Review Article

The scoping review of Chinese and Western medicine treatment of diabetic foot in Asia

Abstract

Background of Study: Diabetic foot is a common and serious chronic complication of diabetes due to the simultaneous occurrence of diabetic peripheral neuropathy and vascular lesion. Among all complications, foot ulcers in diabetic ulcers account for the first place among the reasons for hospitalization and treatment of diabetic patients. 15% of diabetic patients may have foot diseases, and 85% of patients may have foot ulcers as the cause of amputation. Diabetic foot seriously affects the quality of life of patients. Although there are many methods to treat diabetic foot, the therapeutic effect of diabetic foot is not ideal in general. **Objectives:** The main purpose of this scoping review was analyzing the existing loopholes of researches on diabetic foot in Asia. Methods: Used PubMed, CNKI, Wangfangdata, CQVIP to search and select 5 traditional Chinese medicine literature and 5 western medicine literature, through the comparison of various conditions between literature to analyze the lack of research. Result and Discussion:10 pieces of literature were retained through 183 records and included 9 drugs or decoction, they were adipose-derived stem cell-hydrogel complex, hyperbaric oxygen therapy, Xenogeneic(poreine) acellular dermal matrix, alprostadil, salvia miltiorrhiza polyphenols for injection and collagen sponge, TaohongSiwu Decoction, Simiao Yong ' an Decoction, Jiawei Simiao Yong ' an Decoction, HuangqiGuizhiWuwu Tang, and WuweiXiaodu Drink. The obvious problems found by this scoping review were the quantity and quality deficiency of the research in the diabetic foot in Asia. Conclusion: Scoping review is an effective method of evidence identification and synthesis, which can provide a basis for the further development of acertain field. In the further study of the diabetic foot, more attention should be paid to the verification of experimental data as well as the feasibility of the researches on oral drugs.

Keyword: Diabetic foot, Traditional Chinese medicine, Western medicine

Introduction

Diabetic foot is a general term for a series of clinical manifestations of the foot caused by peripheral vascular disease, neuropathy, and infection in diabetic patients due to long-term and chronically elevated blood glucose, including foot ulcers, infection and deep tissue destruction. Once diabetic foot occurs, it is difficult to treat, with the high cost and poor prognosis. If the treatment is not timely or adequate, there is often a risk of amputation. Diabetic foot is one of the main complications of diabetes. According to the IDF diabetes map (9th edition), about 463 million adults worldwide suffered from diabetes in 2019, and it is estimated that 700 million people could have diabetes by 2045 Figure 1.1 shows the global distribution of diabetes patients in 20191.

At the national level, the top three countries with diabetes in 2019 were China, India and the United States, with about 118 million, 78 million and 30 million diabetics (20-79 years old), respectively. And China and the United States also have a high expenditure with diabetics.1 Figure 1.2 shows the top 10 countries with the highest number of diabetes in 2019. Based on the above data, the number of diabetes patients in Asia is the first in the world.

In global diabetes research, the study of American forces dominates. Diabetes research in China in the number of papers and patents statistics on has and Germany are close to, or more than these countries, but from the paper quality in new drug research and clinical research level, with Europe and the United States and other developed countries have a certain gap. (2)

Because Asian countries the number of diabetes and the number of concurrent with diabetic foot ulcers are more, but at this stage of clinical research level is not high, the number of new drug research, research direction is fuzzy, therefore decided to Asian drug therapy for diabetic foot ulcers group scoping review, to provide reference for drug treatment for the future research direction.

A scoped review is a design approach "to rapidly map the key concepts underpinning the research field as well as the key resources and types of available evidence" (3). This scoping review is to summarize the current clinical research literature on Chinese medicine and western medicine for diabetic foot treatment in Asia, so as to provide new ideas for the future research direction of diabetic foot medicine in Asia and facilitate the further development of research.

Methodology

Eligibility Criteria

Literature including nearly five years (2015-2019) in four databases of free text articles. Literature type limited to the clinical trial. Intervention limited to drugtreatment. Study design limited to randomized controlled trials. The population limit for Asians, any gender, most was over 35 years old, and the study population was limited to 30 or more (N > 30).

Search Strategy

The databases used are CNKI (China National Knowledge Infrastructure), PubMed and CQVIP (China Science and Technology Journal Database), Wanfang Data Knowledge Service Platform.

In the process of visual search of three databases, the search of the literature will be stopped until the literature in the database does not conform to the content of the literature required by the re problem which is researched.

