenges due to a shortage of trained primary care physicians (PCPs), particularly in low-resource settings. Here, to bridge the gaps, we developed an integrated image–language system (DeepDR-LLM), combining a large language model (LLM module) and image-based deep learning (DeepDR-Transformer), to provide individualized diabetes management recommendations to PCPs. In a retrospective evaluation, the LLM module demonstrated comparable performance to PCPs and endocrinology residents when tested in English and outperformed PCPs and



10.45 – 11.45 A.M.

DR. P. SIVASUBRAMANIAM ORATION

**‘EMERGING GLUCOSE METRICS & MEASURES OVER TWO DECADES OF RESEARCH IN
DIABETES AND ITS IMPACT ON METABOLIC COMPLICATIONS’**

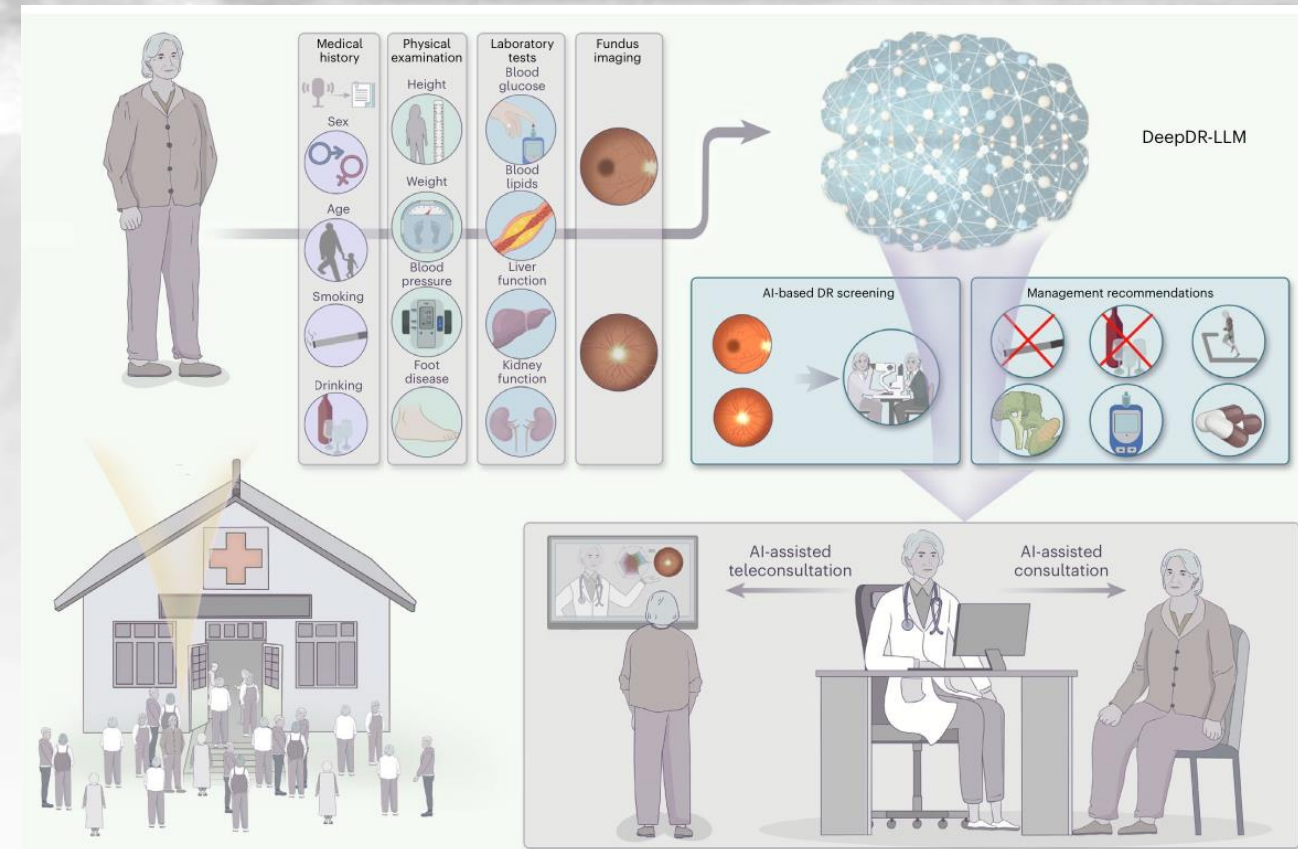
Dr. Uditha Bulugahapitiya

*MBBS (SL), MD (Col), MRCP (UK), FRCP (Edin) FACE (USA),
FCCP (SL), FSLCE (SL)*

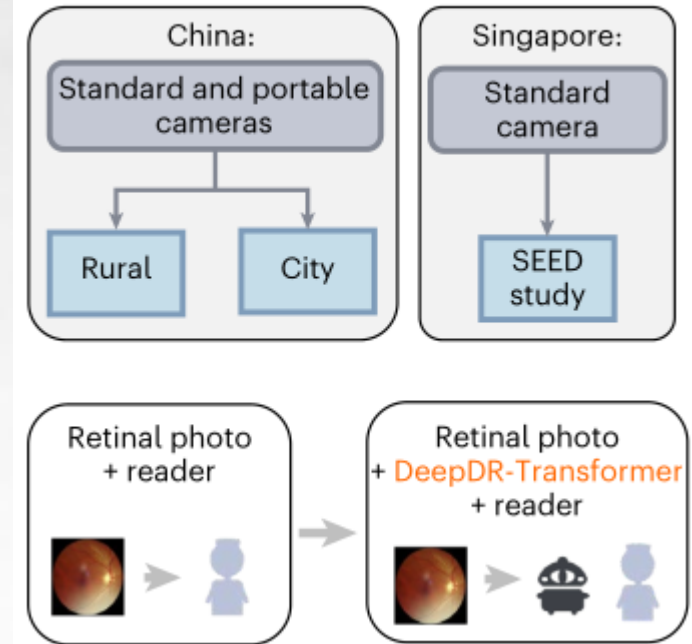
*Consultant Endocrinologist
National Hospital of Sri Lanka*



The future of patient care ?



