

Nutritional Management of Diabetes Mellitus: An appraisal of the role of Medicinal plants

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Summary

Diabetes mellitus is a heterogeneous metabolic syndrome characterized by chronic hyperglycemia with partial or total lack of insulin secretion and insulin resistance. The most common symptoms are polydipsia, polyuria, blurred vision, slow healing sore, nerve damage. Diabetes, being a metabolic, endocrine disorder is directly connected to carbohydrates, lipid, and protein metabolism. As a result, nutrition therapy forms an integral part of diabetes management. Daily caloric intake of 50% - 55% carbohydrates is recommended. Carbohydrate with low glycaemic index is preferred to those with high glycaemic index. Nonnutritive sweeteners are also encouraged for people with diabetes to add increased variety to their food choices. The protein requirement for persons with diabetes is not different from the general population, 15-20 % of total caloric intake. Both soluble and insoluble fibers are encouraged in amounts similar to the recommendations for the general population (20-35 g). Fibers are useful as they prolong gastric emptying, prevent constipation, lower serum cholesterol level, and reduce nutrients diffusion rates, thus reducing blood glucose response. A low-fat diet is advisable for diabetics in order to reach and maintain good weight and health. As part of a healthy diet, 30% of daily calories should come from fats, and of these less than 10% should be saturated fat, less than 10% polyunsaturated fat and 10-15% monounsaturated fat. As the general population, people with diabetes have no need for vitamin and mineral supplementation when the dietary intake is adequate. Despite much research on nutritional factors in the etiology and management of diabetes, the risk associated with several individual nutrients is not entirely clear. In order to achieve maximum benefit from nutritional interventions in the management of diabetes changes in government policies and legislation will be needed in addition to individual and community-based programs.

Keywords: Diabetes mellitus, Nutritional management, Medicinal plant, hyperglycemia

1.0 Introduction

Food supply and intake is very important to the body regardless of age, sex, religion, occupation or season. Food must be taken with consciousness for well-being of the body. Too much or too little food intake may cause serious damage to the body system. This can result to various diseases - among which is diabetes (Belinda, 2004).

Diabetes mellitus is a combination of heterogeneous disorders commonly present with episodes of hyperglycaemia and glucose intolerance, as a result of lack of insulin, defective insulin action, or both (Sicree *et al.*, 2006). Such complications arise due to derangements in the regulatory systems for storage and mobilization of metabolic fuels, including the catabolism and anabolism of carbohydrates, lipids and proteins emanating from

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defective insulin secretion, insulin action, or both (Votey and Peters, 2004). In addition to insulin, aging, over weight and several other hormones affect blood glucose level thereby preventing glucose from entering the cells (Clavel, 2005). This leads to hyperglycaemia, which may result in acute and chronic complications such as diabetic ketoacidosis, coronary artery disease, cerebrovascular disease, kidney and eye diseases, disorders of the nerves and others (Iwueze, 2007).

Classification of diabetes mellitus is based on its aetiology and clinical presentation. As such, there are four types or classes of diabetes mellitus viz; type 1 diabetes, type 2 diabetes, gestational diabetes, and other specific types (Sicree *et al.*, 2006). Type 1 diabetes is said to account for only a minority of the total burden of diabetes in a population, although it is the major type of the diabetes in younger age groups developed countries (Sicree *et al.*, 2006). Type 2 diabetes accounts for 85 to 95% of all diabetes in developed countries. It is intimately associated with improper utilisation of insulin by target cells and tissues. It is currently a common and serious health concern globally (Votey and Peters, 2004).

The dietary management of diabetes mellitus is a complement of lifestyle management. It has a positive effect on long term health and quality of life. Dietary management aims at optimal metabolic control by establishing a balance between food intake, physical activity, and medication to avoid complications. In type 2 diabetes, the dietary objective is for improved blood glucose and lipid levels and weight loss as appropriate (Piero, 2006). Because Diabetes Mellitus is a disease directly related to carbohydrate, lipid, and protein metabolism, nutrition has always had an integral role in its management. The contemporary term used to describe the dietary prescriptions is medical nutrition therapy (MNT) (American Diabetes Association, 2002). Before the advent of insulin therapy in the early 20th century, MNT was the only form of therapy for diabetes mellitus (Piero, 2006). The focus of nutritional recommendations at the turn of the last century was on type 1 diabetes mellitus: on control of the severe hyperglycaemia to which these patients are vulnerable as well as on prevention of what was then, before the advent of insulin therapy, commonly a mortal episode of ketoacidosis (Piero, 2006).

Medicinal plants play a significant role in the treatment of diabetes mellitus which is a serious metabolic disorder.

Traditional plants are reported to have significant anti-diabetic properties with no harmful side effects. They are rich sources of anti-diabetic compounds such as flavonoids, alkaloids, phenolic and tannins that improve the efficiency of pancreatic tissues by increasing the insulin secretion or decreasing the intestinal absorption of glucose (Bindu & Narendhirakannan, 2019; Kooti *et al.* 2016)

1.1 Biochemical Background of Diabetes Mellitus

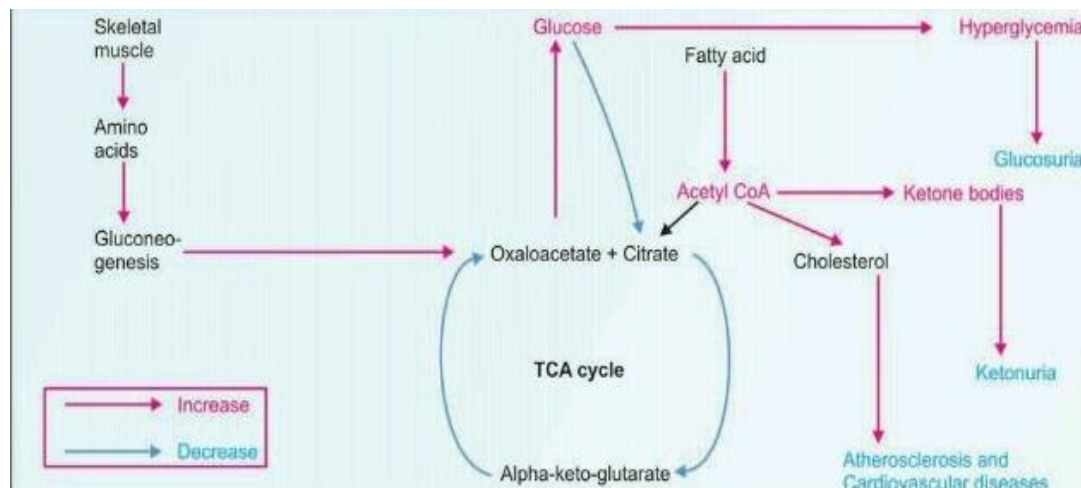
Diabetes mellitus is a group of diseases characterised by high levels of glucose in the blood resulting from defects in insulin production (insulin deficiency), insulin action (insulin resistance), or both. Insulin is a hormone produced by the pancreas. After eating, the carbohydrates are metabolised to glucose that enters the bloodstream. Insulin is needed for the uptake glucose into the liver where it is used for energy, and excesses are stored in the liver and fat cells. Insufficient amounts of working insulin cause blood sugar levels to rise and large amounts of glucose are excreted in the urine (Kibiti, 2006).

