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Why Miss the Chance? Incidental Findings while Telescreening for Diabetic Retinopathy

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ABSTRACT

Purpose: To report on incidental pathological findings met while screening for Diabetic Retinopathy (DR) in Diabetes Clinics (DC) by ophthalmologist-graded digital fundus imaging.

Methods: At the DC of Pescara (central Italy), for 3,859 eyes of 1,930 consecutive patients having not undergone fundus examination in the last year, two mydriatic fundus digital images, taken with a CenterVue DRS Digital Retinal Camera, were sent along with Best Corrected Visual Acuity, on a “store-and-forward” basis, to an ophthalmologist trained in DR screening, and graded according to the UK Diabetic Eye Screening Programme. Incidental fundus abnormalities other than DR were reported.

Results: No adverse event to mydriasis was reported. One hundred and eighty eyes (4.66%) were ungradable. Among the 3,679 gradable ones, 1,105 (30.04%) showed different degrees of DR (R1 to R3), and 126 (3.42%) maculopathy (M1). Any Age-Related Macular Degeneration was present in 387 eyes (10.52%), any optic disc and parapapillary area features suspect for glaucoma in 562 eyes (15.27%), any hypertensive retinopathy in 1,263 eyes (34.33%), vitreoretinal interface disease in 252 eyes (6.84%), myopic choroidopathy in 92 eyes (2.50%), disc pallor in 31 eyes (0.84%). Mean time was 5 min for screening, 2 min for grading.

Conclusion: Teleretinography is a well-established, cost-effective procedure in DR screening. Along with increased attendance, locating a digital camera in a DC with a retina-specialist grader results in finding fundus pathologies also beyond DR, very similarly to fundus examination in an outpatient ophthalmic setting.

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Introduction

Diabetes Mellitus (DM) is a common and rapidly increasing disease¹ and a main cause of blindness worldwide.² Diabetic retinopathy (DR) is both a frequent complication^{3,4} and a marker of disease control⁵ extremely useful to diabetologists, but the attendance of diabetic patients to periodical fundus examination is universally low,^{6,7} although the need for screening programmes has been stated long since.⁸

Therefore, the application of telemedicine to DR screening by remote digital fundus images evaluation⁹⁻¹¹ has revolutionized both the attendance to screening¹² and the natural history of such complication,¹³ allowing a better knowledge of DM course and early identification of DR, amenable to treatment resulting in a strong decrease in DR-related blindness. In his review of the results of the English National Screening Programme for DR in the years 2003–2016, Scanlon¹² reported a decrease in prevalence of certifiable blindness from 4,200 to less than 1,000, and the

finding that DR was no more the leading cause of certifiable blindness amongst working age adults in UK. In England, a well-established DR screening strategy, recognized as one of the 15 essential health-care checks by Diabetes UK, produced outstanding results: between 1 April 2017 and 31 March 2018, 2,700,774 people with DM were offered eye screening, 2,232,797 people were screened, 63,675 people were referred for follow-up tests or treatment; every year, around 7,000 people with sight-threatening diabetic retinopathy are referred to hospital eye services for urgent treatment.¹⁴ The positive effect of screening patients in a Diabetes Clinic on attendance and early diagnosis of DR has been reported by Roser.¹⁵ Such efforts meet the priorities reported in the National Eye Institute – National Eye Health Education Program Five-Year Agenda 2012–2017 (at nei.nih.gov/nehep): “Research has shown that early diagnosis and timely treatment can prevent vision loss in more than 90% of people with diabetes, yet approximately half of all people with diabetic

retinopathy are diagnosed at a stage when it is too late for treatment to be effective”.

Nowadays, longer life expectancy in DM causes an increase in age-related and sight-threatening eye diseases (Age-Related Macular Degeneration – ARMD, primary open-angle glaucoma – POAG), and a prolonged effect in pathologies in which DM acts as a contributory cause: Jeganathan¹⁶ reported clinical fundus pictures associated with DM as a cause (anterior ischemic optic neuropathy, diabetic papillopathy), a recognized risk factor (open-angle glaucoma, neovascular glaucoma, ocular ischemic syndrome) or a possible risk factor (retinal vein occlusion, retinal artery occlusion).

As such conditions (and others not related to DM, like choroidal naevi and melanomas) involve the posterior pole, well targeted by teleretinography, we tested the opportunity both of adding screening for other eye conditions to screening for DR, and of putting an ophthalmologist in charge of grading DR, capable of identifying early stages of other retinal/optic nerve diseases too.

In reporting the UK initiatives in screening for DR and detection of glaucoma, macula disease, emergency eye disease in 2016, Sim¹⁷ concluded that “The use of teleophthalmology presents an immense opportunity to manage the steadily increasing demand for eye care, but challenges remain in the delivery of practical, viable, and clinically proven solutions”. This paper aims to show how eye care opportunities can be increased and delivery challenges reduced with affordable “practical, viable and clinically proven solutions”.

Materials and methods

Since 2017, at the Diabetes Clinic of the University of Chieti-Pescara Chair of Endocrinology, patients needing guideline-suggested yearly fundus examination are routinely offered digital photography during a scheduled diabetes check instead of a clinical outpatient exam, aiming to increase patients’ attendance, very low in Italy (8.2% according to the 2019 ARNO report, at www.arno.cineca.it).

Since September 1st, 2017, to October 19, 2019, 1,930 consecutive diabetic patients choosing digital eye examination (Males = 1047, Females = 883, aged 18 to 90) participated in the present study. Patients repeating retinography during the study period were considered only once, to avoid duplicating incidental findings.