C As assistive tool for referable DR detection



Li, J. *et al* Nat Med 2024

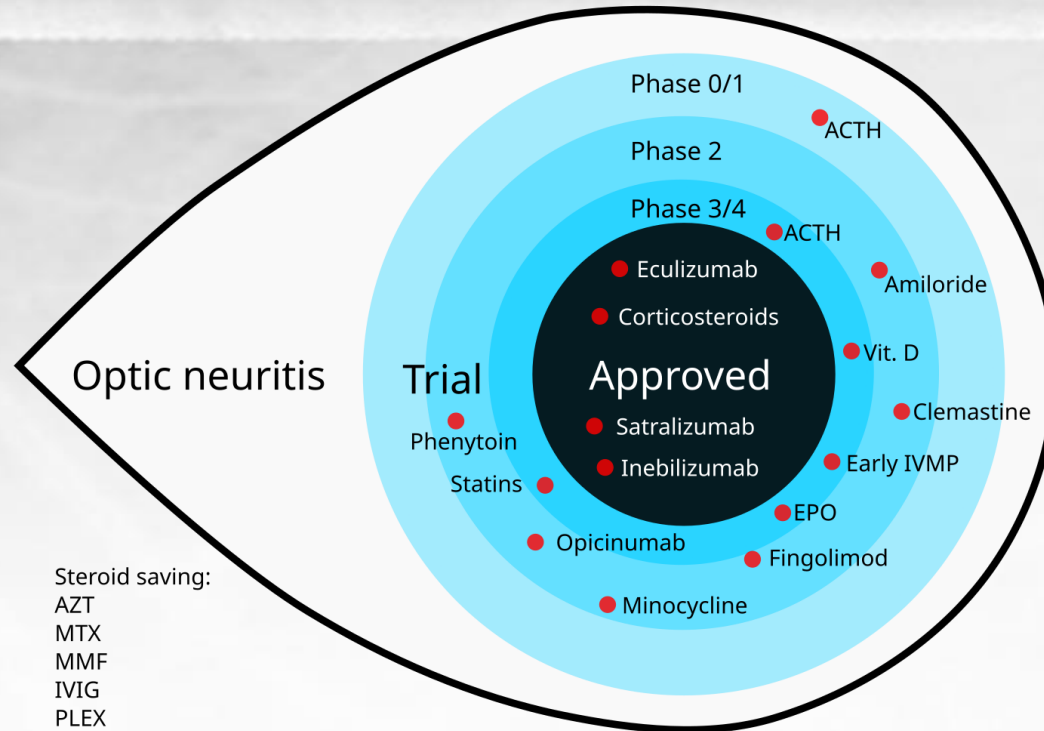


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Update on Treatment





Optic Nerve Repair = Remyelination

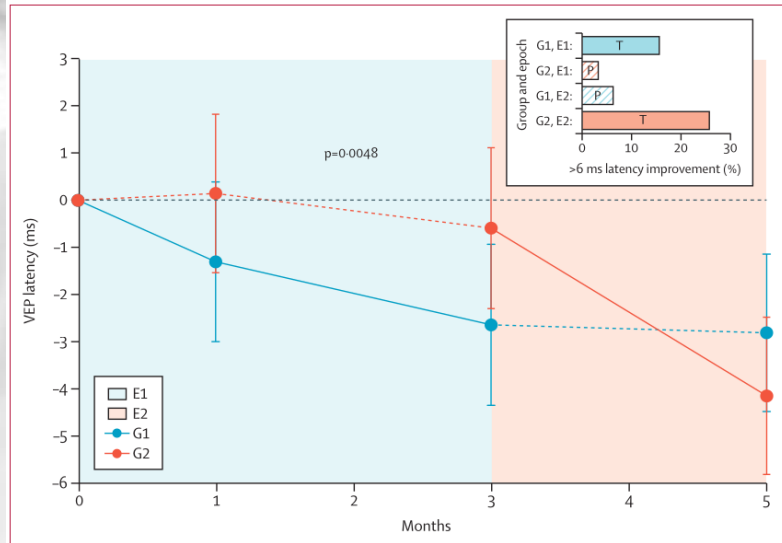
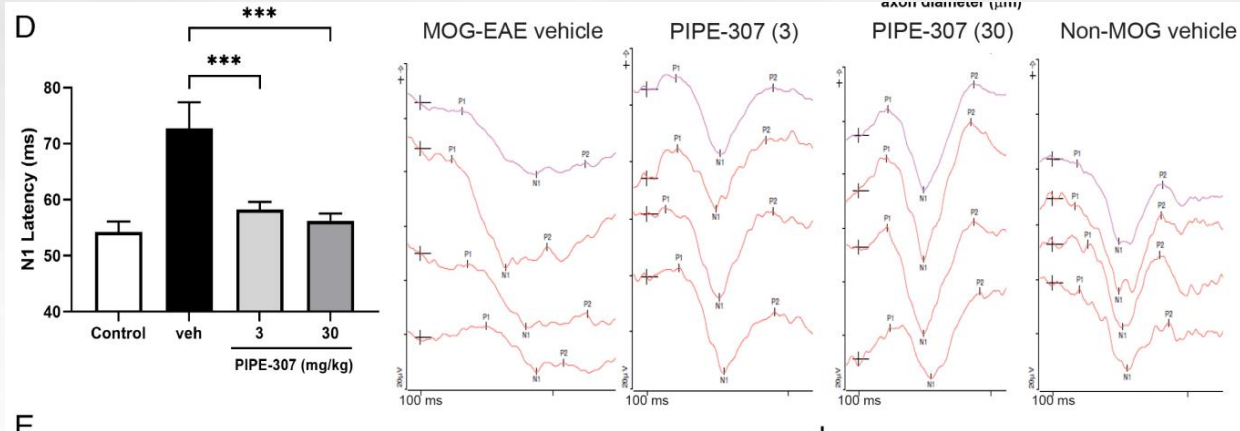


Figure 2: Association of clemastine fumarate treatment with VEP latency delay in patients with chronic optic neuropathy
Change from baseline in latency by group and epoch (model-derived estimates of means are represented by dots with the SE from baseline represented by error bars at each relevant timepoint). Solid line is on-treatment and dashed line is on-placebo. Blue line is group 1, orange line is group 2. Blue shaded area is epoch 1 and orange shaded area is epoch 2. p value is for primary analysis including crossover (with assumption of carryover). The inset is the percentage of patients with more than 6 ms improvement in latency delay. VEP=visual-evoked potential. G1=group 1. G2=group 2. E1=first epoch. E2=second epoch. T=treatment period. P=placebo period.

Is a VEP improvement of ~2 ms relevant? – Probably not.

Is a VEP improvement of ~20 ms relevant? – Probably yes.



Green, AJ *et al* Lancet 2017

Poon, MM *et al* PNAS 2024



Conclusion – Optic Nerve Updates

- Anatomically: novel fluid transport systems (glymphatic) to be taken into account for interpretation of optic atrophy and OCT signal changes
- Imaging: excellent diagnostic performance of novel diagnostic criteria (ROC AUC > 0.99) in MOG-ON
- Treatment: at the dawn of new remyelination agents with a hope to heal



Thank you - Q&A

Please support the ICON survey
Co-authorship as part of the ICON study group name

02-SEP-2024 update