The search strategy is based on the review of the clinical research progress of Chinese and western medicine in treating diabetic foot and the management of diabetic foot ulcers and extends to the differences in the treatment of diabetic foot with Chinese and western medicine. (4)(5) The common terms used in different databases include diabetic foot (including diabetes foot); broad terms such as non-surgical debridement agents; narrower terms such as hydrogels. The literature published after 2015 was searched until all the required literature was retrieved.

Study Selection

Three researchers examined the search records of the first layer cooperatively, at the first place screening by the published date, title and abstract, secondly, by full text. When a disagreement arises between the researchers, the controversial papers should first be retained and then screened according to drug type of the papers as well as a comparison of the literature describing the same classification of drugs.

Charting the Data

Before the data analysis, two articles were used to discuss the process of data analysis. After collecting the data, three researchers summarized, analyzed and compared, and agreed on the types of data items including drug therapy, study method, inclusion and exclusion criteria of participants (not included in the report) and treatment results.

Results

The purpose of this scoping review is to summarize the existing studies on drug therapy (western medicine and Chinese medicine) for diabetic foot patients in Asia, so as to provide references for future research directions on drug therapy and the combination of Chinese and western medicine. Figure 3.1 shows the process of identifying, screening, including, and excluding literature. Ten available literature remained.

Components of Drug Therapy

After screening, 10 articles and 9 drugs for treating diabetic foot ulcers were retained. There were 5 kinds of western medicines, includingAdipose-Derived Stem Cell–Hydrogel Complex, Hyperbaric oxygen therapy, Acellular dermal matrix, Alprostadil, Salvianolate. Chinese medicine prescription 4 kinds, including TaohongSiwu Decoction,Simiao Yong' an Decoction, HuangqiGuizhiWuwu Tang, WuweiXiaodu drink. TaohongSiwu Decoction has two pieces of literature, while other medicines only have one. Table 3.1 shows the nine drugs and their effects in treating diabetic foot.

treating diabetic root.		
Drug therapy	Effects	literature
Adipose-Derived Stem	Promote granulation tissue	Potential of Allogeneic Adipose-
Cell–Hydrogel	formation, epithelialization,	Derived Stem Cell-Hydrogel
Complex	and angiogenesis, thereby	Complex for Treating Diabetic
	promoting wound healing.	Foot Ulcers (6)
Hyperbaric oxygen	Increase tissue oxygen level,	A Pilot Study of Short-Duration
therapy	accelerate wound healing,	Hyperbaric Oxygen
	reduce edema, kill anaerobic	Therapy to Improve HbA1c,
X	bacteria.	Leukocyte, and Serum
		Creatinine in
		Patients with Diabetic Foot
		Ulcer Wagner 3-4 (7)
Xenogeneic(porcine)	Prevent pollution, promote	Application effect of the
acellular dermal	blood supply reconstruction	xenogeneic (porcine) acellular
matrix(ADM)	and increase cell	dermal matrix in the repair of
	regeneration.	diabetic foot ulcer (8)
Alprostadil	By activating adenylate	Analysis of the effect of
	cyclase to promote vascular	alprostadil on diabetic foot (9)
	dilation, inhibit platelet	
	aggregation, improve	
	microcirculation.	
Salvia miltiorrhiza	Effectively improve	Effects of Salvia miltiorrhiza
polyphenols	microcirculation, increase	polyphenols for injection
	microvascular blood flow,	Adipose-Derived Stem Cell-

	and anticoagulation effect.	Hydrogel Complex
		and collagen sponge in the
		diabetic foot (10)
TaohongSiwu	Vasodilation,anti-	The clinical curative effect of
Decoction	inflammation, anti-fatigue,	modified peach Tao Hong Siwu
	anti-shock, regulation of	decoction and lipoic acid
	immune function, lipid	injection on diabetic foot (DF)
	reduction, supplementation	in 50 cases (11)
	of trace elements, anti-	
	allergic effect.	
Jiawei Si Miao Yong	Resist inflammation, regulate	Clinical Observation of Jiawei
An Decoction	blood coagulation and	Si Miao Yong An Decoction in
Or Simiao Yong' an	immunity, reduce blood	the Treatment of Diabetic Foot
Decoction	glucose and inhibit bacteria,	Ulcer of Lower limb (12)
	strengthen erythrocyte	
	deformability, improve	Research of intervention by
	microcirculation and	Simiaoyong'an decoction for
	scavenging free radicals.	vascular endothelial function in
		the diabetic foot(13)
HuangqiGuizhiWuwu	Improve the vascular	Clinical Effect of Modified
Tang	function and hemodynamics	HuangqiGuizhiWuwu Tang on
	of lower limbs, have certain	Patient with Lower-extremity
	anti-inflammatory and	Arterial Disease (14)
	oxidative stress effects and	
	reduce the injury of vascular	
	endothelial cells.	
WuweiXiaodu Drink	Play an active role in	Clinical observation on
	antibacterial and anti-	treatment of type 2 diabetic foot
	inflammatory, and inhibits	ulcer by adding or subtracting
	the inflammatory response of	WuweiXiaodu Drink combined
	patients.	with tissue engineering full-
	_	thickness skin (15)