While accessing the Clinic for scheduled exams, patients underwent best-corrected visual acuity (BCVA) examination with an ETDRS chart, external ocular examination and pupil dilation with one drop of Tropicamide 1% (Visumidriatic 10%, Visufarma, Italy),

by an orthoptist (screener) trained in grossly evaluating anterior chamber’s depth with lateral illumination and in recognizing signs and symptoms of intraocular pressure (IOP) elevation after mydriasis. A pair of 40x45° fundus images, one centered at the fovea and the other at the optic disc, were taken for each eye with a DRS digital retinal camera (CenterVue, Padova, Italy). Patients were informed on how to recognize and report symptoms and signs of IOP elevation, and sent back to continue their scheduled check in the Clinic. Based on a “store-and-forward” model, both images and BCVA for each patient were sent to a server, to be periodically downloaded in a territorial ophthalmic facility by an ophthalmologist (RP), specifically trained in the UK-model DR screening (www.gov.uk/health-and-social-care/population-screening-programmes-diabetic-eye) by achieving a Certificate of Higher Education in Diabetic Retinopathy Screening by the University of Gloucestershire (www.drscreening.org). Images were graded according to the UK NHS Diabetic Eye Screening Programme (DESP) standards^{18,19} (Table 1), accordingly considering BCVA ≤ 6/12 (in the absence of other causes) as a surrogate marker for macular edema; grading, incidental findings, and clinical suggestions were reported into a specific Electronic Health Record (EHR: Smart Digital Clinic, Meteda, San Benedetto del Tronto, Italy), thus being immediately available to both diabetologists and primary care physicians (via a dedicated software connecting the EHR to each one’s personal computer: QUICKConnect, openNETica, Montesilvano, Pescara, Italy). Urgent cases (DR and others), as well as patients needing instrumental exams or surgical/laser therapy were sent to the Excellence Center in Ophthalmology at the University of Chieti-Pescara in an accordingly short time.

The ophthalmologist classified incidental findings according to:

- ARMD: AREDS 2001, in AAO Preferred Practice Patterns: Age-Related Macular Degeneration²⁰ (Table 2);
- Glaucoma suspect: Ocular Hypertension Treatment Study,²¹ European Glaucoma Society²² (Table 3). As the present paper concerns screening, only patients satisfying the parameters of glaucoma suspect were included, while confirmation was obtained by a second-level facility and not reported herein;
- Hypertensive Retinopathy: Keith-Wagener-Barker²³ (Table 4).

Descriptive classifications were provided for pathologies incapable of being classified correctly enough by

Table 1. Diabetic Eye Screening Programme definitions for diabetic retinopathy.^{18,19}

Grade (Retinopathy)	Retinal lesions
R0 (None)	None
R1 (Background)	Isolated (one or more) cotton wool spots (CWS) in the absence of other features of DR Microaneurysm(s) Haemorrhage(s) Any exudate in the presence of other features of DR Any number of cotton wool spots (CWS) in the presence of other features of DR
R2 (Preproliferative)	Venous loops Venous beading (venous beading from ischaemia in diabetic retinopathy does not occur in isolation from multiple blot haemorrhages or IRMA) Venous reduplication Intraretinal microvascular abnormality (IRMA) Multiple blot haemorrhages (if uncertain, refer only in the presence of IRMA that are definitely seen)
R3S (Stable proliferative post-treatment)	Stable pre-retinal fibrosis + peripheral retinal scatter laser Stable fibrous proliferation (disc or elsewhere) + peripheral retinal scatter laser Stable R2 features (from feature based grading) + peripheral retinal scatter laser R1 features (from feature based grading) + peripheral retinal scatter laser
R3A (Active proliferative)	New vessels on disc (NVD) New vessels elsewhere (NVE) New pre-retinal or vitreous haemorrhage New pre-retinal fibrosis New tractional retinal detachment Reactivation in a previous stable R3S eye
Grade (Maculopathy)	Macular lesions (the macula is defined as that part of the retina which lies within a circle centred on the centre of the fovea whose radius is the distance between the centre of the fovea and the temporal margin of the disc)
M0 (No maculopathy)	None Any microaneurysm or haemorrhage within 1 disc diameter (DD) of the centre of the fovea if associated with a best visual acuity (VA) of $\leq 6/12$ where the cause of the reduced vision is known and is not diabetic macular edema
M1 (Maculopathy)	Exudate within 1 DD of the centre of the fovea Group of exudates within the macula: a group of exudates is an area of exudates that is greater than or equal to half the disc area and this area (of greater than or equal half the disc area) is all within the macular area Retinal thickening within 1 DD of the centre of the fovea (if stereo available) Any microaneurysm or haemorrhage within 1 DD of the centre of the fovea only if associated with a best VA of $\leq 6/12$ (if no stereo)

Table 2. Grading definitions for age-related macular degeneration.

NO	No or few small drusen (<63 micron)
Early	A combination of multiple small drusen, few intermediate drusen (63–124 micron in diameter), or mild RPE (retinal pigment epithelium) abnormalities
Intermediate	Numerous intermediate drusen At least one large druse (≥ 125 micron in diameter)
Advanced	Geographic atrophy (a sharply demarcated, usually round or oval, area of atrophy of the RPE not involving the center of the fovea) Geographic atrophy of the RPE involving the foveal center Neovascular maculopathy that includes the following: <ul style="list-style-type: none"> • choroidal neovascularization (defined as pathologic angiogenesis originating from the choroidal vasculature that extends through a defect in Bruch's membrane) • serous and/or hemorrhagic detachment of the neurosensory retina or RPE • retinal hard exudates (a secondary phenomenon resulting from chronic intravascular leakage) • subretinal and sub-RPE fibrovascular proliferation • disciform scar (subretinal fibrosis)

Table 3. Definitions for glaucoma suspect.

