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Patient and Health Care Provider Knowledge of Diabetes and Diabetic Microvascular Complications: A Comprehensive Literature Review

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Abstract

Diabetic retinopathy, neuropathy, and nephropathy occur in more than 50% of people with diabetes, contributing substantially to morbidity and mortality. Patient understanding of these microvascular complications is essential to ensure early recognition and treatment of these sequalae as well as associated symptoms, yet little is known about patient knowledge of microvascular sequalae. In this comprehensive literature review, we provide an overview of existing knowledge regarding patient knowledge of diabetes, retinopathy, neuropathy, and nephropathy. We also discuss health care provider's knowledge of these sequalae given that patients and providers must work together to achieve optimal care. We evaluated 281 articles on patient and provider knowledge of diabetic retinopathy, neuropathy, and nephropathy as well as predictors of improved knowledge and screening practices. Results demonstrated that patient and provider knowledge of microvascular segualae varied widely between studies, which may reflect sociocultural or methodologic differences. Knowledge assessment instruments varied between studies with limited validation data and few studies controlled for confounding. Generally, improved patient knowledge was associated with greater formal education, longer diabetes duration, and higher socioeconomic status. Fewer studies examined provider knowledge of sequalae, yet these studies identified multiple misconceptions regarding appropriate screening practices for microvascular complications and the need to screen patients who are asymptomatic. Further investigations are needed that use well validated measures, control for confounding, and include diverse populations. Such studies will allow identification of patients and providers who would benefit from interventions to improve knowledge of microvascular complications and, ultimately, improve patient outcomes.

Keywords

understanding, nephropathy, neuropathy, retinopathy, nurse, pharmacist

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Abbreviations

ADA, American Diabetes Association

CI, confidence interval

CKD, chronic kidney disease

DKD, diabetic kidney disease

GP, general practitioner

HbA1c, hemoglobin A1C

KAP, knowledge attitudes and practices

OR, odds ratio

PRISMA, preferred reporting items for systematic reviews and meta-analysis

T1D, type 1 diabetes

T2D, type 2 diabetes

UK, United Kingdom

US, United States

1. Introduction

The global prevalence of diabetes has reached epidemic proportions [1]. In 2019, 463 million people worldwide had either type 1 (T1D) or type 2 (T2D) diabetes [1], placing substantial socioeconomic burdens on health care systems globally. If trends persist, 700 million people will be affected by diabetes by 2045 [1]. The reasons for this growth are manifold. An increase in diabetes risk factors, such as obesity and a sedentary lifestyle, against the setting of aging conspire to increase diabetes onset and prevalence. Socioeconomic forces and demographic factors are also contributing to the rise in diabetes. Countries with growing economies, income levels, and populations, such as in the Middle East, North Africa, and Asia, are driving an increase in T2D prevalence [1]. By 2045, Pakistan will overtake the United States as the third largest population with diabetes.

Diabetes causes several microvascular complications, including retinopathy, nephropathy, and neuropathy, which increase morbidity and mortality. Diabetic retinopathy is the leading cause of moderate and severe vision impairment in working age adults [2]. Neuropathy similarly can impact as many as half of individuals with diabetes, which impairs gait and stability and increases the risk of foot ulcers, ultimately leading, if left untreated, to non-traumatic lower amputations [3]. Diabetic kidney disease (DKD), *i.e.*, diabetic nephropathy, has an estimated prevalence of 25% of T1D and 30 to 40% of T2D patients [4], and can lead, in the end-stages, to death. Indeed, in 2019, 4.2 million people worldwide died from diabetes-related complications [1].

Unfortunately, the growing diabetes prevalence is driving an increase in diabetes-related complications. Long-term diabetic complications can be present at the time of or occur shortly after diabetes diagnosis. Since early treatment of diabetes is essential for preventing disability and death, it is important for patients to understand diabetes and its related complications. Knowledge can enhance self-management behaviors, ultimately improving outcomes among patients with diabetes [5]. However, little is known about patients' knowledge of diabetes sequalae, particularly microvascular complications. Health care providers can educate patients and recommend screening and treatment options for microvascular complications, helping patients with their medical care choices [6]. Yet, up to one-third of physicians do not recognize the signs of diabetic peripheral neuropathy, even in symptomatic patients [7]. Thus, characterizing health care provider knowledge of diabetic microvascular complications, particularly in low- and middle-income countries where diabetes prevalence is increasing most rapidly, can identify changes needed to health care systems to improve patient outcomes.

The aim of this literature review is to summarize knowledge of diabetes and diabetic microvascular complications among patients with diabetes and health care providers. This includes recent studies examining predictors of improved patient and provider knowledge as well as predictors of improved diabetic microvascular screening practices. Our goal, also, is to identify gaps in patient and provider knowledge to facilitate further studies of the relationship between patient and provider knowledge of diabetic microvascular complications.

2. Methods

We searched three electronic databases, PubMed, Cochrane, and CINAHL from 14 July 2021 to 20 July 2021 for articles published from the date of database inception until the end of

search period, 20 July 2021. The key terms and synonyms used alone or in combination were: "patient", "caregiver", "individual", "provider", "doctor", "health care worker", "understanding", "knowledge", "diabetes", "neuropathy", "retinopathy", "nephropathy", and "kidney disease". Additional searches were conducted by scanning the references lists and citations of included articles to ensure all relevant studies were identified. Only peer-reviewed articles that investigated patient or health care provider knowledge of diabetic neuropathy, retinopathy, or nephropathy were included. Studies were excluded if the patient population did not have diabetes (*i.e.*, community-wide sample), unless results from patients with diabetes could be separated using the data provided.

The search identified 13,128 potentially eligible studies once duplicates were removed. After screening based on title, abstract, and keywords, the eligibility of 358 full text articles was assessed, of which 102 were excluded. An additional 28 articles were identified by screening reference lists, resulting in a final literature sample of 284, as shown in PRISMA diagram (**Figure 1**).

3. Patient knowledge of diabetes

Diabetes knowledge is generally assessed via multiple choice surveys, either self-administered or interviewer-administered, covering topics ranging from metabolic facts, hypoglycemia and hyperglycemia symptoms, medication usage, diet and exercise, among others [8]. Studies suggest that diabetes knowledge varies widely between study populations. In an early Scottish study, only 27/182 patients (15%) answered 11 questions correctly that investigators deemed essential diabetes knowledge [8]. Participants generally scored better on medication management questions than general metabolic and diabetes facts. The opposite was shown among patients with diabetes in Mexico who had a better knowledge of diabetes concepts than blood glucose self-monitoring and diabetes-related medication [9]. Diabetes knowledge was markedly better in a recent US study of 17 patients, with an average knowledge score of 60% [10]. Of note, however, the study was underpowered to assess predictors of diabetes knowledge.

A good body of evidence suggests that improved patient knowledge of diabetes is associated with better self-management of disease. This has been demonstrated using overall glucose control [11-13] and medication adherence [14] as surrogates of diabetes self-management. In a US study of 44 patients, those with better scores on a diabetes medication knowledge questionnaire had significantly lower HbA1c (p<0.0001) [12]. This relationship between diabetes knowledge and diabetes control may also be upheld in select populations, such as patients with diabetes on dialysis [13]. Diabetes knowledge also aids in stricter adherence to medication, which is critical for disease management [14]. Overall, indications are that improved patient knowledge of diabetes can translate to better glycemic control and medication adherence, which likely reduce disease progression. Therefore, raising awareness and knowledge of diabetes could exert a meaningful and beneficial impact on disease self-management.