A regular energy source is a prerequisite for every cell to function in the human body. Glucose is the body's primary energy source, which circulates in the blood as a mobilisable fuel source for cells (Njagi, 2006). Insulin is a pancreatic hormone responsible for blood glucose level regulation. The hormone binds to its receptor sites on peripheral side of the cell membranes. It affords entry of glucose into respiring cells and tissues via requisite channels. Insulin stimulates catabolism of glucose into pyruvate through glycolysis. It also regulates glycogenesis and lipogenesis from excessive cytosolic glucose and acetyl-CoA respectively. These metabolic events are antagonistic to metabolic events triggered by the hormone glucagon. When glucose levels are at or below threshold, glucose stays in the blood instead of entering the cells (Belinda, 2004).

The body attempts to arrest hyperglycaemia, by drawing water out of the cells and into the bloodstream. The excess sugar is excreted in the urine. This is why diabetes present with constant thirst, drinking large amounts of water, and polyuria as the cells tries to get rid of the extra glucose. This subsequently leads to glucosuria (Piero, 2006).

As hyperglycaemia prolongs, the body cells are devoid of glucose due to the lack of insulin. This forces the cells to seek alternative mobilisable energy sources. In this regard,

diabetes mellitus. Buildup of ketone bodies in the blood produces ketosis. Ketone bodies are acidic in nature and therefore, their build up in blood lowers blood pH, leading to acidosis. A combination of ketosis and acidosis lead to a condition called ketoacidosis. If left untreated, ketoacidosis leads to coma and death (Belinda, 2004).



Type1 diabetes occurs commonly in childhood and adolescence, but can occur at any age (Piero *et al.*, 2014)

In Type I diabetes, the pancreas cannot synthesize enough amount of insulin as require by the body. The pathophysiology of this type of diabetes mellitus suggests that it is an auto immune disease in which the body's own immune system generates secretion of substance that attacks the beta cells of the pancreas. Some researchers suggest that in genetically susceptible people, certain viruses activate the immune system to attack and destroy cells in the pancreas as if they were foreign cells (Piero *et al.*, 2014). In type 2 diabetes, the more common type of diabetes whereby the cells fail to respond to insulin; this condition tend to occur as a consequence of obesity it usually develops in people over 40 years old (Njagi, 2006).

Another type of diabetes which is less common is gestational diabetes which develops in some women during pregnancy (Piero *et al.*, 2014).

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Pathophysiology of Diabetes Mellitus

Some Over-Simplifications

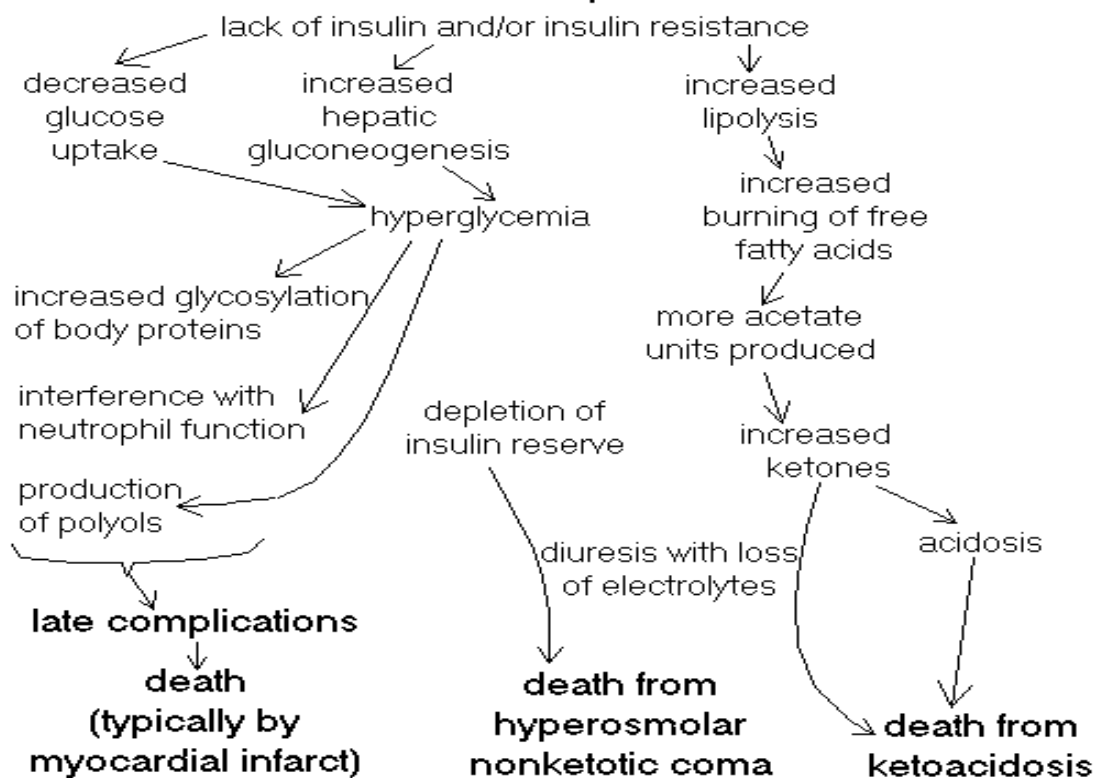


Figure 2: Pathophysiology of Diabetes Mellitus
Source: KrishnaDas, 2008

(ketones are byproduct lipid catabolism), nerve damage, etc. (Taiwo, 2010).

1.3 Symptoms of Diabetes Mellitus

Diabetes symptoms vary depending on blood sugar level of individual with diabetes. Type 2 diabetes, in particular, develops slowly. Many people have type 2 diabetes for as long as eight years before it is diagnosed. When symptoms do develop, they often vary. But two symptoms that occur in many people with the disease are increased thirst (polydipsia) and frequent urination (polyuria) due to excess glucose circulating in the body draws water from the tissues, making diabetics feel dehydrated. To quench the thirst, they drink a lot of water and other beverages, and that leads to more frequent urination (Davidson *et al.*, 2003). Other warning symptoms of diabetes mellitus are fatigue, irritability, blurred vision, slow-healing sores, frequent infections (such as gums or skin infections and vaginal infections), presence of ketones in the urine

1.4 Causes of Diabetes Mellitus

Diabetes mellitus is caused by a complicated interplay of genes, environment, insulin abnormality, increased glucose production in the liver, increased fat breakdown and possible defective hormonal secretion in the intestine. The recent dramatic increase indicates that factors such as obesity and sedentary lifestyle may be particularly important in triggering genetic elements that cause diabetes mellitus (Davidson *et al.*, 2003).

1.5 Classifications of Diabetes Mellitus

Assigning a type of diabetes to an individual often depends on the circumstances present at the time of diagnosis, and many diabetic individuals do not easily fit into a single class.

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For example, a person with gestational diabetes mellitus may continue to be hyperglycaemic after delivery and may be determined to have, in fact, type 2 diabetes. Alternatively, a person who acquires diabetes because of large doses of exogenous steroids may become normoglycaemic once the glucocorticoids are discontinued,

but then may develop diabetes many years later after recurrent episodes of pancreatitis (Baynes, 2015). However, there are two major types of diabetes. These are type 1 and type 2 diabetes. The other types are less common and they include gestational diabetes or pregnancy-induced diabetes and other specific type (Sicree *et al.*, 2006)