- Cup/Disc ratio (increasing risk ≥ 0.3 , depending on disc size)
- Cup/Disc ratio asymmetry (≥ 0.2)
- Disc haemorrhages
- Parapapillary atrophy
- Bending, bayoneting or barring of circumlinear vessels
- Myopic tilted discs
- Irregular ISNT (Inferior-Superior-Nasal-Temporal) nerve fiber layer distribution
- Neuroretinal rim: diffuse narrowing and/or localized notching

means of ophthalmoscopy alone (e.g. Vitreo-Retinal Interface Abnormalities – VRIA).

Patients released informed consent for both the procedure and the personal data management according to the General Data Protection Regulation (European Union Regulation 2016/679). Patients' data were anonymized. Whatever health and safety aspect related to the procedure was compiled. This study followed the principles of the

Table 4. Keith–Wagener–Barker classification of hypertensive retinopathy.²³

Grade	Features
I	Mild generalized retinal arteriolar narrowing
II	Definite focal narrowing and arteriovenous nicking
III	The above and retinal haemorrhages, exudates and cotton-wool spots
IV	Severe grade III and papilloedema

Declaration of Helsinki and was authorized by the Ethical Committee of the University of Chieti-Pescara.

Results

No adverse event to mydriatic drops was recorded. One eye was excluded owing to phthisis bulbi.

Out of the remaining 3,859 eyes, 180 (4.66%) were ungradable, and patients were sent to the ophthalmic facility for clinical examination. These eyes were excluded from the total sample. Among the remaining 3,679 eyes, 1,105 (30.04%) showed different degrees of DR (R1: 840 eyes, 22.83%; R2: 111 eyes, 3.02%; R3A: 9 eyes, 0.24%; R3S: 145 eyes, 3.94%) and 126 showed maculopathy (M1, 3.42%), and patients were suggested an appropriate follow-up or intervention in accordance with the Italian guidelines for Diabetic Retinopathy (AMD-SID: Italian Standards for Diabetes Care 2016, at www.standarditaliani.it).

As for incidental findings, ARMD was present in 387 eyes (10.52%): early: 166 eyes (4.51%); intermediate: 173 eyes (4.70%); advanced: 48 eyes (1.30%); optic disc and/or parapapillary area features suspect for glaucoma in 562 eyes (15.27%); hypertensive retinopathy (degrees 1–2) in 1,263 eyes (34.33%); VRIA in 252 eyes (6.84%), choroidal naevi in 53 eyes (1.44%); any form of venous occlusion in 19 eyes (0.51%); myopic choroidopathy in 92 eyes (2.50%); optic disc pallor without signs of glaucoma suspect in 31 eyes (0.84%).

1,336/1,926 patients (69.36%) reported systemic anti-hypertensive therapy and 119/1,823 (6.52%) topical anti-glaucoma therapy. Not all patients were sure to remember to suffer or not from arterial hypertension or glaucoma.

For each patient, BCVA measurement plus image taking lasted from 4 to 7 min (mean: 5), and grading from 1 to 3 min (mean: 2).

Discussion

Chen²⁴ reported DM acting as a risk factor for ARMD, stronger for late than earlier stages. Chew²⁵ stressed the impending diffusion of ARMD in the near future,

stating that the number of individuals with AMD globally will approach 196 million in 2020 and 288 million by 2040, and evidenced the cost-effectiveness of adding the ARMD screening to the DR one. Our high-ranking percentage (all stages 10.52%) confirms the utility of such opportunity. Periodical screening for ARMD even in asymptomatic patients has provided positive results yet.²⁶ The recent observations by Hallak et al.²⁷ that total en face area of drusen in a circular area 3 mm from the fovea is associated with conversion to the wet form, and by Domalpally et al.²⁸ that reticular pseudodrusen have an increased risk of progression to geographic atrophy, further stress the importance of fundus imaging in early diagnosis. In a recent paper on a remote diagnosis imaging model, Hadziahmetovic²⁹ concluded that “Good operational characteristics found in this study suggest the feasibility of using this model to screen and refer patients to a retinal specialist”.

POAG – another leading cause of blindness worldwide² – affects an estimated 2.2 million people in the United States, and that number is likely to increase to 3.3 million in 2020 as the population ages,³⁰ facilitated by the fact that it often goes undetected due to the absence of warning symptoms or signs in the early stages. It is accepted that diabetic patients undergo POAG more frequently than the general population,³¹ and our percentage of patients in glaucoma therapy (6.52%) seems to support such finding. Gangwani³² reported an overall prevalence of 1.8% for confirmed glaucoma in the population of diabetic patients screened, identifying an abnormal (>0.6) cup/disc ratio (CDR) in 3.7% of 2,182 patients undergoing a DR screening. Ong³³ reported 216 (1.9%) fundus-images based glaucoma suspect out of 11,565 patients undergoing the UK DR screening in 2006–2008. As for the problem of monoscopic (proper of DR screening) versus stereoscopic evaluation of the optic disc, Myers³⁴ reported two studies^{35,36} showing a satisfactory interobserver expert agreement in evaluating the CDR and other glaucomatous characteristics, and Newman-Casey³⁷, reviewing studies about monoscopic photography in glaucoma screening, reported that differences in sensitivity and specificity against both dilated fundus examination by an ophthalmologist and stereoscopic photography are not striking, stating that monoscopic digital photography could be an important component of glaucoma assessment, capable of being provided at the primary care level too. Our high-ranking percentage of glaucoma suspects (15.27%) is likely due to considering each and every sign of suspect disc and parapapillary area (Table 3), and not only the vertical CDR, thus stressing the advantage of an ophthalmologist grading DR images and finding

comorbidities (or suspects). Verifying how many glaucoma suspects turned into glaucoma patients was not an aim of the present study.