To identify patients with diabetes that would benefit from diabetes education, and hence improved disease control, it is essential to identify determinants of low diabetes knowledge. There are several parameters that potentially influence patients' knowledge of diabetes. Age is a well- and long-established predictor of diabetes knowledge. This relationship between greater

diabetes knowledge and younger age has been noted in various populations, including in Mexico [9], Costa Rica [15], Kuwait [16], United Arab Emirates [17], and Singapore [18].

Additional clinical and/or demographic variables are linked to diabetes knowledge. Longer diabetes duration is implicated in better knowledge scores by some studies [17, 19], but not by others [16, 20], possibly due to confounding parameters. There is also discordance in studies investigating the impact of sex; diabetes knowledge was greater in females [21] or males [17, 22-25] depending on the study, and one found no difference [26]. However, overall, studies failed to adjust for education level and other confounding variables, which may have varied in these countries. Race/ethnicity is a possible contributor; studies to date suggest that patients who self-identify as White are likelier to have broader diabetes knowledge compared to some minorities [20, 27-30]. While this may indicate areas to improve through outreach in these populations, it is important to emphasize that most of these studies do not adjust for confounding, particularly by socioeconomic status, which limits interpretation of this association. Further, some studies categorized participants by nationality, rather than by ethnicity [16, 17]. Overall, there is a need for better quality studies that adjust for confounding, to better identify predictive factors affecting diabetes knowledge to launch education campaigns likely to benefit those patients at highest risk.

As might be anticipated, formal education [9, 15-17, 19, 20, 23, 31], income [19], and health literacy [32, 33] influence diabetes knowledge. Importantly, delivering focused diabetes education can raise diabetes knowledge [11], paving a way forward for patients with lower knowledge on a trajectory for better disease control.

4. Knowledge of diabetic complications

Uncontrolled diabetes, *i.e.*, elevated HbA1c and fasting blood glucose, is a significant risk factor for developing microvascular complications, including retinopathy, nephropathy [34], and neuropathy [3]. Several additional risks factors, such as obesity and dyslipidemia [35-37], which are frequent comorbidities in patients with diabetes, also raise the risk of microvascular complications, especially in patients with T2D. Therefore, lack of knowledge of diabetes leading to suboptimal self-care and poor disease control could potentially also increase the risk of diabetic complications. Indeed, just as patient knowledge of diabetes is generally suboptimal, recognition of diabetic complications, including microvascular complications, was less than 50% in some settings [23, 38].

In addition, and compounding the issue, providers do not universally address diabetic microvascular complications in patients [7, 39-42], even in patients clearly exhibiting symptoms e.g., neuropathy [7]. Since provider recommendations significantly influence patient care choices in diabetes microvascular complications [6, 43], it is important to understand the level of provider knowledge of microvascular complications. However, just as for patients, there is no comprehensive review of provider knowledge and predictors of provider knowledge of these diabetic complications. Herein, we provide our findings from a thorough literature review of both patient and provider knowledge, covering retinopathy, neuropathy, and nephropathy.

4.1 Retinopathy

Diabetic retinopathy is a leading global cause of preventable vision impairment and blindness. The damaging effects of diabetes on the eye can be prevented by early detection of retinopathy through screening and timely treatment of sight-threatening complications. Therefore, it is critical to identify approaches that improve patient knowledge leading to better health choices and care seeking behavior. We found literature regarding patient knowledge of retinopathy from many world regions, although the preponderance of data was from Saudi Arabia and India [40] with some studies from the US [44]. They highlight critical gaps in patient knowledge. Even in countries with provider guidelines, for instance by the American Diabetes Association (ADA) [45], to ensure retinopathy is evaluated in timely manner, patient adherence to screening is suboptimal [43], of which lack of knowledge of the potential harms of retinopathy may be a contributing cause. In a nationwide US study of 204,073 patients with diabetes, only 71.1% adhered to the retinal screening recommendations during a median 4.8-year follow-up [46]. Therefore, it is important to characterize the determinants leading to poor adherence, such as knowledge of retinopathy.

4.1.1 Instruments to assess retinopathy knowledge

Several formal Knowledge, Attitudes, and Practices (KAP) surveys have been developed to assess patient knowledge of retinopathy. A KAP survey is a quantitative or qualitative method to address a predefined question, what are patient and provider understanding of diabetic complications in this instance, through a standardized questionnaire. The goal of KAP surveys is to reveal misconceptions or misunderstandings, which pose an obstacle to desirable activities or behaviors, i.e., screening, self-care, and appropriate management of diabetic complications in this review. KAP surveys can be structured, e.g., multiple choice, guided questions, or semistructured, i.e., relatively more open-ended interviews. The KAP survey may be conducted by the investigator, either in-person or over the phone, or may be self-administered, online or by mail. To assess whether the KAP surveys will evaluate the intended topic, they are tested for internal consistency, reliability, and face validity and frequently pretested in a pilot group characteristic of the target population. Cronbach's alpha coefficient is a measure of test reliability or internal consistency for a set of scales or test items [60]. The coefficient ranges from 0 to 1; the coefficient is 0 when test items are independent from one another, but approaches 1 when test items have high covariances, i.e., they measure the same underlying concept, KAP in these instances. A Cronbach's alpha coefficient minimum of 0.65 and 0.8 is recommended.

KAP surveys for retinopathy span structured and semi-structured self- and investigator-administered questionnaires [47-55]. The internal consistency of some KAP surveys has been assessed and found to be acceptable with Cronbach's alpha coefficients ranging from 0.6 to 0.8 [56-59]. Some studies have evaluated face validity by consulting an expert panel [61, 62], whereas other have developed questionnaires based on reviews of the literature [62, 63]. Pretesting by leveraging a pilot group of volunteers outside of the study area or study cohort is a relatively widely adopted validation method [55, 58, 62]. Unfortunately, little information using the same retinopathy KAP survey in multiple settings or in multi-center studies is available therefore it is unclear how these measures perform across different patient populations [43].

Far less literature has been published regarding KAP surveys of retinopathy knowledge among providers. A search of the literature yields both self- [64-66] and investigator- [40, 67]

administered structured questionnaires, all pretested in separate participant groups. One study evaluated KAP survey internal consistency with a moderate Cronbach's alpha coefficient (0.64) [65]. More commonly, informal surveys of provider knowledge of retinopathy using short one- or two-question queries are documented in the literature [68-70]. A couple of reports lack details regarding the instrument used [71, 72]. Thus, overall, there are few KAP survey instruments developed to assess provider knowledge of retinopathy.

4.1.2 Patient knowledge of retinopathy

KAP surveys of patient knowledge of diabetic complications suggest that retinopathy is the most recognized complication [31, 49, 63, 73]. Among Irish patients with diabetes, 92% of those with T1D and 83% with T2D knew retinopathy was a diabetes-related complication, compared to 71% of T1D and 53% of T2D for neuropathy [73]. Despite this, some populations, such as American Indians and Alaskan Natives, were unaware of the connection between retinopathy and diabetes [44] whereas other populations held misconceptions regarding causes, e.g., watching too much TV [6] or bad luck [52] (Table 1). There is also a limited understanding by patients that retinopathy can be asymptomatic [48, 55, 74-76], including in select populations, such a 50% of females surveyed in New York City [77]. This belief may hold patients back from attending screening if they are unaware that they may be in the early asymptomatic stages of retinopathy. Indeed, in this survey of 150 low-income diabetic females from New York City, a fifth were unfamiliar with the type of provider required for an eye exam and 17% were unaware that annual eye exams were recommended [77]. Of those aware of the annual screening recommendation, only approximately a quarter had knowledge regarding the need for dilation of the pupils as a critical component of the eye exam. In a survey of patients with diabetes in rural India, less than a third were aware that eyes must be assessed on a regular basis [61].