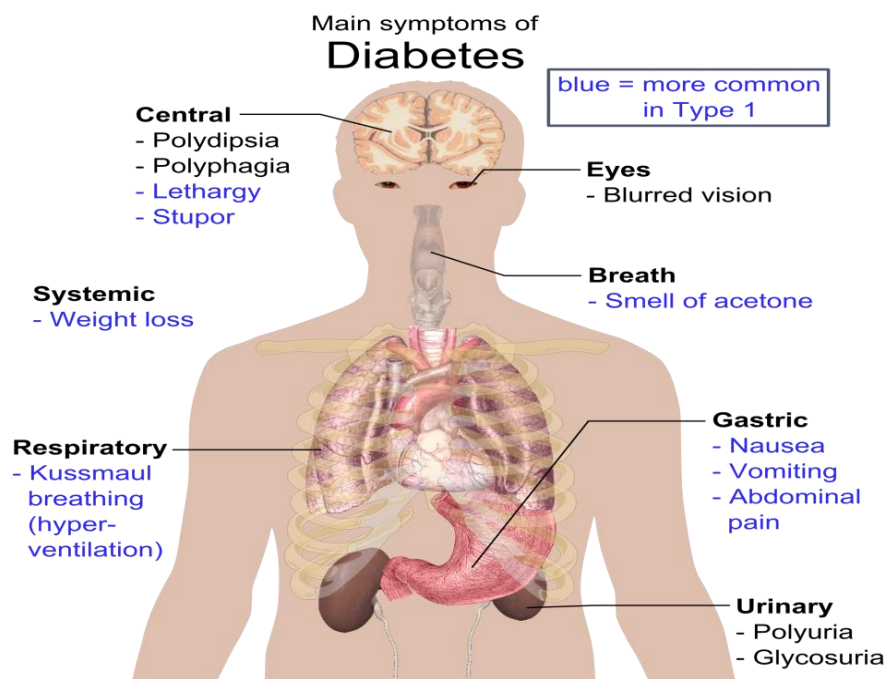


Figure 3: Overview of the most significant symptoms of diabetes

all type1 diabetic patients will require insulin therapy to maintain normoglycaemia (Baynes, 2015).

Sources: Davidson *et al.*, 2003

1.5.1 Type 1 Diabetes Mellitus

Type 1 diabetes is often known as insulin-dependent diabetes. Since it starts in children and in the young, it is also termed juvenile diabetes or early-onset diabetes. Type 1 diabetes mellitus is characterized by beta cell destruction caused by an autoimmune process, usually leading to absolute insulin deficiency (Kumar and Clark, 2002). Type 1 is usually characterized by the presence of anti-glutamic acid decarboxylase (anti-GAD) antibodies, islet cell or insulin antibodies that target an enzyme called glutamic acid carboxylase. This enzyme converts glutamic acid to gamma amino butyric acid (GABA), a neurotransmitter found in high concentration in the cerebellum. Eventually,

1.5.2 Type 2 Diabetes Mellitus

Type 2 diabetes is otherwise called non-insulin dependent diabetes mellitus (NIDDM). Type 2 diabetes is a heterogenous disorder caused by a combination of genetic factors related to impaired insulin secretion, insulin resistance and environmental factors such as obesity, over eating, lack of exercise, and stress as well as aging (Kaku, 2010). It is typically a multifactorial disease involving multiple genes and environmental factors to varying extents (Holt, 2004). Type 2 diabetes is the predominant form of diabetes and accounts for at least 90% of all cases of diabetes mellitus (Gonzalez *et al.*, 2009). The rise in prevalence is predicted to be much greater in developing than in developed countries (69% versus 20%) (Shaw *et al.*, 2010). This increase in type 2 diabetes is inextricably linked

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to changes towards a western lifestyle (high diet with reduced physical activity) in developing countries and the rise in prevalence of overweight and obesity (Chan *et al.*, 2009). The American Diabetes Association (ADA) recommends that testing to detect prediabetic should be considered in adults who are overweight or obese and have one or more risk factors for diabetes (Diabetes Prevention Program Research Group, 2009).

1.5.3 Gestational Diabetes Mellitus or Pregnancy-Induced Diabetes Mellitus

Gestational diabetes mellitus (GDM) is associated with women who develop Type 1 diabetes mellitus during pregnancy and women with undiagnosed asymptomatic Type 2 diabetes mellitus that is discovered during pregnancy. In most women who develop GDM; the disorder has its onset in the third trimester of pregnancy (Baynes, 2015).

1.5.4 Other specific type (Monogenic diabetes)

Types of diabetes mellitus of various known etiologies are grouped together to form the classification called "Other Specific Types". This group includes people with genetic defects of beta-cell function (this type of diabetes was formerly called MODY or maturity-onset diabetes in youth) or with defects of insulin action; people with diseases of the exocrine pancreas, such as pancreatitis or cystic fibrosis; people with dysfunction associated with other endocrinopathies (e.g. acromegaly); and people with pancreatic dysfunction caused by drugs, chemicals or infections and they comprise less than 10% of DM cases (Baynes, 2015).

1.6 Diagnoses of Diabetes Mellitus

The starting point for living well with diabetes is an early diagnosis – the longer a person lives with undiagnosed and untreated diabetes, the worse the health outcomes are likely to be. Easy access to basic diagnostics for diabetes is therefore essential and diagnosis should be available in primary health-care settings (American Diabetes Association, 2000).

Type 1 diabetes often presents with symptoms that prompt the patient to contact health services –thirst, weight loss and copious urination. Type 2 diabetes often shows no

symptoms, and some patients contact health services because of a complication such as vision loss, heart attack or limb gangrene. Type 2 diabetes develops slowly and there is often a very long period of time in which the disease is present but undetected (American Diabetes Association, 2011).

The identification of patients with diabetes or pre-diabetes by screening allows for earlier intervention, with potential reductions in future complication rates (Baynes, 2015). The 1997 American Diabetes Association (ADA) recommendations for diagnosis of diabetes mellitus focus on Fasting Plasma Glucose (FPG), while WHO focuses on the Oral Glucose Tolerance Test (OGTT) (Gillett, 2009). A clinician draws blood after a night of fasting and measures an indicator of blood glucose to determine whether it falls within normal range. Diabetes mellitus can be screened for diagnosis by one of the 3 methods;

Random Blood Glucose Test: Diabetes is diagnosed on the basis of a single abnormal plasma glucose reading. When taken randomly at any time of the day, the levels are considered high if they are 11.1 mmol/L or higher (Gillett, 2009).

Fasting Blood Glucose (FBG): This is usually taken after an overnight fast. Blood glucose level of 7mmol/L (126mg/dL) or higher is considered positive for diabetes when other diabetic symptoms such as thirst, increased urination, recurrent infections, weight loss or drowsiness are present (Gillett, 2009).

Oral Glucose Tolerance Test (OGTT): This blood test evaluates body's response to glucose and require the patient to fast for at least eight hours but not more than sixteen hours (overnight fast). A fasting blood sugar level is measured, then 75 gm glucose or any sugary drink is given to the patient and the blood sugar levels are tested periodically (every 30 minutes to 1 hour) for 2 hours. A blood sugar level less than 140 mg/dL (7.8 mmol/L) is normal, 140 gm/dL – 199 mg/dL (7.8 mmol/L -11.0 mmol/L) indicates prediabetes, and a reading of 200 mg/dL (11.1 mmol/L) or higher after 2 hours indicates diabetes (Mayo Foundation for Medical Education and Research, 2016). An abnormal screening test result should prompt clinician to repeat the same test or perform another test to confirm the diagnosis.

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1.7 Management of Diabetes Mellitus

Currently, the goal of diabetes management is to avoid or minimize chronic diabetic complications, as well as to avoid acute problems of hyperglycaemia or hypoglycaemia. Adequate control of diabetes leads to lower risk of complications associated with unmonitored diabetes including kidney failure (requiring dialysis or transplant), blindness, heart disease and limb amputation. The most prevalent form of medication is hypoglycaemic treatment through either oral hypoglycaemics and/or insulin therapy (Tuomilehto, *et al.*, 2001).