Vitreoretinal interface³⁸ is a leading factor in interpreting and treating diabetic maculopathy as well as other pathologies like vitreomacular traction or macular hole, and its abnormalities (VRIA) are more frequent in diabetic eyes.³⁹ VRIA are finely diagnosed by OCT, but fundus images allow diagnosis of some presentations: in a UK experience,⁴⁰ photography allowed diagnosing epiretinal membrane (ERM) and full-thickness macular hole, but missed other forms of VRIA needing OCT for diagnosis: vitreomacular traction at the fovea or elsewhere, partial thickness macular hole, and foveoschisis. A review by Xiao⁴¹ stressed the variability in grading different kinds and stages of ERMs, but reported an overall usefulness of monoscopic photography. Our percentage (6.84%) is not far from the overall incidence of 8.4% (cellophane macular reflex + preretinal macular fibrosis) in 2,476 electronically imaged patients examined by Yang,⁴² who attributes to diabetes (defined as fasting plasma glucose \geq 7 mmol/L) a 1.41 odds ratio, to DR prevalence a 1.39 odds ratio, and to DR stage an 1.13 odds ratio in VRIA development.

Long⁴³ reported that up to 75% of people with DM have arterial hypertension (69.37% of our patients reported anti-hypertensive therapy), and it is well known that hypertension has long been followed up by fundus examination too. In hypertensive retinopathy, fundus examination is a well-established diagnostic mainstay (https://eyewiki.aao.org/Hypertensive_retinopathy), with some doubts emerging in the last years related to its value as a marker of cardiovascular risk in elder people⁴⁴. We found 34.33% of eyes showing grades 1–2 of the Keith–Wagener–Barker’s classification.

Other pathologies that can be reported in DR screening are: retinal vein occlusion (RVO, prevalence: 0.5–2% for branch RVO and 0.1–0.2% for central RVO; estimated 15-year incidence rate: 1.8% for branch RVO and 0.2% for central RVO⁴⁵); central retinal artery occlusion (incidence: 1/100,000⁴⁶); non-arteritic anterior ischemic optic neuropathy (annual incidence: 2.3–10.2/100,000 in the USA⁴⁷); arteritic anterior ischemic optic neuropathy (5–10% of all anterior ischemic optic neuropathies⁴⁸); diabetic papillopathy (prevalence: 1.4% in diabetic patients⁴⁹); signs of optic neuritis in multiple sclerosis patients (initial presentation in ~20% of patients, potentially occurring during the course of the disease in 50% of patients⁵⁰); choroidal naevi (prevalence in USA adults aged > 40: 5%⁵¹); choroidal melanoma (overall incidence in the USA: 5/1,000,000⁵²). Among all these, we reported 17 eyes with any form of venous occlusions (0.48%) and

39 eyes with posterior pole choroidal naevi (1.10%); other fundi amenable of clinical diagnosis were the 35 (0.98%) showing optic disc pallor not suitable of being glaucoma suspects and the 92 (2.50%) showing myopic choroidopathy: both categories were sent to the ophthalmic facilities owing to the possibility of systemic or ocular complications.

Two more advantages of screening performed by an ophthalmologist concern population eye needs assessment.

Consider the suggested periodicity of eye examinations in a general population. The American Academy of Ophthalmology (AAO) Preferred Practice Pattern Diabetic Retinopathy⁵³ suggests that “People with Type 1 diabetes should have annual screenings for diabetic retinopathy beginning 5 years after the onset of their disease, whereas those with Type 2 diabetes should have a prompt examination at the time of diagnosis and at least yearly examinations thereafter”; the AAO Preferred Practice Pattern Comprehensive Adult Medical Eye Evaluation⁵⁴ states: “The suggested fundus examination periodicity for people with no risk factors is: under 40: 5–10 years; 40–54: 2–4 years; 55–64: 1–3 years; 65 or older: 1–2 years (for all classes: moderate quality, strong recommendation). For people with risk factors for glaucoma: 1–3 years for 40–54, 1–2 years for others, depending on the ophthalmologist’s judgement (for all classes: moderate quality, strong recommendation)”. Plotting the above recommendations suggests that undergoing yearly DR telescreening in people otherwise free of ocular symptoms and signs provides patients with the opportunity of a frequent fundus examination (in younger age classes even more frequently than suggested), thus allowing an early identification of whatsoever pathologic findings in the posterior pole and glaucoma suspects.