In addition to knowledge gaps concerning causes and screening for retinopathy, there is a lack of knowledge regarding treatment options. Early surveys of urban populations suggest a significant proportion of patients with diabetes are unfamiliar with treatment options, with only around a fifth of respondents demonstrating correct knowledge [6]. This proportion was even lower, 5%, in a recent rural study in India [61]. It is unclear from our literature review whether knowledge has improved in urban populations in recent years. Furthermore, there is a low level of knowledge regarding the preventative, rather than curative, nature of treatments, such as laser and glucose control, which are erroneously thought to be curative [76, 78].

Previous studies also examined predictors of retinopathy knowledge, spanning demographic, clinical, and socioeconomic factors. Studies are discordant regarding age, with findings of greater knowledge in younger [69, 79] versus older [47, 61] patients with diabetes. In surveys that examined sex, being female was associated with greater knowledge of retinopathy [50]. In addition, patients with a longer duration of disease were more likely to understand diabetic retinopathy, and this was confirmed in multiple recent surveys [48, 56, 58, 69, 80]. Further, patients with a prior eye exam also had a greater knowledge of retinopathy [79]. Examination of socioeconomic factors reveals some anticipated correlations with greater retinopathy knowledge, such as higher formal education [48, 56, 59, 61, 69, 77, 81-83], literacy [63], urban residency [58, 81], and income [58, 69, 83]. Additionally, speaking English, were English was not the persons first language, was also linked to greater knowledge of retinopathy [59, 79]. Only ten studies controlled for confounding. Among those that did, the following

predictors of retinopathy knowledge remained significant: younger age [79], duration of diabetes [58, 84], a prior eye exam [79, 84-86], higher income [58], urban residency [58, 79], and greater formal education [61]. Six studies found no significant demographic or clinical predictors of increased retinopathy after controlling for confounding [47-49, 70, 74, 87].

Moreover, surveys have examined predictors of engaging in retinopathy screening behavior, which were multifactorial. From the literature we identified the main determinants for attending screening were younger age, female sex, and White ethnicity [88]. Diabetes characteristics also play a role in whether patients seek screening, such as more severe diabetes and comorbidities such as hypertension and hyperlipidemia, [85] and, overwhelmingly, longer disease duration [48, 74, 85, 89], although one study noted the opposite [90]. As might be expected, previous attendance for an eye exam [87], receiving a physician recommendation to attend for screening [43, 62, 75, 76, 85], and better knowledge of diabetes [53, 85, 91] and retinopathy [43, 51, 53, 57, 91, 92] also increase screening adherence. Lastly, as for determinants of retinopathy knowledge, greater formal education [48, 89, 90], urban residency [88], higher income [92], and linkage into care/health insurance [92] was associated with an increased likelihood of screening attendance. Again, few studies adjusted for potential confounding.

Overall, our literature review identifies crucial gaps in patients' diabetic retinopathy knowledge. These span lack of awareness of the relationship of retinopathy to diabetes, occurrence of early asymptomatic disease, misconceptions regarding causes, and paucity of knowledge regarding therapies. Surveys have revealed several determinants of low retinopathy knowledge and screening, such as demographics (age, sex, ethnicity), diabetes duration, prior behavior, physician recommendations, and various socioeconomic factors. This could help identify patient populations, which would benefit from retinopathy education and outreach. Of note, several studies identified from urban, Western countries were conducted over two decades ago, and we could not ascertain whether patient KAP have since changed in the intervening years.

4.1.3 Provider knowledge of retinopathy

Provider knowledge of retinopathy is crucial for ensuring patients' optimal eye care because multiple studies support that physician recommendations are strong determinants of patients' adherence to screening guidelines [43, 62, 75, 76, 85]. Sixty to 100 percent of physicians [39, 66] and 50%-75% of nurses and midlevel providers [67, 93] know diabetes can adversely affect the eyes. However, overall retinopathy knowledge can be poor among providers in some geographic areas. A survey of private sector non-ophthalmic providers (n=355) in Saudi Arabia found a good level of diabetic retinopathy knowledge was only present in 54.3% of interviewees, along with a positive attitude among 31.3% and excellent practice among only 40.8% of interviewees [40]. We did not identify any studies that compared provider knowledge globally. We did, however, find evidence that ophthalmic specialists outperform non-specialists for detecting proliferative retinopathy from seven-view stereo fundus photographs and review of medical charts [94]; therefore, suboptimal retinopathy knowledge may be more of an issue among general doctors than eye experts. Since most patients receive their medical care first from their primary care physician, they must be knowledgeable of retinopathy to determine when a referral to an ophthalmologist is necessary.

Our search of the literature identified several points of provider knowledge limitations. These included a lack of awareness concerning what part of the eye diabetes affects [95], uncertainty regarding the tests used to diagnose retinopathy [64, 65, 72], as well as misconceptions regarding contraindications to diabetic fundoscopic exams, e.g., hypertension [96, 97]. As we had identified for patients, a small KAP survey also found lack of knowledge among providers regarding the existence of asymptomatic disease. This was noted by a small study of physicians (n=22) and village health workers (n=25) in rural China, which found most providers did not conduct a pupil dilation exam if the patient had no symptoms [97]. Similarly, a KAP survey of primary care physicians in Saudi Arabia (n=216) found that only 46% were aware that patients initially exhibit no symptoms in the early stages of retinopathy [64]. An early investigation of KAP among Canadian general practitioners (n=1,038) found that 27% overestimated the benefits of treatment, i.e., a false belief that laser photocoagulation improved rather than stabilize disease progression [66]. Lastly, we identified provider gaps in knowledge regarding gestational diabetes. Family-practice physicians (n=224) were more likely to examine the eyes of patients with gestational diabetes for retinopathy compared to obstetrics/gynecology physicians (n=184), as surveyed by mail [98, 99]. In

We found only scant information regarding predictors of greater provider knowledge of retinopathy. As might be anticipated, specialist training correlates with greater knowledge or ability to detect retinopathy, e.g., retinal specialists versus internists, diabetologists, and medical residents [94] or additional subspeciality training [64]. Longer duration of practice was also a determinant of greater knowledge [64]. Patient characteristics also contributed, with providers demonstrating better knowledge regarding the connection of retinopathy and T2D versus T1D and, as a result, providers more frequently referred patients with T2D versus T1D to ophthalmology [101]. In a KAP survey of medical students in Saudi Arabia, males scored higher on knowledge and practice whereas females scored better on attitude [102].

Cumulatively, our search of the literature revealed some investigation of provider KAP, although recent studies were limited in scope and geographic location. Moreover, carefully adjusted studies for confounding factors are scarce.

4.2 Neuropathy

Peripheral neuropathy is an injury of the nerves, generally in a symmetric distal to proximal fashion, initiating in the feet and progressing to the calves [3]. In the later stages, the hands may also be affected. Neuropathy can impair gait and stability, increasing susceptibility to falls and secondary injury. Moreover, peripheral neuropathy can lead to non-healing foot ulcers, which may ultimately require lower limb amputation. Thus, it can significantly increase disability and lower quality of life, making it essential for patients to understand neuropathy. We searched the literature for studies that examined patient knowledge of neuropathy. Most studies were conducted in India and China, although studies were conducted across multiple other countries [103-105]. Patient populations comprised both inpatients with diabetic ulcers as well as outpatients with diabetes lacking neuropathy symptoms. One 2000 US study of patients who were ADA members in an urban setting found that 27% of respondents reported they had not been advised or educated on diabetic neuropathy and foot complication by their health care provider [103]. Thus, gaps in patient knowledge of neuropathy may be substantial, even in patients belonging to an organization advocating and supporting diabetes research.