Table 3.1Nine drugs or prescriptions and their effects in treating diabetic foot.

Experimental design

The experimental designs were based on the route of drug use, different experimental sites, and the degree of disease. Comparing different experimental designs needed to be done from different aspects, thus the following table 3.2 and table 3.3 discussed the ten literature from the number of participants, age of participants, experimental location, country, degree of the lesion in patients and specific experimental operation.

ald)	Number of participants	e	Experimental location	Country	Degree of lesion of patients
------	---------------------------	---	-----------------------	---------	------------------------------------

		1	r		1
Adipose-	44	18-80	1. Yonsei University	Korea	Type of
Derived Stem			Severance Hospital		diabetes: type
Cell–Hydrogel			2. Eulji General		1 or type 2
Complex (6			Hospital		diabetes;
			3. Asan Medical		Ulcer duration:
			Center		longer than 4
			4. Korea University		weeks for the
			Guro Hospital		history of ulcer
					at screening;
					Wound size:
					between 1 and
					25 cm2;
					Wound depth
					of Wagner
					grade 1 and 2
Hyperbaric	30	20-79	Institutional Review	Indonesia	Diabetic foot
Oxygen Therapy	50	20-79	Board of Medical	muonesia	ulcer (DFU)
(7)			Faculty of Udayana		Wagner 3-4
(7)			University and		wagner 5-4
			Sanglah General		
V	7(10 (0	Hospital Denpasar Shizuishan first	China	Distration for at
Xenogeneic(por	76	18-68		China	Diabetic foot
cine) acellular			people's hospital of		ulcer (DFU)
dermal matrix			Ningxia hui		Wagner 2-3
(8)			autonomous region		DFU
					Wagner2: 47
					DFU
					Wagner3: 29
Alprostadil (9)	100	Male: 48-81	Shanghai Ruijin	China	\backslash
	Q.	Female: 43- 78	rehabilitation hospital		
Salvia	115	44-72	Metabolic disease	China	Diabetic foot
miltiorrhiza			hospital of Tianjin		ulcer (DFU)
polyphenols for			medical university		Wagner 2-3
injection and					
collagen sponge					
(10)					
TaohongSiwu	100	44-76	Pinghu hospital of	China	Diabetic foot
Decoction (11)	100	,0	traditional Chinese	Jiiiiu	ulcer (DFU)
			medicine		Wagner 0-1
			medicine		wagner 0-1
Simiao Yong'	50	>60	Henan Province	China	
an Decoction			Hospital of TCM		
(13)			-		
Jiawei Simiao	86	58-74	People's hospital of	China	Diabetic foot
Yong' an			Wuhan university	Junia	ulcer (DFU)
e			wanan aniversity		Wagner 1-4
Deposition (12)	l – – – – – – – – – – – – – – – – – – –	1			
Decoction (12) HuangqiGuizhi	128	64-74	Weifang People's Ho	China	Fontaine stage

Wuwu Tang (14)			spital		I to III
WuweiXiaodu drink (15)	150	41-63	Department of dermatology, Cangzhou central hospital, Hebei province	China	Diabetic foot ulcer (DFU) Wagner 3-4

Table 3.2 The basic information of all the experiments.