Moreover, telescreening can strongly contribute in building up a whole integrated care path for DR involving primary care providers (PCPs). Liu et al.⁵⁵ reported that their recommendations to diabetic patients to undergo teleretinal screening are, along with convenience (e.g., same-day scheduling, location, being a quick procedure) and comprehension of the importance, the most important facilitator to participate in screening initiatives. From the PCPs’ viewpoint, the main facilitators to join a screening initiative were ease of referral process and results communication, perceived benefits to patients (e.g., convenience, cost), improved adherence, benefits to health-care organization (e.g., increased reimbursement for improved quality metrics). PCPs nevertheless play a central role in motivating citizens to take care of their own eyes, what they often neglect by themselves: a 2016 UK Royal

National Institute for the Blind survey (available at www.rnib.org.uk/stateofthenation) evidenced that 27% of the general population aged 18 and above had not undergone even a simple sight test in the last two years. This confirms that citizens frequently do not attend scheduled ophthalmic examinations, thus often missing the early diagnosis of potential causes of low vision and blindness. Obvious consequences both on each one's health (not only eye health, as low vision strongly affects quality of life,^{56,57} is a factor in falls and fall-related fractures⁵⁸ and a cause of depression^{59,60}), and on the nations' social security systems result from such health literacy weakness.

There is a long-standing debate about who should be a grader (trained nonprofessional, optometrist, orthoptist, family physician, diabetologist, ophthalmologist), with recent encouraging results provided by optometrists supervised by retina specialists.⁶¹ In view of the above considerations, and along with both Boucher⁶² (reporting a much longer than ours series of incidental findings), and the AAO Telemedicine Statement⁶³ (point F – Reading: “An ophthalmologist or team of image readers under ophthalmologist supervision should perform image grading, image reading with recommendations, and results reporting.”), we believe that an ophthalmologist (better, a retina specialist, as shown for DR⁶⁴) can identify not only DR but each and every incidental finding, and collaborate with neurologists to identify risk factors for cerebrovascular disease^{65,66} and microvascular findings related to cognitive impairment.⁶⁷ A non-ophthalmologist undergoing both an accurate training for DR-specific and other types of retinal/optic nerve abnormalities, and a strict quality assurance control (as in the UK Diabetic Eye Screening Programme), could act as a first grader in identifying normal/any abnormal images.

With a retina-specialist examining images, our percentages in reporting pathological findings are slightly higher than those reported by Park and Mansberger,⁶⁸ possibly owing to different population and methodology characteristics (undilated patients, only vertical C/D ratio for glaucoma suspect, training of graders), and strongly support their findings that one in four eyes had at least one eye disease, and that 17% of eyes had diseases other than DR, thus confirming the necessity to widen the diagnostic spectrum when examining fundus images.

In our experience, we gave DR screening the meaning of providing diabetologists with a tool to monitor the disease, as it happened and still happens when the diabetologist him/herself examines the posterior pole by direct ophthalmoscopy, thus making fundus examination diabetologist-oriented more than ophthalmologist-oriented. Such pathway completely satisfies both the Wilson and Jungner's⁶⁹ and the Andermann's⁷⁰

criteria for screenings, excellently adapted to DR by Das.⁷¹

Comparing our results (in terms of: patients' uptake; screening worktime – about 5 min for the screener, about 2 min for the grader for each patient; time between screening, grading and results' availability in the EHR; timeliness of referral to ophthalmic facilities for a thorough diagnosis and therapy) to the setting of outpatient clinical fundus examination, suggests that shifting fundus examinations for DR from the ophthalmic outpatient to the in-the-Diabetic-Clinic teleretinography setting, could become a cost- and health-effective procedure, with the side effect of “not to miss the chance” both to identify incidental findings and to improve health literacy (making patients aware of their pathologies and suggesting them to undergo regular complete eye examinations).

The “World Health Organization Universal Eye Health: a Global Action Plan 2014–2019” (who.int/blindness/actionplan/en/) states that – Developing and implementing national policies and plans for the prevention of avoidable visual impairment remain the cornerstone of strategic action. Some programmes against eye diseases have had considerable success in developing and implementing policies and plans, however, the need remains to integrate eye disease control programmes into wider health-care delivery systems, and at all levels of the health-care system. In increasing numbers, countries are acquiring experience in developing and implementing effective eye health services and embedding them into the wider health system. These experiences need to be better documented and disseminated so that all countries can benefit from them. Operational research will provide evidence on ways to overcome barriers in service provision and uptake, and improvements in appropriate cost-effective strategies and approaches for meeting ever-growing public health needs for improving and preserving eye health in communities.

Thus, each effort to make citizens' access to health easier and to improve literacy is a holy one, and teleretinography, as above discussed, has all the characteristics to be a preeminent one, helping to bring fundus examination to patients instead of waiting for patients to attend an outpatient ophthalmic facility.

Declaration of interest statement

None of the authors have whatsoever financial interest in this study.

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Disclosure of interest

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Publication elsewhere

This submission has not been published elsewhere previously and is not simultaneously being considered for any other publication.