The management of diabetes mellitus includes; diet, physical exercise, insulin therapy and oral hypoglycaemic agents such as sulphonylureas, Biguanides, thiazolidinediones, alpha glucosidase inhibitors and Metformin. These drugs work in either two ways: They improve sensitivity of body cells to insulin or Reduce synthesis of glucose in the liver. If a person with Diabetes mellitus cannot secrete enough insulin, the patient must take daily injections of insulin (American Diabetes Association, 2000).

Patients with type 1 diabetes mellitus require direct injection of insulin as their bodies cannot produce enough (or even any) insulin. As of 2010, there is no other clinically available form of insulin other than injection for patients with type 1: injection can be done by insulin pump, by jet injector, or any of several forms of hypodermic needle. Non-injective methods of insulin have been unattainable as the insulin protein breaks down in the digestive tract (Mealey, 2006). For type 2 diabetics, diabetic management consists of a combination of diet, exercise, and weight loss, in any achievable combination depending on the patient. Obesity is very common in type 2 diabetes and contributes greatly to insulin resistance. Weight reduction and exercise improve tissue sensitivity to insulin and allow its proper use by target tissues (Mealey, 2006).

Diet is an important component of diabetes treatment. To maintain near-normal blood glucose levels, diet is designed to deliver the same amount of carbohydrate each day, spaced evenly throughout the day. People with diabetes should follow most of the same dietary guideline recommended for those without diabetes. The difference is that people with diabetes mellitus may need to eat less

carbohydrate, slightly more fat or protein to help regulate blood glucose level (American Diabetes Association, 2000).

2.0 The Concept of Nutrition

Nutrition is the science that interprets the interaction of nutrients and other substances in food in relation to maintenance, growth, reproduction, health and disease of an organism. It includes food intake, absorption, assimilation, biosynthesis, catabolism and excretion. The diet of an organism is what it eats, which is largely determined by the availability, the processing and palatability of foods. A healthy diet includes preparation of food and storage methods that preserve nutrients from oxidation, heat or leaching, and that reduce risk of foodborne illness. A poor diet may cause health problems, causing deficiency diseases such as blindness, anemia, scurvy, preterm birth, stillbirth and cretinism; health-threatening conditions like obesity and metabolic syndrome; and such common chronic systemic diseases as cardiovascular disease, diabetes, and osteoporosis. A poor diet can cause the wasting of kwashiorkor in acute cases, and the stunting of marasmus in chronic cases of malnutrition (Whitney and Sharon, 2013).

A nutrient is a component in foods that an organism uses to survive and grow. Macronutrients provide the bulk energy an organism's metabolic system needs to function while micronutrients provide the necessary cofactors for metabolism to be carried out. Both types of nutrients can be acquired from the environment. Micronutrients are used to build and repair tissues and to regulate body processes while macronutrients are converted to, and used for, energy. Methods of nutrient intake are different for plants and animals. Plants take in nutrients directly from the soil through their roots and from the atmosphere through their leaves. Animals and protists have specialised digestive systems that work to break down macronutrients for energy and utilize micronutrients for both metabolism and anabolism (constructive synthesis) in the body (Whitney and Sharon, 2005).

A healthy and balanced diet should provide enough calories for the daily energy requirement to maintain or achieved reasonable body weight, to provide for the needs of pregnant and lactating women, to allow for normal growth in children and adolescents and to satisfy the needs of aged people.

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2.1 Types of Nutrients

Nutrients can be classified in two ways;

Classification based on their synthesis: based on this classification, nutrients can be classified into essential and non-essential nutrients. These are those that cannot be synthesised in the body (either at all, or in sufficient quantities), and so must be consumed by an organism from its environment. For humans, these include essential fatty acids, essential amino acids, vitamins, and certain dietary minerals. Oxygen and water are also essential for human survival, but are generally not considered "food" when consumed in isolation. There are no "essential carbohydrates", animals can synthesise all the types of carbohydrates needed for growth. Humans can derive energy from a wide variety of fats, carbohydrates, proteins, and simple chemicals such as ethanol and acetic acid (John *et al.*, 2009). The nonessential nutrients are those nutrients that can be synthesized in the body; they may often also be absorbed from consumed food. The majority of animals ultimately derive their essential nutrients from plants. Interest has recently increased in phytochemicals, which include many non-essential substances which may or may not have health benefits (John *et al.*, 2009).

2. Classification based on the quantity required: nutrients are classified into macronutrients and micronutrients.

2.1.1 Macronutrients

Macronutrients are defined as a class of chemical compounds which humans consume in the largest quantities (must be above a threshold amount) and which provide humans with the bulk of energy. The macronutrients are carbohydrates, fiber, fats, protein, and water (Fuhrman, 2014). The macronutrients (excluding fiber and water) provide structural material (amino acids from which proteins are built, and lipids from which cell membranes and some signaling molecules are built) and energy. Carbohydrates and proteins provide 17 kJ approximately (4 kcal) of energy per gram, while fats provide 37 kJ (9 kcal) per gram (Jeffrey, 2006), though the net energy from either depends on such factors as absorption and digestive effort, which vary substantially from instance to instance. Vitamins, minerals, fiber, and water do not provide energy, but are required for other reasons (Berg *et al.*, 2002).

2.1.1.1 Carbohydrate

Glucose is an essential metabolic substrate of all mammalian cells. D-glucose is the major carbohydrate presented to the cell for energy production and many other anabolic requirements. Glucose and other monosaccharides are transported across the intestinal wall to the hepatic portal vein and then to liver cells and other tissues. There they are converted to fatty acids, amino acids, and glycogen, or are oxidized by the various catabolic pathways of cells (Bjornholm and Zierath, 2005). Most tissues and organs, such as the brain, need glucose constantly, as an important source of energy. The low blood concentrations of glucose can cause seizures, loss of consciousness, and death. On the other hand, long lasting elevation of blood glucose concentrations, can result in blindness, renal failure, vascular disease, and neuropathy. Therefore, blood glucose concentrations need to be maintained within narrow limits. The process of maintaining blood glucose at a steady-state level is called glucose homeostasis (Guettier and Gorden, 2006).

This is accomplished by the hormonal regulation of glucose uptake and hepatic glucose production. This maintenance is achieved through a balance of several factors, including the rate of consumption and intestinal absorption of dietary carbohydrate, the rate of utilization of glucose by peripheral tissues and the loss of glucose through the kidney tubule and the rate of removal or release of glucose by the liver and kidney. To avoid postprandial hyperglycaemia (uncontrolled increases in blood glucose levels following meals) and fasting hypoglycaemia (decreased in blood glucose levels during periods of fasting), the body can adjust blood sugar levels by a variety of cellular mechanisms. Important mechanisms are conveyed by hormones, cytokines, and fuel substrates and are sensed through of cellular mechanisms (Bjornholm and Zierath, 2005).

2.1.1.2 Protein

Proteins consist of long chains of subunits called amino acids. As the name implies, each amino acid contains an amino group (NH₂) on one end of the molecule and a carboxylic acid group (COOH) on another end. There are approximately twenty different amino acids, each with a distinct structure and chemical properties that are used to build proteins (Mathews *et al.* 2000). When amino acids are joined, the hydrogen from the amino end of one amino acid

combines with the hydroxyl group of carboxylic end of another amino acid. The bond between adjacent amino acids is called a peptide bond, and the compound formed is called a peptide. Two amino acids bound together is called a dipeptide; three, a tripeptide (Manninen, 2002).

The most critical aspects of protein metabolism that occur in the liver are: Deamination and transamination of amino acids, followed by conversion of the non-nitrogenous part of those molecules to glucose or lipids. Several of the enzymes used in these pathways (for example, alanine and aspartate aminotransferases) are commonly assayed in serum to assess liver damage (Felig 2005).