	
4.1	Specific experimental operation
Adipose-	Of the 44 participants, 25 were assigned to the experimental group and 19
Derived Stem	were assigned to the matched group. 3 participants in the experimental
Cell-Hydrogel	group and 2 participants in the matched group were excluded because they
Complex (6)	did not reach high blood glucose levels (>450). Patients of both groups
	were treated with 3% hydrogen peroxide and a saline solution to remove
	dirt and other debris from their ulcers, and both groups were covered with
	polyurethane foam. The difference between the two groups' patients was
	that the experimental group used allograft ASC sheets as the primary
	dressing and the matched group used mesh polyurethane films with
	silicone adhesives as the primary wound dressing. In both groups, drugs
	were changed once a week. If necessary, patients could go to the hospital
	to check the wound two or three times a week. If there were any adverse
	reactions, the auxiliary dressing could be replaced.
Hyperbaric	Blood tests were performed on all patients to detect HbAlc levels, white
Oxygen	blood cell counts and serum creatinine levels before debridement, and they
Therapy (7)	were divided into two groups. Among them, the experimental group
	conducted ten courses of HBOT(One session of HBOT uses oxygen at 2.4
	ATA for 90 minutes per day at several hyperbaric chambers) and the
	matched group adopted the standard treatment. Five courses of treatment
	per week, it took two weeks altogether.
Xenogeneic(po	All participants were given anti-infection, nutritional support treatment,
rcine) acellular	diabetes specialist guidance diabetic diet, insulin treatment, monitoring and
dermal matrix	regulating blood sugar, wound secretion culture-sensitive antibiotics
(8)	guide treatment as well as other comprehensive systemic treatment. The
	wound surface was washed with 1.5% hydrogen peroxide solution, iodine-
	volt solution and 0.9% sodium chloride solution in turn, and the wound
	surface was wrapped and fixed with gauze, and the dressing was changed
	once a day.
	Experimental group: One week later, under subarachnoid anesthesia and
	spinal anesthesia, the Sonoca180 low-frequency ultrasonic debridement
	device (produced by Germany surin-cieser company) was applied to
	debridement of the diabetic foot, and the wound surface was expanded and
	created stealthily. Clipping heterogeneous (swine) ADM size bigger than
	wound for $2 \sim 3$ cm, then used 0.9% sodium chloride solution cleaning the
	wound for three times, the duration of cleaning was about 3 min, then soak
	in dilute iodine volts fluid (liquid iodine volts: 0.9% sodium chloride
	in anute toume vons thuta (inquita toume vons. 0.9% soutum chioride

	solution is 1:1) for 1 min, guarantee level toward the wound in the
	heterogeneous skin, fully cover the wound, external use sterile dressings
	bandage fixation after six floors, 1 week to replace one.
	Matched group: The matched group was treated with sulfadiazine silver
	cream (1% sulfadiazine silver, 60 g, Guangdong Hengjian pharmaceutical
	co., LTD.).8
Alprostadil (9)	Of the 100 participants, 50 were assigned to the experimental group and 50
	to the matched group. Patients in both groups received conventional
	treatment and were treated with hypoglycemic drugs. The wounds were
	washed with normal saline and hydrogen peroxide, and the wound was
	wrapped with iodoform gauze. Both groups dissolved 20ml tanshinone
	injection in normal saline, and the patients were given intravenous infusion
	once a day. The difference between the two groups was that the
	experimental group was treated with alprostadil injection, 100mg of
	alprostadil injection was dissolved in saline, and the patients were given
	intravenous infusion once a day. Patients were treated for 2 weeks and the
	results were reviewed.
Salvia	Of the 115 participants, 60 were assigned to the matched group and 55 to
miltiorrhiza	the experimental group. Patients in both groups were treated with
polyphenols	conventional type 2 diabetic foot therapy, wound cleaning, and spraying
for injection	0.9% sodium chloride solution on the wound with the same type of
and collagen	antibiotic. The difference between the two groups was that the patients in
sponge (10)	the experimental group were treated with collagen dressing, dissolved 200
	mg Danshen polyphenols in 5% glucose injection, and given intravenous
	infusion once a day. For four weeks of one course of treatment, patients'
	ulcer recovery before and after treatment and during treatment were
	observed after two courses of treatment.
TaohongSiwu	All participants received basic treatment for diabetes, including diet
Decoction (11)	control, insulin control of blood glucose, and decompression therapy. Foot
	superficial ulcer was disinfected with iodophor, local antibiotics were
	given wet compress, and on this basis, the experimental group was given
	Tao Hong Siwu Decoction(Tao Ren 30g, Hong Hua 30g, Dang Gui 30g,
	Chuanxiong 20g, Shu Di Huang 20g, Shen Jin Cao 30g, Hai Feng Teng
	30g, Ai Ye 30g), which was fried and soaked in feet once a day for 15
	minutes each time. At the same time, the blood flow velocity of the dorsal
	foot artery and sensory conduction velocity of the common peroneal nerve
	and superficial peroneal nerve were measured before and after treatment.
Simiao	All patients were given diabetes education and a diabetic diet, and the
Yong' an	experimental group was given Simiao Yong' an Decoction (Dang Gui
Decoction (13)	30g, Xuan Shen 15g, Huang Qi 15g, Jin Yin Hua 15g, Ji Xue Teng 15g,
Decoction (13)	Gan Cao 6g) which was fried in water, one dose per day and separated one
	dose for two times. In the meantime, the matched group was not given any
	medication.
Jiawei Simiao	Of the 86 participants, 43 were assigned to the experimental group and 43
Yong' an	to the matched group. Patients in the two groups were first given
Decoction	conventional treatment to control their blood glucose, and the same
(12)	antibiotics were selected to control infection. Physical methods and drugs
	were used to improve the blood circulation in the lower extremities of the