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References

- World Health Organization. Diabetes. <https://www.who.int/news-room/fact-sheets/detail/diabetes> (accessed march 16, 2019)
- World Health Organization. Blindness and vision impairment. <https://www.who.int/news-room/fact-sheets/detail/blindness-and-visual-impairment> (accessed march 16, 2019)
- Lee R, Wong TY, Sabanayagam C. Epidemiology of diabetic retinopathy, diabetic macular edema and related vision loss. *Eye and Vision*. 2015;2:17. doi:10.1186/s40662-015-0026-2.
- Sabanayagam C, Banu R, Chee ML, et al. Incidence and progression of diabetic retinopathy: a systematic review. *Lancet Diabetes Endocrinol*. 2019 Feb;7(2):140–149. doi:10.1016/S2213-8587(18)30128-1.
- American Diabetes Association. Diabetic retinopathy: a position statement by the American diabetes association. *Diabetes Care*. 2017;40:412–418. doi:10.2337/dc16-2641
- Scanlon PH, Stratton IM, Leese GP. et al. Screening attendance, age group and diabetic retinopathy level at first screen. *Diabet Med*. 2016;33:904–911. doi:10.1111/dme.12957.
- Lawrenson JG, Graham-Rowe E, Lorencatto F, et al. Interventions to increase attendance for diabetic retinopathy screening. *Cochrane Database Syst Rev*. 2018; (1):Art. No.: CD012054 doi:10.1002/14651858.CD012054.pub2.
- Kohner EM, Porta M. Protocols for screening and treatment of diabetic retinopathy in Europe. *Eur J Ophthalmol*. 1991 Jan-Mar;1(1):45–54. doi:10.1177/112067219100100109.
- Scanlon PH, Dirani M, van Wijngaarden P. Screening for sight-threatening diabetic retinopathy: an update. Symposium - Diabetic Retinopathy Update. *Egypt Ret J*. 2014;2(1):3–18. doi:10.4103/2347-5617.152479.
- Tozer K, Woodward MA, Newman-Casey PA. Telemedicine and diabetic retinopathy: review of published screening programs. *J Endocrinol Diab*. 2015;2(4):1–10. doi:10.15226/2374-6890/2/4/00131.
- Panwar N, Huang P, Lee J, et al. Fundus photography in the 21st century—a review of recent technological advances and their implications for worldwide healthcare. *Telemed J E Health*. 2016;22(3):198–208. doi:10.1089/tmj.2015.0068.
- Scanlon PH. The english national screening programme for diabetic retinopathy 2003–2016. *Acta Diabetol*. 2017;54:515–525. doi:10.1007/s00592-017-0974-1.
- Shi L, Wu H, Dong J. et al. Telemedicine for detecting diabetic retinopathy: a systematic review and meta-analysis. *Br J Ophthalmol*. 2015;99:823–831. doi:10.1136/bjophthalmol-2014-30563.
- Public Health England. *Breaking down barriers to diabetic eye screening attendance*, <https://phscreening.blog.gov.uk/2018/11/14/breaking-down-barriers-to-diabetic-eye-screening-attendance/> (accessed march 16, 2019)
- Roser P, Kalscheuer H, Groener JB, et al. Diabetic Retinopathy screening ratio is improved when using a digital, nonmydriatic fundus camera onsite in a diabetes outpatient clinic. *J Diabetes Res*. 2016. doi:10.1155/2016/4101890. [Article ID 4101890].
- Jeganathan VSE, Wang JJ, Wong TY. Ocular associations of diabetes other than diabetic retinopathy. *Diabetes Care*. 2008;31(9):1905–1912. doi:10.2337/dc08-0342.
- Sim DA, Mitry D, Alexander P, et al. The evolution of teleophthalmology programs in the United Kingdom: beyond diabetic retinopathy screening. *J Diabetes Sci Technol*. 2016;10(2):308–317. doi:10.1177/1932296816629983.
- NHS Screening Programmes – Diabetic Eye. Diabetic eye screening feature-based grading forms. Guidance on standard feature-based grading forms to be used in the NHS Diabetic Eye Screening Programme. Version 1.4, 1 November 2012 https://www.gov.uk/government/uploads/system/uploads/attachment_data/file/402295/Feature_Based_Grading_Forms_V1_4_1Nov12_SSG.pdf
- NHS Public Health England - NHS Diabetic Eye Screening Programme. Grading definitions for referable disease. https://assets.publishing.service.gov.uk/government/uploads/system/uploads/attachment_data/file/582710/Grading_definitions_for_referrable_disease_2017_new_110117.pdf Published January 2017
- American Academy of Ophthalmology Retina/Vitreous Panel. *Preferred Practice Pattern® Guidelines. Age-Related Macular Degeneration*. San Francisco, CA: American Academy of Ophthalmology; 2015. www.aao.org/ppp.
- Gordon MO, Beiser JA, Brandt JD, et al. The ocular hypertension treatment study. baseline factors that predict the onset of primary open-angle glaucoma. *Arch Ophthalmol*. 2002;120:714–720. doi:10.1001/archophth.120.6.714.
- European Glaucoma Society terminology and guidelines for glaucoma, 4th edition 2014 supported by the European Glaucoma Society Foundation: 48–54. www.eugs.org. doi:10.1136/bjophthalmol-2016-EGSguideline.001.
- Keith NM, Wagener HP, Barker NW. Some different types of essential hypertension: their course and prognosis. *Ann J Med Sci*. 1939;191:332–343. doi:10.1097/00000441-193903000-00006.
- Chen X, Rong SS, Xu Q, et al. Diabetes mellitus and risk of age-related macular degeneration: a systematic review and meta-analysis. *PLoS ONE*. 2014;9(9): e108196. doi:10.1371/journal.pone.0108196.
- Chew EY, Schachat AP. Should we add screening of age-related macular degeneration to current screening

