## What's next in Al for glaucoma screening? The REFUGE challenge outcomes

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## **Opportunities for AI in glaucoma**

Screening

Follow up / prognosis Treatment planning

**Disease understanding** 

## **Opportunities for AI in glaucoma**

### Screening

Follow up / prognosis Treatment planning

**Disease understanding** 

### **Types of screening**

Population based (universal) Case detection (opportunistic)

#### **COST EFFECTIVENESS!**



## How to reduce the cost of glaucoma screening using AI?

# The Diabetic Retinopathy experience



## **Grand challenges**





#### **Diabetic Retinopathy Detection**

Identify signs of diabetic retinopathy in eye images \$100,000 · 661 teams · 4 years ago



Overview	Data	Notebooks	Discussion	Leaderboard	Rules	Join Competition

#### **Overview**

#### Description

Evaluation

Prizes

Timeline

**Diabetic retinopathy** is the leading cause of blindness in the working-age population of the developed world. It is estimated to affect over 93 million people.



The US Center for Disease Control and Prevention estimates that 29.1 million people in the US have diabetes and the World Health Organization estimates that 347 million people have the disease worldwide. Diabetic Retinopathy (DR) is an eye disease associated with long-standing diabetes. Around 40% to 45% of Americans with diabetes have some stage of the disease. Progression to vision impairment can be slowed or averted if DR is detected in time, however this can be difficult as the disease often shows few symptoms until it is too late to provide effective treatment. **Original Investigation** | Innovations in Health Care Delivery

December 13, 2016

## Development and Validation of a Deep Learning Algorithm for Detection of Diabetic Retinopathy in Retinal Fundus Photographs

Varun Gulshan, PhD<sup>1</sup>; Lily Peng, MD, PhD<sup>1</sup>; Marc Coram, PhD<sup>1</sup>; <u>et al</u>

» Author Affiliations | Article Information

JAMA. 2016;316(22):2402-2410. doi:10.1001/jama.2016.17216

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B Messidor-2: AUC, 99.0%; 95% CI, 98.6%-99.5%



+ Home / News & Events / FDA Newsroom / Press Announcements / FDA permits marketing of artificial intelligence-based device to detect certain diabetes-related eve problems

**FDA NEWS RELEASE** 

#### FDA permits marketing of artificial intelligencebased device to detect certain diabetes-related eye problems

	Image: Formation of the second se	
G More Press Announcements	For Immediate Release: April 11, 2018	Content current as of:
Press Announcements	Español	04/12/2018
	The U.S. Food and Drug Administration today permitted marketing of the first medical device to use artificial intelligence to detect greater than a mild level of the eye disease diabetic retinopathy in adults who have diabetes.	Follow FDA Follow @US_FDA C Follow FDA C Follow @EDAmedia (
	Diabetic retinopathy occurs when high levels of blood sugar lead to damage in the blood vessels of the retina, the light-sensitive tissue in the back of the eye. Diabetic retinopathy is the most common cause of vision loss among the more than 30 million Americans living with diabetes and the leading cause of vision impairment and blindness among working- age adults.	
	"Early detection of retinopathy is an important part of managing care for the millions of people with diabetes, yet many patients with diabetes are not adequately screened for	

diabetic retinopathy since about 50 percent of them do not see their eye doctor on a yearly having " and Makring Eudolmon M.D. director of the Division of Onbthalmic and Far





#### Autonomous AI that instantly detects disease

Increase patient access to early disease detection.

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#### https://www.eyediagnosis.co

### But...

#### Glaucoma features less evident in fundus images Glaucoma as a multimodal disease

## High cost of building (huge) reliable data sets

Clinical records based labels + Scrapping on clinical databases





#### **Retinal Fundus Glaucoma Challenge**

20<sup>th</sup> September, Granada, Spain OMIA Workshop, MICCAI 2018



## The largest available data set of fundus pictures for glaucoma assessment

1200 images (10% glaucoma) divided in training, validation and test sets

#### Reliable annotations for glaucoma classification and optic disc/cup segmentation

Glaucoma labels retrieved from clinical records and optic disc/cup segmentations produced by +7 experts

#### A common evaluation framework to compare fundus image analysis methods for glaucoma

A test set with private labels + evaluation routines to report performance and prevent overfitting on test data

Zeiss Visucam 500	Cannon CR-2	Cannon CR-2
400 images	400 images	400 images
Training	Offline evaluation (labels are now available)	On-site evaluation (private labels)

### 12 participating teams All the teams proposed deep learning based solutions!



Medical Image Analysis Volume 59, January 2020, 101570



### REFUGE Challenge: A unified framework for evaluating automated methods for glaucoma assessment from fundus photographs

José Ignacio Orlando <sup>a</sup>, Huazhu Fu <sup>b</sup>, João Barbosa Breda <sup>c, d</sup>, Karel van Keer <sup>d</sup>, Deepti R. Bathula <sup>e</sup>, Andrés Diaz-Pinto <sup>f</sup>, Ruogu Fang <sup>g</sup>, Pheng-Ann Heng <sup>h</sup>, Jeyoung Kim <sup>i</sup>, JoonHo Lee <sup>j</sup>, Joonseok Lee <sup>j</sup>, Xiaoxiao Li <sup>k</sup>, Peng Liu <sup>g</sup>, Shuai Lu <sup>l</sup>, Balamurali Murugesan <sup>m</sup>, Valery Naranjo <sup>f</sup>, Sai Samarth R. Phaye <sup>e</sup>, Sharath M. Shankaranarayana <sup>n</sup> ... Hrvoje Bogunović <sup>a</sup>

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https://doi.org/10.1016/j.media.2019.101570

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## **Optic disc/cup segmentation**



### **Glaucoma classification**



#### Still a long way to go to ensure that the problem is solved

#### Future initiatives must consider

(I) Interpretability of the results  $\rightarrow$  Class activation maps

#### Interpretability of the results

LAG database: 11.760 images + image level annotations + attention maps (from "Attention Based Glaucoma Detection: A Large-Scale Database and CNN model")



### Interpretability of the results



### Interpretability of the results





Jun et al., BMVC, 2018

#### Future initiatives must consider

(I) Interpretability of the results  $\rightarrow$  Class activation maps

(II) Larger / heterogeneous data sets (age, gender, ethnicities, acquisition quality, multiple devices, comorbidities, disease stages)

## Larger + more heterogeneous data sets







http://blog.kaggle.com/2015/08/10/detecting-diabetic-retinopathy-in-eye-images/

#### Future initiatives must consider

(I) Interpretability of the results ightarrow Class activation maps

(II) Larger / heterogeneous data sets (age, gender, ethnicities, acquisition quality, multiple devices, comorbidities, disease stages)

(III) Annotations for other relevant structures (OD hemorrhages, RNFL defects, PPA)

#### Annotations for other clinical signs



(a) Cupping in the optic nerve head (ONH)



(b) Peripapillary hemorrhages





(c) Retinal nerve fiber layer (RNFL) defects

Lesions in images + Clinical parameters + Demographics + Why not clinical records?



#### We need to go beyond just the optic disc

Target other features or even clinical parameters

#### Al is guided by data, so we need data!

Multicenter collaborations + Not just images

#### Advances in DR were boosted by open science We need to (carefully, but still) release data and code!

#### Building a data set has to be the very first step

Invest time/money on preparing good data, then invest of developing the models!

## That's all!





## yotiris





https://yatiris.github.io



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# Thanks for your attention!

If you have any further questions



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