	patients. The ulcer of the patients was changed once a day, and the patients were given cilostazol tablets twice a day, 100mg each time. The difference between the two groups of patients was that the experimental group took
	modified Simiao Yong' an Decoction orally (Yin Chen 15g,Chui Pen Cao
	30g, Pu Huang 10g, Sheng Mu Li 30g, Dang Gui 10g, Xuan Shen 15g, Chi
	Shao 10g, ChuanNiu Xi 15g, Tu Fu Lin 30g, Yi Yi Ren 30g, Xi Lian Cao
	30g, Huang Lian 9g, Rou Gui 4.5g), once a day, and took 200ml in the
	morning and evening.
HuangqiGuiZh	Of the 128 participants, 64 were assigned to the experimental group and 64
iWuwu Tang	to the matched group. Two groups of patients simultaneously received non-
(14)	drug treatment, control of blood glucose, blood pressure, blood lipid, and
	other drug treatment. Patients were given probucol tablets at breakfast and
	dinner with 0.5g each time. Oral aspirin enteric-coated tablets were taken
	100mg per day for 3 months. Intravenous infusion of alprostadil injection,
	10 mg a day, plus 10ml normal saline, continuous injection for 15 days, 15
	days rest for a course of treatment, a total of 3 courses of treatment. The
	difference between the two groups was that the experimental group was
	given orally modified HuangqiGuizhiWuwu soup (Huang Qi 30g, GuiZhi
	10g, Bai Shao 15g, Sheng Jiang 10g, Da Zao 10g, Tian Hua Fen 15g, Tai Zi Shan 20g, Di Lang 10g, Ji Yang Tang 20g, Dan Shan 20g, Orang Xia 5g
	Zi Shen 20g, Di Long 10g, Ji Xue Teng 30g, Dan Shen 20g, Quan Xie 5g,
	Gan Cao 5g). Fu Ling 20g, Kun Bu 10g, Bai Jiang Can 10g were added to
	patients with phlegm, which were prone to fatigue and physical weakness.
	Patients who did not want to speak were added to Dang Shen 15g, Chao
	Bai Zhu 15g. Patients with hot chest, palm and foot were added to Xuan
	Shen 15g and Mu Dan P115g. Soak the medicine in water for 30min and
	cook it twice a day. Take 150ml each morning and evening. Both groups
	received continuous treatment for 3 months.
WuweiXiaodu	Of the 150 participants, 75 were assigned to the experimental group and 75
Drink (15)	to the control group. Patients in the two groups were given insulin injection
	at the same time, 20 units a day, and subcutaneously injected at 3 times in
	the morning, the middle and the evening 30 minutes before the meal, and
	the patients were given tissue engineering full-layer skin treatment. The
	difference between the two groups was that the experimental group was
	treated with WuweiXiaodu drink (Jin Yin Hua 30g, Ye Ju Hua 30g, Pu
	Gong Ying 30g, Dang Gui 20g, Tian Hua Fen 20g, Lian Qiao 15g, Dong
	Kui Zi 15g, Mu Dan Pi 15g, Niu Xi 15g, Bai Shao 15g, Bai Zhi 15g,
	Huang Qin 15g, Zi Hua Di Ding 10g, San Qi Fen 3g). For patients with
	pain, add 10g Sheng Ru Xiang and 10g Sheng Mo Yao. For patients with
	blood stasis, add Chi Shao 10g and Ji Xue Teng 15g. Take 150ml each
	morning and evening. Both groups were treated for 8 weeks.
	Table 3.3 The details of all the experimental designs.

