Abstract

Introduction: Diabetic Retinopathy is the leading cause of blindness in working adults, and remains a serious public health concern, globally. Whilst extensive research has been done in this area, the prevention of diabetic retinopathy is still very poor. Contributing factors in previous studies have included shortage of Human Resources for Eye Health in primary health centres, and a shortage of technology and skill in most countries. In some countries, teleophthalmology has been introduced to assist in the management of diabetic retinopathy by capacitating Eye Health professionals in primary health care settings since diabetes is diagnosed at this level. Where implemented, management of diabetic retinopathy through teleophthalmology has further contributed to reduction in preventable blindness while minimizing turnaround times as patients no longer have to wait for an Ophthalmologist appointment. The main objective of this scoping review is to map evidence on the use of teleophthalmology in the management of diabetic retinopathy globally.

Methods and Analysis: Methodology will be guided by Arksey and O’Malley scoping review framework. We will search for literature from the following EBSCO Host databases: MEDLINE, Health source – Consumer, CINAHL and Academic Search Complete. PubMed, Google Scholar and Science Direct will also be searched for studies. Where articles are not accessible online, the author will be contacted for the article, failing which the University of KwaZulu-Natal library services will be asked for assistance. The literature search will be done for the period from January 2014 to October 2019 where all studies in English will be accepted. After the initial search, title, abstract and full text screening will be conducted by two independent reviewers. To assess the methodological rigour of included studies, the Mixed Method Appraisal tool (MMAT 2018) will be used. Study results will be presented using Preferred Reporting Items for Systematic Reviews and Meta-Analyses (PRISMA) chart.

Discussion: It is anticipated that this scoping review will identify suitable literature on diabetic retinopathy and its management through teleophthalmology. The findings of this review will be disseminated through peer-reviewed journals and information summarized will be used to guide future research.

Keywords: Diabetic retinopathy, teleophthalmology, telemedicine, Low-to-middle-income countries, Diabetic Eye Disease

1. Introduction

Diabetic retinopathy is the main cause of new onset blindness in many developed countries (Organization, 2006), and a common cause of blindness in many low-to-middle-income (LMIC) countries (Klein et al., 1992). Named one of the World Health Organisation’s (WHO) priority eye diseases leading to preventable visual impairment and blindness, diabetic retinopathy (DR) is a vascular disease of the retina which affects patients with long-standing diabetes (Organization). Due to the high blood glucose levels in diabetics’ bloodstream, damage is caused to the blood vessels in the retina causing diabetic retinopathy, which further results to diabetic macula oedema, and ultimately blindness (Organization, 2006). With a global estimate of 652.1 million in 2016 to a predicted 1.42 billion by 2045 (Cho et al., 2018), the WHO estimated that the number of people with DR who will progress to
vision-threatening diabetic retinopathy (VTDR) will increase from 37.3 million to 56.3 million in 2030, making diabetic retinopathy the leading cause of new-onset blindness in industrialised countries, and a growing concern in middle income countries (Organization, 2006).

The reality of the insidious onset of Type 2 diabetes results in many patients remaining undiagnosed for years and presenting with DR at the time of diagnosis (Cook, 2014). Type 1 diabetics, on the other hand, are diagnosed early in the course of their disease, and they typically do not develop retinopathy until years after the diagnosis is made. One of methods used to diagnose DR is teleophthalmology. This is an area of telemedicine, where digital images from a patient’s eye are taken by non-ophthalmologists and forwarded to Ophthalmology professionals for expert guidance on interventions and patient management (Sreelatha & Ramesh, 2016). With a severe shortage of Human Resources for eye Health (HReH), especially in Low-to-Middle income countries (LMIC) such as India, this method of intervention has largely relieved the eye health system. Not only has teleophthalmology reduced waiting periods and delays in the referral system, reduced costs it has further ensured that tests are not repeated, and interventions for ophthalmology-related conditions are implemented earlier (Grisolia et al., 2017).

Tele-ophthalmology has been utilised in many areas to successfully improve the reach of specialist services (Prathiba & Rema, 2011) whilst improving rate of patient screening, turnaround times for retinal disease management and access to eye health for those in remote areas (Gupta et al., 2017), ultimately resulting in the prevention of vision loss overall (Silva et al., 2016). Various reviews have been conducted to ascertain the accuracy, specificity and reliability of teleophthalmology in general eye conditions including Retinopathy of Prematurity, neuroophthalmology, glaucoma and diabetic retinopathy (Whited, 2006; Sreelatha & Ramesh, 2016; Bahaadinbeigy & Yogesan, 2011).

This scoping review will specifically focus on literature pertaining screening and management of diabetic retinopathy through teleophthalmology. Interventions used to facilitate screening practices will emerge and assist in screening strategies that enable an earlier detection of DR. The aim of the study is to map evidenced information on the use of Teleophthalmology in the management of diabetic retinopathy. It is hoped that this literature will help with exploring strategies that will enable earlier disease diagnosis, management and treatment in order to avoid unnecessary vision loss from untreated diabetic retinopathy. It is further anticipated that the emerging literature will highlight research gaps to guide future research in this area.