- programs for diabetic retinopathy? *Ophthalmology*. 2015;122(11):2155–2156. doi:10.1016/j.ophtha.2015.08.007.
26. Tamura H, Goto R, Akune Y, et al. The clinical effectiveness and cost-effectiveness of screening for age-related macular degeneration in Japan: a Markov modeling study. *PLoS ONE*. 2015;10(7):e0133628. doi:10.1371/journal.pone.0133628.
 27. Hallak JA, de Cisternes L, Osborne A, et al. Imaging, genetic, and demographic factors associated with conversion to neovascular age-related macular degeneration. secondary analysis of a randomized clinical trial. *JAMA Ophthalmol*. 2019;137(7):738–744. doi:10.1001/jamaophthalmol.2019.0868.
 28. Domalpally A, Agron E, Pak JW et al. Prevalence, risk and genetic association of reticular pseudodrusen in age-related macular degeneration. AREDS2 report 20. Published online July 29, 2019 DOI: 10.1016/j.ophtha.2019.07.022
 29. Hadziahmetovic M, Nicholas P, Jindal S, et al. Evaluation of a remote diagnosis imaging model vs dilated eye examination in referable macular degeneration. *JAMA Ophthalmol*. 2019 May 16. doi:10.1001/jamaophthalmol.2019.1203.
 30. American Academy of Ophthalmology. *Preferred Practice Pattern® Guidelines. Primary Open Angle Glaucoma*. San Francisco, CA: American Academy of Ophthalmology; 2016. www.aao.org/ppp.
 31. American Academy of Ophthalmology. *Preferred Practice Pattern® Guidelines. Primary Open Angle Glaucoma Suspect*. San Francisco, CA; American Academy of Ophthalmology; 2016. doi:10.1016/j.ophtha.2015.10.055.
 32. Gangwani RA, McGhee SM, Laj JS, et al. Detection of glaucoma and its association with diabetic retinopathy in a diabetic retinopathy screening program. *J Glaucoma*. 2016;25(1):101–105. doi:10.1097/IJG.000000000000138.
 33. Ong HS, Levin S, Ahmed F, et al. Is glaucoma screening using national diabetic retinopathy programme images effective? *ARVO Ann Meet Abst IOVS*. 2009;50(13):4090.
 34. Myers JS, Fudemberg SJ, Lee D. Evolution of optic nerve photography for glaucoma screening: a review. *Clin Exp Ophthalmol*. 2018;46:169–176. doi:10.1111/ceo.13138.
 35. Parkin B, Shuttleworth G, Costen M. et al. A comparison of stereoscopic and monoscopic evaluation of optic disc topography using a digital optic disc stereo camera. *Br J Ophthalmol*. 2001;85:1347–1351. doi:10.1136/bjo.85.11.1347.
 36. Chan HH, Ong CN, Kong YX, et al. Glaucomatous optic neuropathy evaluation (GONE) project: the effect of monoscopic versus stereoscopic viewing conditions on optic nerve evaluation. *Am J Ophthalmol*. 2014;157(5):936–944. doi:10.1016/j.ajo.2014.01.024.
 37. Newman-Casey PA, Verkade AJ, Oren G, Robin AL. Gaps in Glaucoma care: a systematic review of monoscopic disc photos to screen for glaucoma. *Expert Rev Ophthalmol*. 2014;9(6):467–474. doi:10.1586/17469899.2014.967218.
 38. Agarwal D, Gelman R, Prospero Ponce C, et al. The vitreomacular interface in diabetic retinopathy. *J Ophthalmol*. 2015. [Article ID 392983]. doi:10.1155/2015/392983.
 39. Liu L, Yue S, Wu J, et al. The prevalence and distribution of vitreoretinal interface abnormalities among urban community population in China. *J Ophthalmol*. 2015. [Article ID 742686]. doi:10.1155/2015/742686.
 40. McKibbin M, Farragher T, Shickle D. UK biobank eye and vision consortium. vitreoretinal interface abnormalities in middle-aged adults with visual impairment in the UK Biobank study: prevalence, impact on visual acuity and associations. *BMJ Open Ophth*. 2017;1:e000057. doi:10.1136/bmjophth-2016-000057.
 41. Xiao W, Chen X, Yan W. et al. Prevalence and risk factors of epiretinal membranes: a systematic review and meta-analysis of population-based studies. *BMJ Open*. 2017;7:e014644. doi:10.1136/bmjopen-2016-014644.
 42. Yang Y, Yan YN, Wang YX, et al. Ten-year cumulative incidence of epiretinal membranes assessed on fundus photographs. The Beijing Eye Study 2001/2011. *PLoS ONE*. 2018;13(4):e0195768. doi:10.1371/journal.pone.0195768.
 43. Long AN, Dagogo-Jack S. The comorbidities of diabetes and hypertension: mechanisms and approach to target organ protection. *J Clin Hypertens (Greenwich)*. 2011;13(4):244–251. doi:10.1111/j.1751-7176.2011.00434.x.
 44. Cuspidi C, Sala C, Grassi G. Updated classification of hypertensive retinopathy: which role for cardiovascular risk stratification? *J Hypertens*. 2015;33:2204–2206. doi:10.1097/HJH.0000000000000733.
 45. Laouri M, Chen E, Looman M. et al. The burden of disease of retinal vein occlusion: review of the literature. *Eye*. 2011;25:981–988. doi:10.1038/eye.2011.92.
 46. Varma DD, Cugati S, Lee AW. et al. A review of central retinal artery occlusion: clinical presentation and management. *Eye*. 2013;27:688–697. doi:10.1038/eye.2013.25.
 47. Chen T, Song D, Shan G, et al. The association between diabetes mellitus and nonarteritic anterior ischemic optic neuropathy: a systematic review and meta-analysis. *PLoS ONE*. 2013;8(9):e76653. doi:10.1371/journal.pone.0076653.
 48. [https://eyewiki.aao.org/Arteritic_Anterior_Ischemic_Optic_Neuropathy_\(AAION\)](https://eyewiki.aao.org/Arteritic_Anterior_Ischemic_Optic_Neuropathy_(AAION))
 49. Al-Hinai AS, Al-Abri MS, Al-Hajri RH. Diabetic papillopathy with macular edema treated with intravitreal bevacizumab. *Oman J Ophthalmol*. 2011;4(3):135–138. doi:10.4103/0974-620X.91270.
 50. Kale N. Optic neuritis as an early sign of multiple sclerosis. *Eye Brain*. 2016;8:195–202. doi:10.2147/EB.S54131.
 51. Chien JL, Sioufi K, Surakiatchanukul T, et al. Choroidal nevus: a review of prevalence, features, genetics, risks, and outcomes. *Curr Opin Ophthalmol*. 2017;28(3):228–237. doi:10.1097/ICU.0000000000000361.
 52. Kranz BA, Dave N, Komatsubara KM. et al. Uveal melanoma: epidemiology, etiology, and treatment of primary disease. *Clin Ophthalmol*. 2017;11:279–289. doi:10.2147/OPH.S89591.
 53. American Academy of Ophthalmology Retina/Vitreous Panel. Preferred Practice Pattern® Guidelines. *Diabetic Retinopathy*. San Francisco, CA: American Academy of Ophthalmology; 2017:4. www.aao.org/ppp