4.2.1 Instruments to assess neuropathy knowledge

We identified several instruments assessing neuropathy knowledge in the literature. The majority were KAP surveys focused on foot care and foot ulcer knowledge and practice, rather than neuropathy more broadly. KAP surveys were both in structured [106-108] and interview format questionnaires [109], either self- or investigator-administered [110]. In one study, the questionnaire was investigator-administered when the respondent was illiterate or physically unable to complete the survey but self-administered by the remainder of participants [111]. A few administered KAP surveys were adapted from prior surveys [112, 113], whereas a few were utilized in multiple studies [106, 114] or used prior instruments [107, 115]. Moreover, we found a KAP instrument that split the survey into basic and extended foot care practices [116]. The Patient Interpretation of Neuropathy (PIN) questionnaire evaluated both misperceptions about foot complications, patient knowledge of neuropathy and its link to complications, and foot self-care efficacy beliefs, among other concepts related to patient understanding of neuropathy [117].

A few studies evaluated parameters of KAP surveys for capturing patient knowledge. Several KAP surveys we identified were pretested [106, 107, 109, 113, 118], although one study was pretested in medical students instead of a population meeting the criteria of the study population [110]. Regarding, internal consistency of KAP surveys, they had Cronbach's alpha coefficients ranging from 0.72 to 0.86, which is rated as acceptable [116, 119, 120]. A couple of studies assessed face validity of the utilized KAP survey by a panel of medical experts [108, 109]. In addition to KAP surveys, we also found papers that leveraged scoring and/or scaling instruments to assess patient knowledge of diabetic foot care. These included diabetic knowledge [121] and foot care scores [104, 114, 121-125]. Finally, one study report provided no information regarding the employed instrument [126].

We found far fewer neuropathy KAP instruments for providers; however, they spanned structured and semi-structured questionnaires, which were self- [127-129] or investigator- [39, 93] administered. We noted some surveys were pretested [39], were assessed for face validity by experts, and evaluated for internal consistency by Cronbach's alpha coefficients (0.72 for junior doctors, 0.81 for nurses) [128]. Additionally, studies used previously validated instruments about KAP towards diabetes more broadly, *e.g.*, Diabetes Self-Report Tool, Diabetes Basic Knowledge Tool [42, 129, 130].

4.2.2 Patient knowledge of neuropathy

Overall, evidence suggests patient knowledge of neuropathy ranges from 10-60% compared to 60-92% for retinopathy [73, 131], which may also be the case in providers, e.g., nurses [93]. Of the papers we assessed, we found a broad range of patient level knowledge and practice behaviors in neuropathy, i.e., diabetic foot (Table 2). Many reported less than adequate foot care behavior in diverse populations worldwide, urban and rural [105, 111, 123]. Moreover, some studies highlighted a disconnect in knowledge and practice. In a Saudi Arabian study of patients with T2D (n=360), although 70% had knowledge of diabetic foot care, only 41.7% examined their feet, 41.4% washed them with warm water, 31.4% carefully dried them between the toes, and 33.1% used moisturizer [132]. We also noted some misconceptions regarding foot

care; for example, qualitative interviews with people with diabetes in Jordan revealed the belief that there is no need to examine the feet if participants had no ulcers [133]. Appropriate education on diabetic neuropathy can have tangible effects on care adherence. A study of T2D patients with diabetic neuropathy (n=104) found that foot care education enhanced attendance at yearly check-ups, as well as moisturizer use and appropriate shoe wear (all p<0.05) [134]. Another study in Saudi Arabia similarly found that foot care practice was superior in T2D patients that received physician recommendations to examine their feet [132]. Therefore, it is essential for patients to understand neuropathy to adopt practices that improve foot care.

Across the studies, we identified multiple predictors of patient knowledge, which included demographic, clinical, and socioeconomic factors. The literature findings regarding sex were mixed. We found reports that found neuropathy knowledge was greater in females [114] and, conversely, in males [118], and one study that did not find a relationship between neuropathy knowledge and sex [135]. Older age also associated with deeper knowledge of diabetic foot and neuropathy [109, 115]. Additionally, in a large Chinese study of patients with T2D (n=5,961), disease characteristics had an influence on patient knowledge, including positive correlations with diabetes duration and regular diabetes care following multiple regression analysis [115]. Prior foot complications may impact neuropathy knowledge. In a Thai study, knowledge was lower among T2D patients with (n=55) versus without ulcers (n=110), which did not correlate with either foot care score or diabetes duration [121]. Conversely, a UK study of amputees at a foot clinic found a high level of foot care knowledge, which did not differ between patients with unilateral (n=121) or bilateral (n=22) amputations [122]. No differences were noted in KAP between patients with (n=89) versus without (n=121) amputation in a St. Kitts and Nevis [136]. We also identified that prior education on foot care [114, 115] and prior physician advice [132] was associated with greater neuropathy knowledge. As might be anticipated, multiple studies also found that higher levels of formal education enhanced knowledge of diabetic foot disease [106-109, 115, 126]. Lastly, higher socioeconomic status was linked to a greater knowledge [107, 126].

Although knowledge scores can correlate with practice behavior [137], as noted above, better neuropathy knowledge does not always lead to better foot care [132]. Thus, we also combed the literature for determinants of good foot self-care. Female sex was associated with greater foot self-care (p<0.035) [138], an association that persists after controlling for confounders in the US, China, and Ethiopia [115, 138, 139]. Younger age is also a predictor of better care [104, 140], though one study noted no effect of age based on a 50-year-old cutoff [107]. Studies that identified predictors of greater knowledge through multiple regression analysis indicated a weak association of age with knowledge in a US study [104], but a strong association in other studies [115, 139]. A few studies examined the influence of ethnicity; analysis of T2D US participants from the Diabetes Attitudes, Wishes and Needs 2 study found that Black Americans spent more time on foot self-exam per week versus White or Chinese Americans, after controlling for income, age, education and diabetes type (p<0.05) [141]. A US study of veterans with diabetes confirmed this association, with higher adherence to foot care among Black Americans as well as Hispanic patients when compared to White patients, in multiple regression analysis [104], as did a UK study across the general T2D population [30].

As with neuropathy knowledge, previous experiences with foot ulcers or amputations may also influence self-care practice [142]. In the study of US veterans, neuropathy symptoms, a foot ulcer in the previous year, or a prior amputation independently predicted more meticulous

foot care [104]. Several studies highlighted that better diabetic foot education [104, 134, 138] and attention from a foot care professional [104, 138] also improved self-care practices. Finally, socioeconomic forces played a role in self-care behavior, including a higher formal education [107], urban residency [79, 137, 139], and income [142]. Factors may modify these relationships. For instance, in the US veteran study, years of schooling did not remain significant after multiple regression analysis [104], although it did in a Chinese study [115]. This emphasizes the importance of correcting for confounding factors in KAP surveys, as well the presence of additional potential contributors that may explain study differences.