2. Methods

This systematic scoping review will be guided by the Arksey and O'Malley scoping review framework (Arksey & O'Malley, 2005) as per the Joanna Briggs Institute manual (Peters et al., 2015). The following 5 stages will be followed in the study:

1. Identifying the search question
2. Identifying relevant studies
3. Study selection
4. Charting the data
5. Collating, summarizing and reporting the results.

A quality assessment of the included studies will be conducted in this scoping review, to assess methodology of included studies, following recommendations by Levac et al. (Levac et al., 2010).

2.1 Identifying the Research Question

The main research question is: What is the available evidence on the use of teleophthalmology in the early management of diabetic retinopathy?

2.2 Eligibility of the Research Question

The study will employ the Population, Concept and Context (PCC) framework to determine eligible studies that respond to the research question. The framework has been shown in Table 1.
Table 1. PCC model

| Population                                                                 |
|---------------------------------------------------------------------------|
| Diabetic patients, Ocular manifestations of diabetes, Ocular complications of diabetes, diabetic retinopathy, retinal complications of diabetes. |

| Concept     |
|-------------|
| Tele-ophthalmology or Telemedicine |

| Context                       |
|------------------------------|
| All available studies (globally) |

2.3 Identifying Relevant Studies

Completed primary studies will be searched for using the following databases: PubMed, Google Scholar, Science Direct and EBSCO Host databases MEDLINE, Health source: Nursing/Academic Edition, Health source – Consumer, CINAHL and Academic Search Complete. Qualitative, quantitative and mixed method studies, published in English, will be included in the scoping review. The search will be carried out using keywords as well as Medical Subject Headings (MeSH) terms and Boolean terms. “AND” will be used when separating concepts, whilst “OR” will be used to separate synonyms.

To evaluate feasibility, a pilot study was conducted using the following search terms:

**Search:** Diabetes mellitus OR diabetic AND ocular manifestations OR ocular complications OR retinopathy OR retinal diseases AND Telemedicine OR Teleophthalmology.

2.4 Study Selection

In order to answer the research question, and select only the relevant studies, careful study selection is required. To ensure this, various inclusion and exclusion criteria will be adhered to.

2.4.1 Inclusion Criteria

Studies meeting the following criteria will be included:

- All studies available from 2014 - 2019
- Studies on diabetic human subjects with no gender restriction
- Studies on ocular manifestations of diabetes or diabetic retinopathy and telemedicine or teleophthalmology

2.4.2 Exclusion Criteria

The following exclusion criteria will be used:

- All studies from prior to 2014
- Studies on animal subjects
- Studies on non-diabetic subjects
- Studies that are not primary studies eg review articles
- Studies that only show evidence on one phenomenon and not both diabetic retinopathy and teleophthalmology

Table 2. Pilot study search results

| Date of search | 8 October 2019 |
|----------------|---------------|
| Keywords used  | diabetes mellitus OR diabetes AND ocular manifestations OR ocular complications OR retinopathy OR retinal disease AND telemedicine OR teleophthalmology |
| Period searched| 01/01/2014 – 08/10/2019 |
| Search engine  | Retrieved Sources |
| Pubmed         | 393 |
| Google scholar | 3050 |
| EBSCOHost      | 217 046 |
| Science Direct | 144 028 |
Study selection will follow thorough screening by two independent screeners, using a three-stage strategy. The Primary investigator (PI) will create screening tools for title, abstract and full text screening using Google forms. All tools will be piloted prior to being used.

The PI will initially search for studies and store them in a library in Endnote x8 created especially for this study. Thereafter, duplicates will be removed and screening will begin.

Title screening: The PI will conduct this alone to ensure that the selected studies are all in the area of interest. After finalising this, the endnote library with then be shared with the second reviewer.

Abstract screening: Adhering to specified inclusion and exclusion criteria, two independent reviewers will conduct abstract screening using piloted tools. All relevant articles that adhere to stipulated criteria. On agreement of the final articles, the next step follows.

Full text screening: Using the piloted tools to exclude articles not meeting the inclusion criteria, both independent screeners will conduct full article screening in order to decide on the final articles to be included in the study. Where the full article cannot be found online, University of KwaZulu Natal (UKZN) library services will also be used to find articles that are needed or contact authors to request full articles. Where full articles cannot be found despite all attempts to obtain them, the affected studies will be excluded. If there is any discrepancy between the reviewers, this will be resolved by a discussion, and the final articles included.