2.1.1.3 Lipids

Fatty acid (FA) is the major component of almost all lipid classes (except sterols) and also serves as carbon sources for energy production through β -oxidation and oxidative phosphorylation on shortage of glucose. Triacylglyceride (TAG), one of the major neutral lipids, is composed of three fatty acids (FAs) esterified to a glycerol backbone. The three FAs influence the chemical and physical characteristics of TAG by the length of carbon chains and degree of desaturation. The esterification happens between three hydroxyl groups (-OH) on the glycerol molecule and carboxyl groups (-COOH) on each of the three FA molecules. The most important function of TAGs is to efficiently store energy in the form of FAs, thereby preventing the potential toxicity of excessive free fatty acids (FFAs). Assembled and secreted in mammal intestines and livers, TAGs are also important for transporting FAs to different tissues throughout the body (Mathews *et al.* 2000).

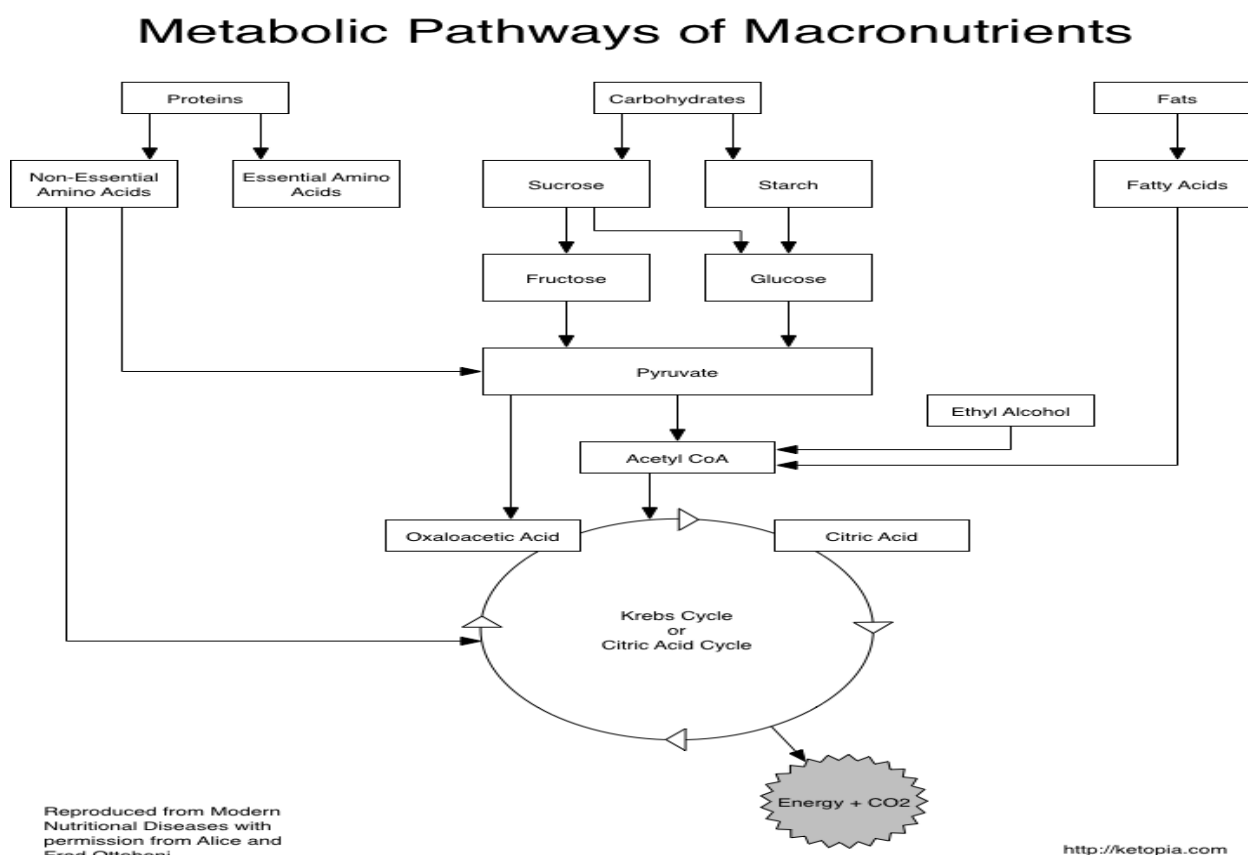


Figure 4: metabolic pathways of macronutrients
Source: Alice and Fred, 2013

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2.1.2 Micronutrients

Micronutrients are essential dietary components and play fundamental role in the prevention of disease. Micronutrients are vitamins and minerals that our bodies require in small quantities for specific functions. They most commonly function as essential coenzymes and cofactors for metabolic reactions and thus help support basic cellular reactions (i.e., glycolysis, the citric acid cycle, lipid and amino acid metabolism) required to maintain energy production and life (Belinda, 2001). Micronutrients have an array of biochemical functions which are fundamental in the homeostatic regulation of body function. Micronutrients can be obtained from different sources such as those that are intrinsic within the tissue matrix and fluid of edible plant and animals. Those that are from extrinsic sources will include nutrients that are added to foods such as with fortification of specific nutrients (thiamine to white flour, iodine added to salt) (Belinda, 2001).

2.1.2.1 Minerals

They are essential micronutrients that are required in diet in relatively small quantities from few micrograms to few milligrams (mg) per day by the body. Minerals that are considered essential are calcium, phosphorus, magnesium, sodium, potassium, iron, zinc, and fluoride (semi-essential). Trace elements considered essential are copper, chromium, manganese, molybdenum, selenium, and iodine (Rita, 2015)

2.1.2.2 Vitamins

They were first identified from the overt deficiency states that resulted from inadequate intakes causing conditions such as scurvy, beriberi and pellagra. Suboptimal intakes can be seen in sub-groups of populations and can cause adverse effects on health, particularly long term. Vitamins are subdivided into two groups: the fat soluble vitamins which include vitamins A, D, E and K and the water soluble vitamins which include B vitamins and vitamin C (Rita, 2015)

3.0 Nutritional Management of Diabetes

Nutrition is often said to be the cornerstone of diabetes care. The nutritional management of diabetes can affect long term health and quality of life. The goal for nutritional management is optimal metabolic control through a

balance between food intake, physical activity, and if necessary, medication to avoid complications (American Diabetes Association, 2008).

The major goal for diabetes care is to improve glycaemic control by balancing food intake with endogenous and/or exogenous insulin levels. For people with type 1 diabetes, insulin doses need to be adjusted to balance with nutritionally adequate food intake and physical activity. For individuals with type 2 diabetes, impaired glucose tolerance or impaired fasting glucose, attention to food portions and weight management combined with physical activity may help improve glycaemic control. Nutrition and all forms of diabetes management should be individualised (Smart *et al.*, 2009).

Evidence available from America from the Diabetes Control and Complications Trial in Type 1 diabetes (The Diabetes Control and Complications Trial Research Group, 1993) and also from the United Kingdom Prospective Diabetes Study in Type 2 diabetes (UK Prospective Diabetes Study (UKPDS) Group, 1998) suggests that normalisation of metabolic markers like blood glucose levels and management of blood pressure constitute key aims of the nutritional management of diabetes. Other aims of nutritional management of diabetes are:

1. Encourage appropriate eating behavior and healthy lifelong eating habits whilst preserving social, cultural and psychological well-being;
2. Three balanced meals a day, with appropriate healthy snacks (if necessary), will supply all essential nutrients, maintain a healthy weight, prevent bingeing and provides a framework for regular monitoring of blood glucose levels;
3. Provide sufficient and appropriate energy intake and nutrients for optimal growth, development and good health;
4. Achieve a balance between food intake, metabolic requirements, energy expenditure and insulin action profiles to attain optimum glycaemic control;
5. Prevent and treat acute complications of diabetes such as hypoglycaemia, hyperglycaemic crises, illness and exercise-related problems;

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3.1 Dietary Recommendation for Diabetics

3.1.1 Macronutrient Requirements

The macronutrient requirements for diabetic patients are carbohydrate, protein and lipid.