4.2.3 Provider knowledge of neuropathy

Our comprehensive literature review found there were far fewer KAP studies of provider neuropathy knowledge versus of patients, and some studies were relatively small. Overall, the studies revealed significant knowledge gaps and misconceptions. A nationwide US study of health care professionals, which comprised general doctors (n=250), specialists (n=150), and nurses and/or physician assistants (n=100), found 53% of survey participants held the belief that adequate glucose control could reverse peripheral neuropathy [127], despite the progressive nature of the disease and the presence of additional risk factors, such as central obesity [37]. Encouragingly, however, over half of providers expressed a desire for more information regarding several aspects of neuropathy, including its cause and how it induces pain or numbness. In a UK study, junior doctors and nurses scored poorly with regards to foot care, although they scored well on general diabetes knowledge [128], indicating a potential disconnect in understanding the link between diabetes and neuropathy. Moreover, neuropathy was the least recognized diabetes complication out of several micro- and macrovascular complications, including retinopathy and nephropathy, by nurses in Australia [93]. Conversely, a study of nurses in Saudi Arabia found neuropathy was recognized 76% of participants, and most by nurses belonging to critical care units [42].

Our survey of the literature also uncovered a few trends in practice. A small 1997 study in Cape Town found that doctors and primary health care nurses (n=22) did not usually assess for peripheral neuropathy (insensate foot, ulcers), unless the patient voiced a complaint [39]. A more recent KAP survey of pharmacists in Qatar found most counselled patients on foot exams and screening for neuropathic pain [129]. In a study that showed footage of a "patient" displaying signs of emerging peripheral neuropathy, only 42.2% of participating US primary care physicians (n=192) indicated they would perform all essential components of a foot examination, whereas 21.9% stated they would perform none [7]. Additionally, providers were more likely to recommend all parts of a foot exam in male versus female, older versus younger, higher versus lower socioeconomic status patients, and in patients with signs of neuropathy compared to those without signs of neuropathy. We could not ascertain more current practices overall due to a lack of studies.

4.3 Nephropathy

Objectively, nephropathy, otherwise known as diabetic kidney disease (DKD), may be considered the most serious of diabetic microvascular complications. DKD is the progressive loss of kidney function secondary to diabetes, which manifests as microalbuminuria and renal

inflammation [143]. In very advanced disease, the so-called end-stage renal disease, it can require renal replacement therapy, and, in the cases of failure, lead to death. Therefore, it is essential for patients with diabetes to be aware of DKD and take measures to prevent onset and/or slow progression. Unfortunately, our literature search did not yield many studies of nephropathy KAP, either among patients or providers. Therefore, this is a significant knowledge gap that requires addressing.

4.3.1 Instruments to assess nephropathy knowledge

Since we identified a few KAP studies of nephropathy in patients with diabetes in the literature, there were only a few instruments, some of which had been previously used to assess knowledge regarding kidney disease not necessarily related to diabetes [144, 145]. Only one mentioned face and content validity and internal consistency [144]. The scenario was similar for provider KAP instruments; some noted pretesting and assessment of internal consistency [146] and employed a previously published tool [147], but there was little data available overall. The lack of validated instruments hinders our ability to accurately assess KAP related to DKD and compare across populations.

4.3.2 Patient knowledge of nephropathy

Among the sparse studies we found, evidence suggests patient knowledge of nephropathy is less than that of retinopathy [73]. Moreover, there was a lack of studies regarding KAP in patients with diabetes and none adjusted for potential confounding (Table 3). A study in Malaysia of patients diagnosed with diseases at risk of chronic kidney disease (diabetes, hypertension, heart diseases, obesity, n=103), nephropathy knowledge was associated with being male, younger, formally educated, married, and of higher income (all p<0.05), although none except marital status remained significant for practice behavior [148]. However, it was unclear from that study what the level of nephropathy knowledge and practice was specifically among patients with diabetes. A Fiji study of T2D patients with chronic kidney disease (n=225) found KAP to be relatively good among participants, with high knowledge, attitude, and practice scores in 61.8%, 63.6%, and 88.4% survey respondents, respectively [144]. It is possible KAP scores were high due to the selected nature of participants, involving those with known T2D and nephropathy recruited from study site providing care for these specific conditions. In fact, patient KAP overall may be low. A study in India of only T2D patients (n=323) found nephropathy knowledge was poor in 79% of survey respondents, and was associated with poor literacy, low socioeconomic status, and limited family income [145]. An Ethiopian study of patients with diabetes and hypertension (n=208) found nephropathy knowledge to be low in 63.5% of participants [149]. Finally, one Australian study of patients with T1D (9%) and T2D (88%) and chronic kidney disease (n=316) investigated the barriers to seeking appropriate care, which included inadequate knowledge of diabetes and nephropathy [150].

4.3.3 Provider knowledge of nephropathy

Nephropathy is relatively well recognized as a diabetes complication by Australian nurses (75% of survey participants), nearing their knowledge of retinopathy (89%) and far outpacing

that of neuropathy (48%) and foot ulcers (43%) [93]. A KAP survey of Ethiopian health care professionals (n=326) indicated 91% were aware of the association of diabetes and hypertension with chronic kidney disease, although there were some gaps, such as only 59% were aware that assessment of enhanced glomerular filtration rate was superior to serum creatinine alone for assessing nephropathy severity [151]. However, the KAP instrument used by this study only had a Cronbach's alpha coefficient of >0.62; therefore, it may not have accurately captured KAP. The association between diabetes and nephropathy was also recognized by 88% of general practitioners in a Pakistani study [147]. Overall, there is a lack of studies investigating DKD KAP in providers and only one adjusted for potential confounders using logistic regression [145].

With regards to DKD practices, in a small Cape Town study, 82% of doctors and primary health care nurses assessed nephropathy by urine protein, whereas only 27% assessed serum creatinine [39]. Moreover, providers were unaware that controlling hypertensive nephropathy is essential for reducing the risk of DKD. A 1999 US study of primary care physicians recruited from the American Medical Association database (n=211) found nearly 98% of physicians assessed proteinuria and microalbuminuria at least as frequently or more frequently as the recommended guidelines at the time of the study, yet 39% chose an inappropriate test for monitoring [146]. In addition, an Australian study found patients with DKD did not always receive the guideline recommended care, with nearly 40% of patients with a blood pressure >140/90 mmHg despite strict blood pressure recommendations among patients with DKD [150]. However, further investigation into the reasons for this, including patient compliance with physician recommendations, was not provided.

5. Conclusions

Microvascular complications contribute to substantial morbidity and mortality among patients with diabetes, yet our comprehensive literature review found that studies examining patient and health care provider knowledge of these complications varies widely between microvascular complication and settings. Retinopathy had the largest number of studies and appears to be the most widely studied diabetic microvascular complication, yet nephropathy, which is a significant driver of diabetes-related mortality, is the topic of substantially fewer studies. Addressing this knowledge gap is essential to reduce mortality among patients with diabetes.

The current literature does offer insight into possible interventions for this patient population. Our literature review found that patients and providers often did not see the need to seek healthcare or screen for microvascular complications unless there are symptoms clearly consistent with diabetic sequalae [7, 64, 74]. This is a clear missed opportunity to reduce morbidity among patients with diabetes. As patients frequently cite health care providers as sources of information [85, 152], improving provider knowledge of diabetic microvascular complications and addressing barriers to patient education and risk factor modification may provide one avenue for improving patient outcomes.

Yet, it is challenging to know whether these interventions would be effective as our literature review noted significant discordance in findings across studies. This could be due to the relatively small sample sizes of most studies, lack of adjusting for potential confounding parameters in most cases, or sociocultural differences between study sites. While there were studies that recruited from multiple cities or from organizational or national databases [114, 115, 153, 154], we did not identify any studies that recruited from more than one country. Therefore, it is difficult to draw firm conclusions whether differences in patient and provider knowledge are

setting specific or related to methodology. Instruments used to assess patient and provider knowledge of microvascular complications varied substantially between studies and, while some authors included detailed regarding instrument validation, this was only moderately implemented. Consistent use of well-validated measures between study sites would address concerns regarding methodology as well as begin to highlight sociocultural differences between settings. Lastly, studies conducted in high-income settings have not been updated and there are few studies that have included the same study site over time. Therefore, we were unable to draw conclusions regarding temporal changes in patient and provider knowledge. To move forward and identify patient and provider populations that would benefit most from educational programs, updated studies using well-validated KAP surveys with results analyzed using more refined statistical tools are essential.