Table 3.3 The details of all the experimental designs.

Discussion

Nine different Chinese medicines and western medicines were used to treat diabetic foot. In the above paragraphs, different Chinese medicines and western medicines were compared one by one. Next, this section will analyze 10 articles used to determine the research direction of treating diabetic foot in the future.

In "The Potential of Allogeneic Adipose-Derived Stem cells - Hydrogel Complex for Treating Diabetic Foot Ulcers"(6), the author said research is likely to be

the first report to prove Allogeneic Adipose-Derived Stem Cell for the effectiveness of the treatment of diabetic foot ulcers. So, the study has limitations, the action mechanism of allogeneic adipose-derived stem cells still need to be further confirmed. In the study of short-duration hyperbaric oxygen therapy(7), the author found that the glucose metabolism and the mechanism of improving human renal function were still unclear, and further studies were needed to determine the effect of short-duration hyperbaric oxygen therapy on diabetic foot ulcer patients. When using the xenogeneic(porcine) acellular dermal matrix to repair diabetic foot ulcers(8), it can not only shorten the wound healing time, reduce the number of dressing changes, but also reduce the psychological and economic burden of patients and reduce the infection rate. At the same time, it has wide sources, simple operation, and low price, and is worthy of clinical promotion. In the study of alprostadil(9), it has been determined that alprostadil has a better clinical effect and higher safety. Research shows that alprostadil is worthy of extensive application in clinical practice. In the process of studying salvia miltiorrhiza polyphenols for injection and collagen sponge(10), the direct therapeutic effect of salvianolic on diabetic foot still needs further study, but it can improve microcirculation, prevent microthrombosis, regulate vascular endothelial dysfunction and other effects have been able to play a positive role in the treatment of diabetic foot. The combination of salvianolic and collagen sponge played an active role in the treatment and healing of diabetic foot. In the process of studying of the TaohongSiwu Decoction(11), it has proved flavoredTaohongSiwu Decoction combined thioctic acid injection on diabetic foot has good effect, but the author use samples is limited, did not add the diabetic foot higher severity of patients, so in further research, more data are needed to determine the curative effect of TaohongSiwu Decochion. In the two articles of Simiao Yong' an Decoction, Jia Wei Simiao Yong an Decoction can improve the microvascular lesions of diabetic foot by repairing the damaged intima of arteries(12), inhibiting inflammation(13), and so on so that diabetic foot can be relieved. However, the literature on Simiao Yong an Decoction is still very limited, and the molecular mechanism of Simiao Yong an Decoction remains to be further studied. In the study of HuangqiGuizhiWuwu Tang(14), it was proved that HuangqiGuizhiWuwu drink could improve the vascular function and hemodynamics of the lower extremities, reduce the injury of vascular endothelial cells, and thus play a role in alleviating the progression of diabetic lower extremity vascular diseases. It is worthy of clinical promotion and application of such drugs. In the process of studying WuweiXiaodu drink combined with tissue engineering full-thickness skin (15), it has been determined that it can improve the efficacy, improve the patient's condition and play an anti-inflammatory role without obvious adverse reactions. Therefore, it is a drug worthy of clinical promotion.

Drug therapy	Total effective rate
Adipose-Derived Stem Cell–Hydrogel	Matched group: 47%(8/17)
Complex (6)	
	Experimental group: 73% (16/22)
Hyperbaric oxygen therapy (7)	Matched group:
	HbA1c levels:
	Baseline: 10.98± 2.37
	Effect size: 1.28± 1.54