3.1.1.1 Carbohydrates

Carbohydrates are the body's main energy source; 50% - 55% of the daily caloric intake should be provided by them (American Diabetes Association, 2002). Based on the individual's glucose needs, weight status, lipid profile, and eating habits, carbohydrate intake should be individualised. It is important that people with either type 1 or type 2 diabetes consume the right amount of carbohydrates, as they are the primary energy source for the central nervous system which depends on blood glucose. Carbohydrates also have the role of 'protein sparer', preventing the use of proteins for energy purposes, allowing them to perform their real role in tissue building and as metabolic primers for fat metabolism. The amount of carbohydrates in the diet regulates the levels of the intermediate products of fat metabolism, ketones. If the amount of carbohydrates is too low or unavailable, fat is oxidized for energy purposes with an increase of ketones resulting in keto-acidosis, which is a common problem in patients with type 1 diabetes (Arnoldi, 2004).

Simple sugars are thought to be absorbed quickly, thus rapidly increasing the level of blood sugars and so were forbidden. However, the ADA Position Statement 2002 has concluded that for people with diabetes it is the total amount of carbohydrates in meals and snacks, rather than the type, that determines the glycaemic response. Sugars, however, should be restricted in the diet, as a high intake usually increases triglyceride levels in the blood and may also contribute to the development of dental caries (American Diabetes Association, 2002). Moreover, sugars are usually associated with high-calorie foods, which should be limited in order to maintain a healthy body weight. Their amount should not be more than 10 per cent of the daily total energy intake. Polysaccharides are preferred, especially those containing fibre, since the more fibre food contains, the more slowly it is digested, raising blood sugar levels at a slower rate (European Diabetes Policy Group, 1999).

A system for classifying carbohydrates, known as the glycaemic index (GI), measures the effect that a food has on blood sugar levels. Foods that have a high glycaemic index cause a rapid and strong rise in blood sugar levels; diets filled with these foods have been linked to an increased risk for both diabetes and heart disease. Various factors, including the degree of processing, physical form and fibre content, determine a food's glycaemic index. Therefore, the glycaemic index relates to the quality and not the quantity of carbohydrates consumed (Arnoldi, 2004).

Foods that contain complex carbohydrates, such as potatoes, quickly raise blood sugar levels, whereas foods that contain simple carbohydrates, such as whole fruit, raise blood sugar levels more slowly. Fruits and milk have been shown to have lower glycaemic responses than most starches (Leonid, 2009). Sucrose produces a response similar to common starches like rice, potato, and bread. Although various starches per gram of carbohydrate do have different effects on blood sugar, for purposes of dietary planning it is the total amount of carbohydrate consumed rather than the source that merits primary consideration (Leonid, 2009).

Another dietary sugar, fructose, produces smaller blood glucose increase than sucrose and has been proposed as a "natural" sweetener. The moderate consumption of fructose-containing foods has presented no problem, but when fructose is consumed in large amounts, up to 20% of the calories, it is known to adversely affect blood lipids (Leonid, 2009). The sweeteners that have calories from carbohydrate like fruit juice concentrates, molasses, honey, and corn syrup have direct effects on blood sugar similar to sucrose and offer no advantage to persons with diabetes. Nonnutritive sweeteners are encouraged for people with diabetes to add increased variety to their food choices (American Diabetes Association, 2008).

The daily intake of carbohydrates will depend upon the type of exercise programme, the amount and timing of daily insulin requirement, the use of an insulin pump, the need to lose weight or for a special diet to reduce cholesterol or blood pressure levels. Food such as honey, syrup, jam, jelly, candy, sweet rolls, regular gelatin, cake with icing, and pie should be avoided, as should soft drinks. The consumption of the so-called products for diabetics, such as chocolate, should also be restricted as they contain excess calories in fat.

Citations: Muhammed D, Adebisi YH, Odey BO, Ibrahim J, Hassan ON, Ugwunnaji PI, and Berinyuy E.B. (2021). Nutritional Management of Diabetes Mellitus: An appraisal of the role of Medicinal plants. *AROC in Natural Product Research*. 01(01), 001-027

3.1.1.2 Fibre

For years fiber has received much attention for its disease prevention effects in the general population. Fibre is the indigestible part of plant foods (Nuttall, 1993). There are two types of fibre: soluble and insoluble. Recently the literature is replete with articles demonstrating improved blood glucose management in both type 1 and type 2 diabetes with high-fiber diets (Wylie-Rosett *et al.*, 2004).

Fiber-rich foods such as beans or cereals with 5 or more grams of fiber per serving and fruits and vegetables are emphasised due to their nutrient content (Liese *et al.*, 2005). Both soluble and insoluble fibers are encouraged in amounts similar to the recommendations for the general population (20–35 g or 14 g/1000 cal) (Liese *et al.*, 2005). Soluble fibres, such as gums and mucilages, form gels with pectin or gums in the intestine and thus slow the rate of nutrient absorption, helping to reduce postprandial glucose levels. Insoluble fibres, such as lignins, celluloses and hemicelluloses, increase bulk and decrease transit time. Both types are useful as they prolong gastric emptying, shorten intestinal transit time, prevent constipation and bind cholesterol, thus limiting its absorption. Food rich in fibre include vegetables, fruits, whole wheat, bran, cereals and legumes. An adequate diet should provide 30 grams of fibre a day (Arnoldi, 2004).

3.1.1.3 Proteins

At the current time no data are available to indicate that the protein needs of persons with diabetes are different from the dietary reference intake (DRI) for the general population, 15-20 % of total caloric intake, 0.8-1 g/kg body weight (Franz, 2002^b). This amount should increase during pregnancy, breast feeding, infancy and childhood, illness and diseases.

Proteins, unlike fats and carbohydrates, contain nitrogen. Proteins are normally not energy suppliers, but in cases of stress, diseases or fasting, when there is either an increased need of energy or the caloric intake is not sufficient to cover the body needs, they are used to provide energy. Thus, a diet with the right amount of carbohydrate has a protein-sparing effect. However, dietary protein intake must be balanced, as large amounts can cause gout or kidney stones in some individuals, as purine breaks down to uric acid, and high concentrations can crystallise in the kidneys and joints.

Moreover, excess nitrogen is a burden for the kidneys and animal food proteins are also rich in fat, especially saturated types, which cause cardiovascular disease and obesity. According to the ADA dietary guidelines, the intake of protein in the usual range does not appear to be associated with the development of diabetic nephropathy, although it is advisable to avoid intakes superior to 20 per cent of the total daily energy (American Diabetes Association, 2011).

3.1.1.4 Lipids

As people with diabetes have an increased risk of heart and blood vessel diseases, a normal to low-fat diet is advisable in order to reach and maintain good weight and health. As part of a healthy diet, 30% of daily calories should come from fats and of these less than 10% should be saturated fat, less than 10% polyunsaturated fat and 10-15% monounsaturated fat (American Diabetes Association, 2008). However, a diet with a low intake of fat can be dangerous since not only is fat a major source of energy for the body as it supplies 9kcal/g, it is also the source of essential fatty acids (EFA), such as linoleic, linolenic and arachidonic acids. These are called 'essential' as the body cannot synthesize them and they are needed for a number of key functions such as transport and metabolism of cholesterol, precursors of prostaglandins and others. Among the EFA, omega-3 fatty acids found in fish, especially salmon, mackerel and herring, have been shown to lower plasma triglyceride levels in patients with type 2 diabetes (American Diabetes Association, 2008).