Author contributions Melissa A. Elafros and Brian C. Callaghan contributed to conceptualization and the literature review. Melissa A. Elafros and Eva L. Feldman wrote the original draft. All authors contributed to writing, review, and editing. All authors have reviewed and approved the submitted draft.

Table 1. Summary of studies that investigated retinopathy knowledge among patients with diabetes and health care providers

Studies arranged alphabetically. Abbreviations: CI, confidence interval; GP general practitioner; KAP knowledge attitudes and practices; OR, odds ratio; T1D, type 1 diabetes; T2D, type 2 diabetes.

Study	Country	Study Type & Population	Measure	Main Findings		
Patients with Diabe	Patients with Diabetes					
Addoor 2011 [83]	Malaysia	Cross-sectional 351 from 1 ophthalmology clinic	Study-created KAP survey	87% aware diabetes affects eye. Predictors of knowledge: duration of diabetes (p<0.01), eye exam in last 6 months (p<0.04)		
Adriono 2011 [84]	Indonesia	Cross-sectional 196 from 3 primary care clinics	Study-created KAP survey	38% aware diabetes causes blindness. Prior exam linked to better knowledge (p=0.002).		
Ahmed 2017 [51]	Bangladesh	Cross-sectional 122 from 1 diabetes clinic	Study-created KAP survey	24% with poor knowledge about the effect of diabetes on the eye.		
Al-Asbali 2020 [79]	Saudi Arabia	Cross-sectional 200 from 1 endocrine and 1 ophthalmology clinic	Study-created KAP survey	45% excellent knowledge. Predictors of knowledge: duration diabetes (p=0.03).		
AlHargan 2019 [82]	Saudi Arabia	Cross-sectional 280 from 2 primary care clinics	Adapted KAP survey [149, 151]	88% knew diabetes affects the retina. Predictors of knowledge: formal education (p<0.01), higher income (p<0.05).		
Almalki 2018 [56]	Saudi Arabia	Cross-sectional 253 T2D from 1 endocrinology clinic	KAP survey adapted from prior study [83]	64% knew diabetes affects the eye.		

Alsaidan 2019 [70]	Saudi Arabia	Cross-sectional 174 T2D from 1 primary care clinic	Details not provided	82% aware diabetes affects eye. Predictors of knowledge: male gender (p=0.045), well controlled T2D (p=0.021).
Alwazae 2019 [57]	Saudi Arabia	Cross-sectional 404 from 4 clinics	Study-created KAP survey	73.5% with adequate knowledge.
Al-Yahya 2020 [152]	Saudi Arabia	Cross-sectional 313 from 52 primary care clinics	Validated KAP survey [53]	53% knew diabetes affects the eye. Predictors of knowledge: higher income (p<0.02).
Alzahrani 2018 [61]	Saudi Arabia	Cross-sectional 377 from 38 primary care clinics	Study-created KAP survey	82% knew diabetes affects the eye.
Al Zarea 2016 [151]	Saudi Arabia	Cross-sectional 439 from 5 clinics	Study-created KAP survey	75% aware diabetes can cause eye disease.
Assem 2020 [58]	Ethiopia	Cross-sectional 230 from 1 diabetes clinic	Study-created KAP survey	52% with poor knowledge. Predictors of knowledge: urban residence (p<0.05), income, diabetes (p<0.05), duration (p<0.01),
Bakkar 2017 [149]	Jordan	Cross-sectional 237 T2D randomly selected from 3 cities	Study-created KAP survey	88% aware diabetes can affect the eyes. Predictors of eye knowledge: more than high school education (p<0.01).
Balasubramanian 2016 [60]	India	Cross-sectional 105 from 1 clinic	Details not provided	76% aware diabetes affects the eye. Predictors of knowledge: education (p<0.05)
Çetin 2013 [48]	Turkey	Cross-sectional 437 seen at 1 ophthalmology and 1 endocrinology clinic	Study-created questionnaire	88% knew diabetes affects eyes. 25% thought eye exams only necessary if having troubled vision or poorly controlled diabetes.
Das 2016 [49]	India	Cross-sectional	Study-created KAP survey	65% knew diabetic retinopathy affects the eyes. 42% disagreed that eyes could be affected, even if blood sugar was

		240 from 1 ophthalmology clinic		controlled. Predictors of eye knowledge: none significant
Duan 2020 [68]	China	Cross-sectional 1972 in 1 community health system	Study-created KAP survey	62% knew diabetes affects eyes. Predictors of knowledge: younger age, male sex, higher education, longer diabetes duration (all p<0.01).
Fallatah 2918 [54]	Saudi Arabia	Cross-sectional 380 from 1 ophthalmology clinic	Study-created KAP survey	92% aware diabetes affects eyes. Predictors of knowledge: formal education (p<0.05), urban residence (p>0.05).
Gillibrand 2000 [67]	UK	Cross-sectional 2,815 community patients not engaged in eye care	One knowledge question	18.3% did not know diabetes affects eyes.
Khandekar 2010 [47]	Oman	Cross-sectional 750 in 1 region	Study-created KAP survey	61% aware diabetes affects eyes.
Konstantinidis 2017 [52]	Switzerland	Cross-sectional 323 recruited from community pharmacies	Study-created questionnaire	96% aware diabetes can cause eye disease. 98% knew good glycemic control could prevent occurrence or deterioration of eyes.
Lian 2018 [74]	Hong Kong	Cross-sectional 2,593 at 2 clinics	Study-created questionnaire	11.5% knew retinopathy could be asymptomatic.
Livingston 1998 [78]	Australia	Cross-sectional 205 urban, 240 rural	Study-created knowledge score	37% aware eye problems can occur. Predictors of increased awareness: younger age: rural OR 2.89 [95% CI 1.36-6.06] urban OR 2.32 [95% CI 1.24-4.22]; eye exam in last 2 years: rural OR 1.89 [95% CI 1.04-3.42] urban OR 2.43 [95% CI 1.29-4.57].
Manu 2018 [75]	India	Cross-sectional 150 T2D from 1 hospital	Details not provided	58% aware diabetes affects the eye. No significant predictors of knowledge.
				