54. American Academy of Ophthalmology. Preferred Practice Pattern® Guidelines. *Comprehensive Adult Medical Eye Evaluation*. San Francisco, CA: American Academy of Ophthalmology; 2016:224. www.aaopt.org/ppp
55. Liu Y, Zupan NJ, Swearingen R. et al. Identification of barriers, facilitators and system-based implementation strategies to increase teleophthalmology use for diabetic eye screening in a rural US primary care clinic: a qualitative study. *BMJ Open*. 2019;9:e022594. doi:10.1136/bmjopen-2018-022594.
56. Swagerty DL Jr. The impact of age-related visual impairment on functional independence in the elderly. *Kansas Med*. 1995;96:24–26.
57. Scott AW, Bressler NM, Folkes S, et al. Public attitudes about eye and vision health. *JAMA Ophthalmol*. 2016;134(10):1111–1118. doi:10.1001/jamaophthalmol.2016.2627.
58. Niihata K, Fukuma S, Hiratsuka Y, et al. Association between vision-specific quality of life and falls in community-dwelling older adults: LOHAS. *PLoS ONE*. 2018;13(4):e0195806. doi:10.1371/journal.pone.0195806.
59. Casten RJ, Rovner BW. Update on depression and age-related macular degeneration. *Curr Opin Ophthalmol*. 2013;24(3):239–243. doi:10.1097/ICU.0b013e32835f8e55.
60. Diniz-Filho A, Abe RY, Cho HJ, et al. Fast visual field progression is associated with depressive symptoms in patients with glaucoma. *Ophthalmology*. 2016;123(4):754–759. doi:10.1016/j.ophtha.2015.12.014.
61. Melles RB, Conell C, Siegner SW, Tarasewicz D. Diabetic retinopathy screening using a virtual reading center. *Acta Diabetol*. Published 03 August 2019. doi:10.1007/s00592-019-01392-9.
62. Boucher MC, Desroches G, Garcia-Salines R. et al. Teleophthalmology screening for diabetic retinopathy through mobile imaging units within Canada. *Can J Ophthalmol*. 2008;43:658–668. doi:10.3129/i08-120.
63. American Academy of Ophthalmology. Task force on telemedicine in ophthalmology. telemedicine in ophthalmology information statement. American Academy of Ophthalmology 2018. <https://www.aaopt.org/clinical-statement/telemedicine-ophthalmology-information-statement>
64. Provetti-Cunha L, Alvernaz Figueredo E, Pereira Araújo H, et al. Non-mydratic fundus retinography in screening for diabetic retinopathy: agreement between family physicians, general ophthalmologists, and a retinal specialist. *Front Endocrinol*. 2018;9:art251 doi:10.3389/fendo.2018.00251.
65. Wang JJ, Baker ML, Hand PJ. et al. Transient ischemic attack and acute ischemic stroke associations with retinal microvascular signs. *Stroke*. 2011;42:404–408. doi:10.1161/STROKEAHA.110.598599.
66. Vuong LN, Thulasi P, Biousse V. et al. Ocular fundus photography of patients with focal neurologic deficits in an emergency department. *Neurology*. 2015;85:256–262. doi:10.1212/WNL.0000000000001759.
67. Lee MJ, Deal J, Ramulu PY, et al. Prevalence of retinal signs and association with cognitive status: the ARIC neurocognitive study. *J Am Geriatr Soc*. 2019 Jun;67(6):1197–1203. doi:10.1111/jgs.15795.
68. Park D, Mansberger SL. Eye disease in patients with diabetes screened with telemedicine. *Telemed Ad e-Health*. 2017;23(2):113–118. doi:10.1089/tmj.2016.0034.
69. Wilson JMG, Jungner G. *Principles and Practice of Screening for Disease*. 1968;WHO Public Health Papers 34, World Health Organization, Geneva.
70. Andermann A, Blancquaert I, Beauchamp S, Revisiting Wilson DV. Jungner in the genomic age: a review of screening criteria over the past 40 years. *Bull World Health Organ*. 2008;86(4):317–319. doi:10.2471/BLT.07.050112.
71. Das T, Raman R, Ramasamy K, Rani PK. Telemedicine in diabetic retinopathy: current status and future directions. *Middle East Afr J Ophthalmol*. 2015;22(2):174–178. doi:10.4103/0974-9233.154391.