Screening results will be recorded using an adapted Preferred Reporting Items for Systematic Review and Meta-Analyses (PRISMA) guideline (Moher et al., 2009), summarised in Figure 1.
Figure 1. Flow chart showing literature search and selection of studies.
2.4 Data Charting

A data charting tool (Table 3) will be formulated and piloted prior to being used. This tool will be piloted by two independent reviewers on 5 randomly sampled, included articles to ensure consistency with the study. The tool will then be modified where necessary, following this exercise. Data charting will then be effected. Where updating becomes necessary during data charting, this will be done frequently. Data will be categorised and arranged to classify the various outcomes and themes to be analysed from the included studies. Extracted data will also be recorded on the data extraction tool.

2.5 Collating, Summarizing and Reporting of Results

As the aim of this study is to map evidence on the use of teleophthalmology in managing diabetic retinopathy, data extracted should be in line with the areas of interest. Thematic content analysis from the extracted data will look at a few pivotal areas in the use of teleophthalmology. Themes to be explored will be focused along the following outcomes: diagnostic accuracy of teleophthalmology, effectiveness of DR screening using specific features in a camera, and teleophthalmology increasing the rate of DR evaluation in rural settings where there is a shortage of human resources for eye health.

Emerging and resulting themes will then be analysed thoroughly to determine their relevance and response to the research question. This will all be done with a careful cognisance of the aim of the study to ensure that outcomes and results adhere to the main aim, whilst also exposing all gaps in the existing pool of knowledge, existing policies and current practices.

2.6 Quality Appraisal

The Levac et al recommendations for quality appraisal of included studies would be performed by two reviewers (Levac et al., 2010). The Mixed Method Appraisal Tool (MMAT) version 2018 is preferred as it is able to assess all common types of study methodologies and design (Hong et al., 2018). Quality assessment will look at the aim of each study, methodology, study design, and participant adequacy in each study. Appraisal of studies will include an actual scoring of study quality, ranging from 25% for low quality 100% for areas in the study that score well, following the guidance of the MMAT tool.

Table 3. Data Extraction table

| Author and Date |
|-----------------|
| Study title     |
| Study design    |
| Country income classification |
| Mean age of participants |
| Duration of condition |
| Digital device used |
| Who takes the picture |
| Technology used to forward picture (e.g. email) |
| Who interprets the picture |
| Type of program used |
| Mydriatic used |
| Intervention strategy communicated |
| Turnaround time |
| Other systemic diseases |
| Other complications |
| Emerging themes |
| Significant findings |
3. Discussion
With a severe shortage of Human Resources for eye Health (HReH) found in the SSA (Palmer et al., 2014), teleophthalmology was introduced in rural health centres to improve quality of medical services and outreach. This was done with the aim to improve services in rural areas to that of services in urban areas (Prathiba & Rema, 2011). Patient screening rate and satisfaction has been improved through the incorporation of teleophthalmo logical screening, using various digital gadgets to capture retinal pictures, and send them to an Ophthalmologist to identify DR and prevent vision loss (Silva et al., 2016). The introduction of teleophthalmology in India has resulted in increased access to eye health in remote areas (Gupta et al., 2017).

As diabetic retinopathy continues to emerge as one of the leading causes of visual impairment and blindness globally (Mabaso, 2008), VISION 2020: Right to Sight goals for reducing preventable blindness need to be evaluated at a local level. As a priority disease, and the third leading cause of preventable blindness in South Africa (Mabaso & Oduntan, 2014), it is vital that earlier screening be prioritised, and earlier interventions be made for patients to be diagnosed prior to complications and development of the disease. Tele-ophthalmology has proven successful in narrowing this gap and enabling earlier diagnosis and management of diabetic retinopathy (Hautala et al., 2014). Earlier interventions and diagnosis at Primary Health Level ensure that the patient is assisted holistically at one clinic visit. It also ensures that Ophthalmology involvement is accelerated, and blindness prevented much earlier in the disease.

4. Conclusion
It is envisaged that this scoping review will help to identify from the existing literature and cost effective methods that could be introduced at primary health care level to reduce the burden of avoidable blindness secondary to DR. According to the authors’ knowledge no previous scoping reviews on this matter have been done in South Africa or Africa. The result of this review will be useful in determining areas needing further research. This study may pave a way towards improving the quality of eye care services at PHC level and it may add to an enhanced diabetic retinopathy screening and management system for LMICs.

4.1 Ethics and Dissemination
No ethical approval is required as the review will be done on published data. The results of the study will published through peer-reviewed journals. All data utilised is already available in the public domain.

Acronyms
PI: Primary Investigator
RE: Refractive error
DR: diabetic retinopathy
HReH: Human Resources for Eye Health
LMIC: Low-to-middle-income countries
SSA: Sub-Saharan Africa
UKZN: University of KwaZulu-Natal

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Authors’ Contributions
TZ and ZXK conceptualized the study. TZ prepared the manuscript under the guidance and supervision of ZXK. Both authors contributed to the development and design of the study. TZ drafted the manuscript and ZXK reviewed it. Both authors contributed to the final version of the manuscript and agree to publication thereof.

Competing Interests Statement
The authors declare that there are no competing or potential conflicts of interest.

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