Myanmar	Cross-sectional 480 cared for by surveyed GPs	Study-created questionnaire	80.6% knew diabetes affects eyes. 90.4% agreed patients with diabetes should see an eye specialist.
Tanzania	Cross-sectional 316 at 1 diabetes clinic	One knowledge question	34% knew diabetes can damage eye.
Nigeria	Cross-sectional 225 at 1 endocrinology clinic	Study-created questionnaire	57% knew diabetes can affect eye.
Ghana	Cross-sectional 360 at 1 diabetes clinic	Study-created questionnaire	49% knew diabetes can affect eye. No significant predictors of knowledge.
US	Cross-sectional 150 women from 1 medical center	Study-created knowledge and belief score	17% did not know required frequency of eye exams. 40% knew controlling glucose was important. Formal education linked to greater knowledge.
Pakistan	Cross-sectional 132 from 1 ophthalmology clinic	Details not provided	57% knew diabetes affects the eye. 22% reported eye exams should occur once vision was affected.
India	Cross-sectional 1,000 at 1 clinic	Study-created awareness score	84% aware diabetes can affect the eye. 46.9% knew related to glucose control. 50% thought routine eye exams not necessary.
Australia	Cross-sectional 68 T1D, 187 T2D in Diabetes Australia	Study-created questionnaire	96.2% knew diabetes causes eye problems.
US	Cross-sectional 2,308 in 1 county	Study-created questionnaire	47% knew eye examinations were needed for people with diabetes.
India	Cross-sectional 288 from 1 ophthalmology clinic	Study created questionnaire	58% had poor knowledge.
	Tanzania Nigeria Ghana US Pakistan India Australia US	Myanmar 480 cared for by surveyed GPs Cross-sectional 316 at 1 diabetes clinic Cross-sectional 225 at 1 endocrinology clinic Ghana Cross-sectional 360 at 1 diabetes clinic Cross-sectional 150 women from 1 medical center Cross-sectional 132 from 1 ophthalmology clinic India Cross-sectional 1,000 at 1 clinic Cross-sectional 1,000 at 1 clinic Cross-sectional 2,308 in 1 county Cross-sectional India Cross-sectional 2,308 in 1 county Cross-sectional 1,001 288 from 1	Myanmar480 cared for by surveyed GPsStudy-created questionnaireTanzaniaCross-sectional 316 at 1 diabetes clinicOne knowledge questionNigeriaCross-sectional 225 at 1 endocrinology clinicStudy-created questionnaireGhanaCross-sectional 360 at 1 diabetes clinicStudy-created questionnaireUSCross-sectional 150 women from 1 medical centerStudy-created knowledge and belief scorePakistan132 from 1 ophthalmology clinicDetails not providedIndiaCross-sectional 1,000 at 1 clinicStudy-created awareness scoreAustraliaCross-sectional 68 T1D, 187 T2D in Diabetes AustraliaStudy-created questionnaireUSCross-sectional 2,308 in 1 countyStudy-created questionnaireIndiaCross-sectional 2,308 in 1 countyStudy-created questionnaireIndiaCross-sectional 2,308 from 1Study created questionnaire

Tajunisah 2011 [62]	Malaysia	Cross-sectional 137 from 1 ophthalmology clinic	Details not provided	86% aware diabetes can affect the eye. Predictors of knowledge: formal education (p>0.05).
Vanugopal 2020 [59]	India	Cross-sectional 350 from 1 hospital	Study-created questionnaire	34% had adequate knowledge of diabetic retinopathy. Predictors of knowledge: formal education (p<0.001).
Walker 1997 [6]	US	Cross-sectional 67 Black Americans with diabetes in New York	Study-created questionnaire	87% believed diabetic eye problems were symptomatic. 21% thought there were effective treatments.
Wang 2010 [85]	China	Cross-sectional 53 T1D 836 T2D from 1 endocrine and 1 general clinic	Study-created KAP survey	77% aware diabetes affects eyes. Prior exam linked to better knowledge (p<0.001).
Whiting 1998 [77]	Australia	Cross-sectional 121 patients with retinopathy from 1 ophthalmology clinic	Study-created questionnaire	95% knew diabetes affects the eyes
Zou 2017 [73]	China	Cross-sectional 519 with diabetes in 1 community	Study-created questionnaire	95% aware diabetes affects the eye, 12% aware it can be asymptomatic.
Health Care Provid	ers			
Abdulsalam 2018[64]	Nigeria	Cross-sectional 105 physicians from 4 hospitals	Study-created KAP survey	36% perform eye exams, 90% do not use dilating eye drops
Abu-Amara 2019 [40]	Saudi Arabia	Cross-sectional 182 GPs, 115 internists	Study-created KAP survey	45% with poor knowledge.
Al Rasheed 2017 [63]	Saudi Arabia	Cross-sectional	Study-created questionnaire	Knowledge linked to: family medicine subspeciality training (p<0.01), years of practice (p<0.01).

	142 family, 10 pediatric, 8 internists, 56 GPs		
Saudi Arabia	Cross-sectional 76 GPs in 1 province	Previously used KAP survey [63]	37% performed dilated fundus exams.
Saudi Arabia	Cross-sectional 141 GPs from 63 centers	Study-created questionnaire	56% with good knowledge.
Saudi Arabia	Cross-sectional 73 medical students	Study-created KAP survey	Moderate overall KAP score, linked to male sex (p=0.02). 66% knew correct timing for eye exams.
New Zealand	Cross sectional 287 nurses	Study-created survey	89% identified retinopathy as a diabetes complication. Predictors of knowledge: level of training (p=0.006).
Canada	Cross-sectional 648 GPs, 96 trainees	Study-created questionnaire	Correct timing for screening in T1D: 74% vs T2D: 82%. 33% knew macular edema could be asymptomatic.
US	Cross-sectional 23 optometrists	Study-created survey	Low level of knowledge regarding need for dilated fundus exams.
India	Cross-sectional 36 optometrists, 241 GPs	Study-created questionnaire	<23% optometrists and <33% GPs had acceptable knowledge regarding risk factors and management of diabetic retinopathy.
South Africa	Cross-sectional 12 doctors, 23 nurses	Study-created survey	100% knew diabetes affected the eye.
Oman	Cross-sectional 42 ophthalmologists, 33 mid-levels, 12 GPs	Study-created questionnaire	Acceptable knowledge: 71% ophthalmologists, 54% mid-levels, 33% GPs.
Myanmar	Cross-sectional	Study-created guestionnaire	Correct timing for screening in T1D: 2% vs T2D: 93%.
	Saudi Arabia Saudi Arabia New Zealand Canada US India South Africa Oman	Saudi Arabia Cross-sectional 76 GPs in 1 province Cross-sectional 141 GPs from 63 centers Saudi Arabia Cross-sectional 73 medical students Cross-sectional 287 nurses Canada Cross-sectional 648 GPs, 96 trainees US Cross-sectional 23 optometrists Cross-sectional 36 optometrists, 241 GPs South Africa Cross-sectional 12 doctors, 23 nurses Cross-sectional 42 ophthalmologists, 33 mid-levels, 12 GPs	Saudi Arabia Cross-sectional 76 GPs in 1 province Cross-sectional 141 GPs from 63 centers Saudi Arabia Cross-sectional 141 GPs from 63 centers Saudi Arabia Cross-sectional 73 medical students New Zealand Cross sectional 287 nurses Cross-sectional 648 GPs, 96 trainees Cross-sectional 23 optometrists Cross-sectional 36 optometrists, 241 GPs South Africa Cross-sectional 12 doctors, 23 nurses Cross-sectional 42 ophthalmologists, 33 mid-levels, 12 GPs Study-created questionnaire Study-created questionnaire Study-created questionnaire Study-created questionnaire Study-created questionnaire Study-created questionnaire

		100 GPs		
Namperumalsamy 2004 [156]	India	Cross-sectional 199 paramedical personnel	Study-created questionnaire	88.5% knew diabetes could affect eyes. 20% knew uncontrolled diabetes is a risk factor. 75.9% unaware of treatments for retinopathy.
Raman 2006 [157]	India	Cross-sectional 159 GPs	Study-created questionnaire	54% aware patients with diabetesshould have annual dilated eye exams.
Wright 2001 [158]	Australia	Cohort 310 optometrists	Study-created questionnaire	74.5% perform dilated exams on new patients with known diabetes.
Yan 2012 [96]	China	Focus groups 22 physicians, 25 village health workers	Study-created interview guide	Good overall knowledge, physicians did not dilate pupils to detect asymptomatic disease.

Table 2. Summary of studies that investigated neuropathy knowledge among patients with diabetes and health care providers.