Г	~ 1
	Leukocyte count:
	Baseline: 14.27± 6.79
	Effect size: 3.26 ± 7.76
	Serum creatinine levels:
	Baseline: 0.73± 0.25
	Effect size: 0 ± 0.13
	Experimental group:
	HbA1c levels:
	Baseline: 9.42± 1.96
	Effect size: 2.34 ± 1.57
	Leukocyte count:
	Baseline: 13.97± 6.24
	Effect size: 5.13 ± 6.72
	Serum creatinine levels:
	Baseline: 2.10± 2.88
	Effect size: 0.05 ± 0.45
Xenogeneic(porcine) acellular dermal	Matched group: 78.95% (30/38)
matrix (8)	Experimental group: 94.74% (36/38)
Alprostadil (9)	Matched group: 78% (39/50)
	Experimental group: 98% (49/50)
Salvia miltiorrhiza polyphenols for	Matched group: 30.91% (22/60)
injection and collagen sponge(10)	Experimental group: 60% (33/55)
TaohongSiwu Decoction (11)	Matched group: 68% (34/50)
	Experimental group: 84% (42/50)
<i>.</i>	
Simiao Yong' an Decoction (12)	Matched group:
	ET-1 (pg/ml): 52.06± 5.32
	NO(μmol/L): 67.15± 2.93
	TXB ₂ (pg/ml): 84.02± 3.99
	6-Keto-PGF1α(pg/ml): 72.55± 3.89
	Experimental group:
	Before the treatment:
	ET-1 (pg/ml): 74.09± 4.87
	NO(μmol/L): 54.18± 7.97
	TXB ₂ (pg/ml): 101.60± 9.78
	6-Keto-PGF1α(pg/ml): 65.07± 5.83
	After the treatment:
	ET-1 (pg/ml): 62.05± 4.78
	$NO(\mu mol/L): 65.81 \pm 3.83$
	$TXB_2(pg/ml): 91.84 \pm 8.90$
	$6-Keto-PGF1\alpha(pg/ml): 70.77\pm 7.00$
Jiawei Si Miao Yong An Decoction (13)	Matched group: 86.05% (37/43)
	Experimental group: 93.02%(40/43)
HuangqiGuizhiWuwu Tang (14)	Matched group: 92.2% (59/64)
	Experimental group: 100% (64/64)
	r · · · · · · · · · · · · · · · · · · ·
WuweiXiaodu drink combined with	Matched group: 73.33%(55/75)

Table 3.4 The total effective rate of all the experiments.

Conclusion

The purpose of the scoping review is to review the general situation of a certain research field, based on this paper, the principal contradiction discussed in this paper is comparing the researches of partial traditional Chinese medicine and western medicine treatment of the diabetic foot. Through studied and thought of these ten pieces of literature, some problems emerged obviously. The first problem was that the literature of diabetic foot in China was deficient. On the one hand, the quantity of literature in China may be considerable, on the other hand, the quality was dissatisfactory. These were two pieces of literature that had some statistical errors, both of these two pieces of literature were written by Chinese researchers. Proportionately, this kind of error maybe not only existed in the 10 literature but had a big amount in the published literature in China. The second problem was that the research of diabetic foot in Asia was not enough. According to the approach of PUBMED searches, a total of 60 articles were excluded from 33 because of geographical conditions. The third problem was that most of the literature that meets the requirements of this theme were cited fewer times, even some literature hasn't been cited before. That confirmed the lack of research on diabetic foot in Asia from another perspective and the low quality of that literature. To sum up, this scoping review proves that there was still a big loophole in the quality of research on diabetic foot in Asia.

Abbreviation:CNKI = China National Knowledge Infrastructure; CQAIP = China Science and Technology Journal Database; WANFANG DATA =Wanfang Data Knowledge Service Platform

References

- 1. IDF. (2019, 12 1) (IDF) Retrieved March 02 2020, from: https://diabetesatlas.org/en/resources
- 2. Lu, Y., & Yu, J. (2012). International R&D trends in diabetes. *International Diabetes Research and Development Trend*, 24(7), 634–639.
- 3. Uçkay, I., Kressmann, B., Malacarne, S., Toumanova, A., Jaafar, J., Lew, D., & Lipsky, B. A. (2018). A randomized, controlled study to investigate the efficacy and safety of a topical gentamicin-collagen sponge in combination with systemic antibiotic therapy in diabetic patients with a moderate or severe foot ulcer infection. 1–9.
- 4.Guo, W. G., &Xue, Z. M.(2019) Clinical research progress of Chinese and western medicine in the treatment of diabetic foot. Jilin journal of traditional Chinese medicine,2019,39(04):548-550.
- Estelle, E., &Nestoras, M.(2019) Update on management of diabetic foot ulcers. Ann N Y Acad Sci . 2018 January ; 1411(1): 153–165. doi:10.1111/nyas.13569.
- Moon, K. C., Suh, H. S., Kim, K. B., Han, S. K., Young, K. W., Lee, J. W., & Kim, M. H. (2019). Potential of Allogeneic Adipose-Derived Stem Cell–Hydrogel Complex for Treating Diabetic Foot Ulcers. *American Diabetes Association*, 68(4), 837-846
- 7. Irawan, H., Semadi, I. N., & Widiana, I. G. R. (2018). A Pilot Study of Short-Duration Hyperbaric Oxygen Therapy to Improve HbA1c, Leukocyte, and