Monounsaturated fat is a type of unsaturated fat that lowers blood low density lipoprotein (LDL) cholesterol levels (Berry *et al.*, 1991). It is also resistant to oxidation. This is important since oxidation enables cells in arteries to absorb fats and cholesterol and helps accelerate the formation of plaques. This type of fat is found in olive oil, avocados, canola oil, etc. In people with diabetes a diet higher in monounsaturated fats and lower in carbohydrates has been shown to reduce postprandial glycaemia and triglyceridaemia, but it is not recommended for someone with type 2 diabetes as it may cause weight gain (Lichtenstein and Deckelbaum, 2001). Monounsaturated fats and carbohydrate intake should be based on dietary goals and metabolic profile of each individual.

The effects of polyunsaturated fat on subjects with diabetes have not been well studied. Compared with

monounsaturated fat, this vegetable fat lowers total blood cholesterol but is susceptible to oxidation, which as previously described may ultimately lead to coronary artery disease. Polyunsaturated fats are found in cottonseed, soybean, sunflower and safflower oils. Saturated fat is an animal fat that raises total blood cholesterol and increases the risk for coronary artery disease (National Cholesterol Education Program, 2001). It is found in hydrogenated vegetable fats, coconut and palm oils, cocoa butter, meat fat, whole milk, butter cream and fatty cheeses. Cholesterol blood levels should be kept to <200 mg/dL and the daily intake should be limited to 300 mg/day. According to the ADA recommendations, diabetics with dyslipidaemia, such as those with high LDL cholesterol, should lower saturated fats to less than 7 per cent and dietary cholesterol to 200 mg/day and increase soluble fibre to 10-25 g/day, together with physical activity (American diabetes association, 2011). Also plant sterol and stanol esters have been shown to have a good effect on blocking the absorption of total and LDL cholesterol (Lichtenstein and Deckelbaum, 2001).

A healthy diet should be rich in fish, poultry (without skin), lean cuts of beef, lamb, veal, pork, skimmed or low-fat milk, low-fat cheeses, low-fat yogurt, olive oil, rice, pasta, bread, whole-grain cereals, legumes, vegetables and fruits, while intake of foods such as animal fats, bacon, sausages, commercially baked goods, cream, cheeses, butter, salad dressings made with cream and eggs and junk food such as potato chips, cookies should be cut down. Particular attention must be made to margarine or shortening, which contain hydrogenated vegetable oils, which consequently form trans-fatty acids. These are highly related with artery disease, raising blood cholesterol and lowering HDL levels (Arnoldi, 2004).

3.1.2 Micronutrient requirements

As the general population, people with diabetes have no need for vitamin and mineral supplementation when the dietary intake is adequate. Only in certain cases such as illness, stress or pregnancy is a higher intake required.

3.1.2.1 Vitamins

Many physicians prescribe a pill containing the reference dietary intake (RDI) of the established vitamins as an "insurance policy." For the elderly with reduced energy

intake, a multivitamin supplement is commonly given. With the increased risk for heart disease and its adverse outcome in persons with diabetes, antioxidants may be prescribed, but there is little evidence that this practice is beneficial (Yusuf *et al.*, 2000).

High doses of dietary antioxidants such as vitamins C and E, beta-carotene and other carotenoids have been prescribed to people with diabetes (Ylonen *et al.*, 2003). Antioxidants are substances that neutralize the action of free radicals, which are molecules that damage cells, and increase the risk of cancer and heart diseases. In this case, they are thought to protect LDL and PUFA from oxidation. Supplementary intakes of vitamin D seem to lead to an improvement in lipid metabolism in type 2 diabetes sufferers, especially in combination with physical exercise. Experimental research produced evidence that vitamin D can help prevent the destruction of insulin-producing pancreatic beta cells and thus combat the onset of type 1 diabetes. It is assumed that this is due primarily to the immunomodulatory action of the vitamin via T-helper cells and to the reduction of pro-inflammatory cytokines (Zipitis and Akobeng, 2008).

The antioxidant status of a diabetic can be significantly improved by administration of vitamin C, which reduces protein glycosylation by competitively displacing glucose from the amino groups of proteins. In this way the vitamin prevents endothelial damage induced by glycosylated reaction products and improves endothelium function (Meredith *et al.*, 2014). Vitamin C also reduces aldose reductase (figure 5) and hence slows the intracellular accumulation of the sugar alcohol sorbitol, which can cause

damage to nerves, eyes and kidneys (Cunningham *et al.*, 1994). In a randomized controlled study with type 2 diabetics, adjunctive administration of 2 x 500 mg vitamin C per day for a period of four months led to a significant decrease in insulin resistance, HbA1C values and plasma levels of total cholesterol, LDL cholesterol and triglyceride, compared with placebo (Paolisso *et al.*, 1995). Further, supplementary administration of vitamin C appears to enhance the metabolic regulatory activity of the anti-diabetic medication metformin (Dakhale *et al.*, 2011) and to have a positive effect on complications like depression (Mazloom *et al.*, 2013) and periodontitis (Gokhale *et al.*, 2013) in diabetics.

As a highly effective antioxidant, vitamin E protects enzymes and hormones, as well as the polyunsaturated fatty acids of biological membranes and LDL, against oxidation by oxygen radicals. During this process vitamin E is oxidized and must be regenerated by vitamin C or flavonoids. In this way the vitamin combats the oxidative degradation of fatty acids (lipid peroxidation) and in particular the oxidative modifications of LDL that contribute to the incidence of atherosclerosis. Moreover, vitamin E lowers thrombocyte aggregation and hence the risk for thrombosis. In addition, by inhibiting enzymes, vitamin E slows inflammatory processes and the proliferation of connective tissue in the blood vessels and therefore reduces the threat or advance of diabetic complications (Montero *et al.*, 2014). However, according to the ADA dietary guidelines (2002), routine supplementation of the diet with antioxidants is not advised because of uncertainties related to long- term efficacy and safety.

As coenzymes the B vitamins play a central role in carbohydrate, protein and lipid metabolism. A diabetic metabolic status is characterized by both higher requirements and increased renal elimination of B vitamins, especially when the diabetes is not well managed. Studies indicate that the majority of type 1 and type 2 diabetics have inadequate supplies of vitamin B1 and impaired thiamine metabolism (Thornalley *et al* 2007). A tissue-specific vitamin B1 deficiency (e.g., in the kidneys) can increase the risk of vascular complications (e.g., nephropathy) (Rabbani and Thornalley, 2011), and ensuring sufficient provision should be a focus of therapy. Further, vitamin B1, B6 and B12 support nervous system functions and helps prevent diabetic neuropathies. A lack of folic acid and/or vitamin B12 leads to impaired metabolism of the amino acid methionine and is frequently accompanied by elevated plasma homocysteine concentrations. Elevated homocysteine levels are regarded as an independent risk factor for stroke, heart attack, dementia and macular degeneration. Compared to non-diabetics, diabetics are three to five times more likely to suffer a stroke (Montero *et al.*, 2013).