Studies arranged alphabetically. Abbreviations: ADA American Diabetes Association; KAP, Knowledge, Attitudes, and Practices; T2D, type 2 diabetes.

Study	Country	Study Type & Population	Measure	Main Findings
Patients with Diabetes				
Abu-Qamar 2014 [105]	Jordan	Qualitative interviews 7 patients with burn injuries	Study-created interview guide	Participants did not believe they needed regular food exams in the absence of ulcers.
Bohorquez Robles 2017 [115]	Mexico	Cross-sectional 200 T2D from 1 primary care clinic	Foot Care Knowledge and Practice Questionnaire [159]	52% had poor knowledge of foot self-care.
Chellan 2012 [108]	India	Cross-sectional 203 from 1 podiatry clinic	Previously validated KAP survey	Patients with foot ulcers more likely to have poor knowledge (p=0.001).
Corbett 2003 [120]	US	Randomized control trial 40 T2D with home care	Foot Care Knowledge Questionnaire [159]	Moderate baseline foot care knowledge. Educational intervention improved knowledge (p<0.01).
De Sá Pilocarpo 2014 [114]	Brazil	Cross-sectional study 85 T2D from 2 primary care clinics	Previously used KAP questionnaire [160]	49.5% with limited foot care knowledge.
Desalu 2011 [103]	Nigeria	Cross-sectional 352 from 3 tertiary hospitals	Pre-tested questionnaire	46% with poor knowledge of diabetic foot care.
Foolchand 2013 [112]	Mauritius	Qualitative interviews 120 from 5 hospitals	Study-created interview guide	75% unaware of need for annual foot screening.

Hanley 2020 [132]	St. Kitts and Nevis	Cross sectional 210 from multiple health care settings	Adapted KAP questionnaire [108]	Average knowledge reported. No difference in knowledge based on amputation status.
Hasnain 2009 [102]	Pakistan	Cross-sectional 150 from 1 diabetic clinic	Study-created questionnaire	29.3% with good knowledge. Predictors of knowledge: formal education (p<0.01)).
Jain 2012 [107]	India	Cross-sectional 251 from multiple hospitals	Study-created questionnaire	62% had poor foot care knowledge
Jinadasa 2011 [106]	Sri Lanka	Cross-sectional 110 with diabetic foot ulcers	Study-created questionnaire	52.7% with good footcare knowledge.
Khamseh 2007 [104]	Iran	Cross-sectional 148 T2D from 1 diabetes clinic	Study-created questionnaire	Predictors of knowledge: higher formal education (p<0.01).
Lamchahab 2011 [122]	Morocco	Cross-sectional 91 hospitalized patients	Details not provided	85% did not pay attention to "warning signs" of foot injuries. Predictors of knowledge: formal education, socioeconomic status (both P<0.01).
Li 2014 [111]	China	Cross-sectional 5,961 T2D from 144 hospitals	Summary of Diabetes Self-Care Activities	Overall medium level of foot care knowledge. Multivariate predictors of knowledge: female sex, older age, formal education, diabetes duration, regular diabetes care, prior education regarding diabetes complications (all p<0.001).
Muhammad-Lutfi 2014 [131]	Malaysia	Cross-sectional 157 admitted with foot infections.	Previously used questionnaire [102]	58% with poor foot knowledge.

Naicker 2009 [121]	Malaysia	Cross-sectional 100 from 1 hospital	Preventative Measure Scale [161]	Poor overall foot knowledge.
Pollock 2004 [110]	UK	Cross-sectional 365 from a population-based diabetes register	Study-created questionnaire	Moderate overall knowledge. Predictors of knowledge: female gender (p=0.04).
Pourkazemi 2020 [133]	Iran	Cross-sectional 375 T2D from 1 clinic	Study-created questionnaire	15% with good knowledge. Predictors of knowledge: female gender, duration of diabetes, urban residents, formal education, prior diabetic foot ulcer, prior amputation (all p<0.05).
Rheeder 2008 [162]	South Africa	Cross-sectional 120 from 1 diabetes clinic	Modified questionnaire [110]	Participants with ulcer at-risk feet were less likely to inspect their feet daily (p=0.025)
Sulistyo 2017 [116]	Thailand	Cross-sectional 81 from 1 clinic	Modified Diabetic Foot Care Knowledge Questionnaire [159]	58% with moderate, 39.5% poor knowledge.
Tuha 2021 [135]	Ethiopia	Cross-sectional 344 from 1 hospital	Details not provided	72.7% knew to inspect their feet for ulcers.
Health Care Providers				
Alotaibi 2017 [42]	Saudi Arabia	Cross-sectional 423 nurses at 1 hospital	Diabetes Basic Knowledge Test [162]	75.6% mean number of questions right regarding diabetic foot care.
El Hajj 2018 [125]	Qatar	Cross-sectional 126 pharmacists	Michigan Diabetes Research and Training Center Diabetes	25% with moderately poor knowledge.

Knowledge Test
[163]

Table 3. Summary of studies that investigated nephropathy knowledge among patients with diabetes and health care providers.

Studies arranged alphabetically. Abbreviations: CKD, chronic kidney disease; GP, general practitioner; T2D, type 2 diabetes.

Study	Country	Study Type & Population	Measure	Main Findings
Patients with Diabetes				
Alvis Zibran 2019 [140]	Fiji	Cross-sectional 225 with T2D and CKD from 1 hospital	Previously used KAP questionnaire [164]	61.8% with high knowledge.
Hussain 2019 [141]	India	Cross-sectional 323 T2D from 1 endocrinology clinic	Adapted CKD awareness questionnaire [165]	21.4% had good knowledge. Predictors of knowledge: literacy, income, socioeconomic status (p<0.05).
Kumela Goro 2019 [145]	Ethiopia	Cross-sectional 208 with hypertension and diabetes from 1 hospital	Study-created questionnaire	63.5% with poor knowledge
Lo 2017 [146]	Australia	Cross-sectional 308 patients with CKD and diabetes from 4 hospitals	Study-created questionnaire	43.5% cited inadequate knowledge of CKD and poor education about CKD as a barrier to care.
Health Care Providers				
Wolide 2020 [147]	Ethiopia	Cross-sectional 325 providers at 1 hospital and 3 private clinics	Study- created questionnaire	Predictors of knowledge: subspecialist provider (p<0.05).
Wong 1999 [142]	US	Cross-sectional 216 GPs	Study- created questionnaire	91.4% with good risk factor knowledge.
Yaqub 2013 [143]	Pakistan	Cross-sectional 232 GPs in 1 city	Study-created questionnaire	80% knew risk factors for CKD, 41% were unsure when to refer to nephrology

Figure Legends

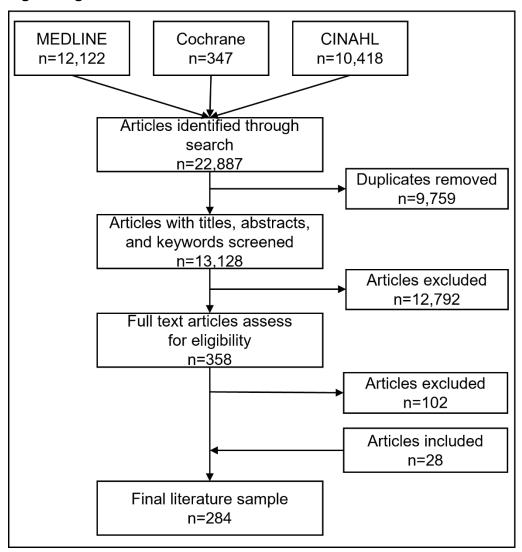


Fig 1 Flow diagram of study inclusions. PRISMA criteria were adopted

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