Serum Creatinine in Patients with Diabetic Foot Ulcer Wagner 3-4. *Scientific World Journal*, 2018. https://doi.org/10.1155/2018/6425857

- Beng, X., W., Zhang, L., Jin, S., H., Q, X., H., & Li, C., J. (2013). Application effect of xenogeneic(porcine) acellular dermal matrix in the repair of diabetic foot ulcer. *Journal of Chemical Information and Modeling*, 53(9), 1689–1699. <u>https://doi.org/10.1017/CBO9781107415324.004</u>
- 9. Chen, F.(2019). Analysis of the effect of alprostadil on diabetic foot. *Contemporary Medical Symposium*, 17 (3), 135-136
- Yao, W. J., Wang, P. H, Xu, J., Li, X. M., & Ding, M. (2016). Effects of salvia miltiorrhiza polyphenols for injection and collagen sponge in diabetic foot. *Chinese journal of biochemical medicine*, 11 (36), 131-137
- Chen, J., Sun, P., &Wang, W. R. (2018). The clinical curative effect of modified peach Tao Hong Siwu decoction and lipoic acid injection on diabetic foot (DF) in 50 cases. *Clinical Journal of Chinese Medicine*, 2018,10(30):59-61.

12. Zhou, T., Hu, T., R., Hu, J., C., Deng, Q., L., & Wu, H. (2016). Clinical Observation of Jiawei Si Miao Yong An Decoction in the Treatment of Diabetic Foot Ulcer of Lower Limb. *Chinese Journal Of Microcirculation*, 26 (3), 49-53

- Li, J., P., Wang, D., W., & Wu, H., L. (2015). Research of intervention by Simiaoyongan decoction for vascular endothelial function in diabetic foot. *China Prac Med*, 17(10), 48-49
- Cheng, Y., Luan, K., D., Ding, H., Y., Tang, Y., Wang, J., & Yu, W., G. (2019). Clinical Effect of Modified Huangqi Guizhi Wuwu Tang on Patient with Lower-extremity Arterial Disease. *Chinese Journal of Experimental Traditional Medical Formulae*, 15(25), 136-141
- 15. Zhang, G. J., Feng, S. J., Wang, Z. X., &Ai, D. F. (2019). Clinical observation on treatment of type 2 diabetic foot ulcer by adding or subtracting Wuwei Xiaodu Drink combined with tissue engineering full-thickness skin. *Hebei Journal of Traditional Chinese Medicine*, 2019,41(02):258-261.



CONFLICT OF INTEREST:We promise that no part of this paper has published or submitted elsewhere and no conflict of interest exists in the submission of this manuscript. There is the permission of using the figures from IDF.

APPENDIX : FIGURES: Figure 1.1 Number of diabetes patients worldwide (20-79 years old) in 2019 (1)

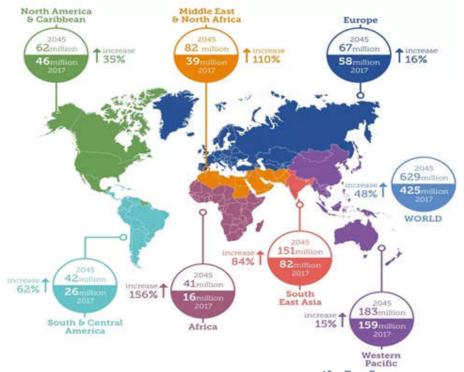


Figure 1.2 Top 10 countries with the highest number of diabetics in 2017 (the red bar represents health spending) (1)

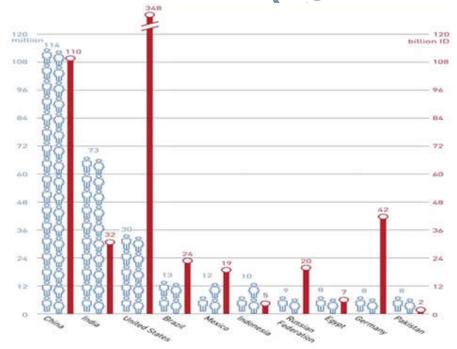


Figure 3.1Process of identifying, screening, including, and excluding literature.