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3.1.2.2 Minerals

The role of microelements, zinc and chromium in glycaemic control is difficult to determine, as they are present in minute amounts and their deficiency is not easily assessed. Of the minerals, calcium supplementation is frequently advised, particularly after menopause for women (with diabetes), since dietary calcium may not be sufficient, but as for all the other minerals the recommendations are similar to those for the general population. The ADA dietary guidelines recommend the intake of 1.0-1.5 mg/day of calcium to older people to reduce the risk of osteoporosis. Chromium has been encouraged because of its positive metabolic role particularly in type 2 diabetes. Chromium's actions have been attributed to an increase in the number of insulin receptors, increased binding of insulin to the insulin receptor, and increased activation of the insulin receptor in the presence of insulin (Belinda, 2001). Chromium supports the activity of insulin and sensitizes the pancreatic beta cells, promoting the expression of insulin. Chromium deficiency can lead to reduced glucose tolerance and disordered glucose utilization (Anderson, 1998). Studies showed that supplementation with chromium (III) could bring about an improvement in glucose and/or lipid metabolism, in particular in diabetes sufferers with insufficient chromium intake (Kleefstra *et al.*, 2007). A meta-analysis of numerous clinical studies with type-2 diabetes patients revealed that supplementary administration of chromium (III) improved fasting blood sugar, as well as lipid metabolic status (Cefalu *et al.*, 2010).

Magnesium is acknowledged for its role in insulin sensitivity and its deficiency can contribute to carbohydrate intolerance. One extensive epidemiological study showed that the risk for both type 1 and type 2 diabetes mellitus and for metabolic syndrome as diabetes precursor rises as magnesium intake falls. According to a meta-analysis the risk for diabetes was reduced by 15% per 100 mg of additional magnesium intake (Larsson and Wolk, 2007). However, only when low serum magnesium levels can be established is repletion with magnesium appropriate.

Sodium is the mineral that receives much attention by both the medical profession and the general public (Leonid, 2009). According to the ADA the amount of sodium in the diet should be limited as it helps reduce blood pressure and the tendency to retain fluids. The main source of sodium in the diet is common table salt. However, as natural foods contain the right amount for the body's need, the general consumption of salt is too high and the recommended goal is to reduce sodium intake to 2400 mg or sodium chloride (salt) to 6000 mg/day (Arnoldi, 2004). Snacks, pickles, bacon, sauces, olives, chips and processed foods in general

contain too much salt and should be cut down. (Loria *et al.*, 2001)

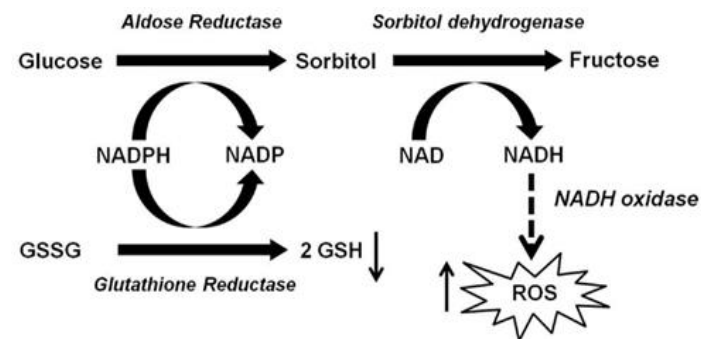


Figure 5: Role of aldose reductase in hyperglycaemia-induced oxidative stress. GSSG-glutathione disulphide, GSH-glutathione, ROS – reactive oxygen species

Source: Tang *et al.*, 2012.

Table 1: Classification of glycaemic index

Class	GI range (%)	Examples
Low GI	55 or less	Beans (black, white, pink, almond, walnut, peanut), most intact whole grains (millet, rice, durum, wheat), fructose, tagatose
Medium GI	56-69	Sucrose, banana, unpeeled boiled potato, strawberry, grape, basmati rice.
High GI	70 and above	Galactose, maltose, maltodextrin, cornflakes, extruded breakfast meal, white bread (only wheat endosperm), white rice (only rice endosperm)

Source: Foster-Powell *et al.*, 2002

There is no sufficient information available to justify the recommendation of herbs and supplements for diabetes care. As with the general population, caution should be advised on the use of supplements due to their lack of standardized ingredients and possible drug interactions (American Diabetes Association, 2008).

3.2 Alcohol Intake

Type 1 diabetes patients must pay particular attention to alcohol, as it can cause some problems such as lowering blood

sugar levels. Otherwise patients with diabetes should take the same precautions as the general population. In fact according to the ADA dietary guidelines (2002), men should have no more than two alcohol-containing drinks per day and no more than one drink for women (there should be no more than 15 g of alcohol per drink). Alcohol should be drunk at mealtimes and never in exchange for regular food in order to balance calories. Alcohol gives 7kcal/g, stimulates appetite and also has an adverse effect on self-control, so people with type 2 diabetes should limit their alcohol consumption to special occasions only, especially when on a weight-loss programme (Arnoldi, 2004). Alcohol also increases serum triglyceride

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levels. In certain conditions such as pregnancy or for medical problems such as pancreatitis, neuropathy and severe hypertriglyceridaemia, total abstinence is advisable. The use of alcohol is also contraindicated with certain medications, particularly metformin which is frequently prescribed for type 2 diabetes, since alcohol can increase the effects of metformin on lactate metabolism which increases the risk of lactic acidosis (Howard *et al.*, 2004).

3.3 Dietary Recommendations for Pregnant and Lactating Women

Nutrients and energy needs increase dramatically during pregnancy and lactation. The time when it is necessary to increase the amount of a specific nutrient varies considerably. Increases in energy needs relate to increased maternal metabolism, added maternal tissues, and growth of the foetus and placenta. Caloric needs during pregnancy may be from 400 to 600 kcal more per day than for the same period in the non-pregnant state (Guthrie and Guthrie, 2009).

Pregnancy is not the time to lose body fat by going on a diet. Calorie restriction is not usually implemented during pregnancy, and there is even controversy about whether women who are obese should gain some weight for optimal fetal development. Restriction of carbohydrates to 35%–40% of calories has been shown to decrease maternal glucose levels and improve maternal and foetal outcomes. A gradual, steady weight gain is optimal and is as important as the total weight gained (Reader *et al.*, 2006).

Intakes of nutrients such as protein, iron, folic acid, minerals, and fat-soluble and water-soluble vitamins increase in diabetic pregnant women. The current American nutritional guidelines for diabetic women during pregnancy and lactation are for a daily protein intake of 0.75 g/kg plus an additional 10 g/day (American Diabetes Association, 2004). Protein needs to be increased to assist with growth, particularly of the maternal breasts, uterus, extracellular fluid, and foetus (Guthrie and Guthrie, 2009).

The goals of dietary management in pregnancy include meeting nutritional requirements while eliminating hyperglycaemia. Distribution of meals and snacks must coincide with peaks of insulin given by injection. Because of nausea and hyperemesis in early pregnancy, the size of snacks may begin to equal the size of meals. Three meals

and three to four snacks per day with equal timing between periods of food intake are optimal. Eating every 3–4 hour may become necessary to prevent ketonemia. Daily calories of 36–50 kcal/kg of ideal body weight should meet a majority of the previously mentioned nutrient needs. This increase may average to about 300kcal/day above the pre-pregnancy meal plan. Alcohol consumption during pregnancy is strongly discouraged because of fetal alcohol syndrome (Djelmis *et al.*, 2005).

On a side note, one study found that milk consumption during pregnancy was associated with a higher birth weight for gestational age which correlated with the protein rather than the fat in the milk (Olsen *et al.*, 2007). What probably is more important, at this point, is that women with diabetes who are obese and are pregnant not only beget obesity in their children, but also a tendency for a multitude of complications (Reece, 2008).

Breastfeeding is recommended for women with pre-existing diabetes and GDM; however, successful lactation requires planning and coordination of care. Breastfeeding lowers blood glucose, and insulin-treated women may require a carbohydrate-containing snack either before or during breastfeeding. Energy requirements during the first 6 months of lactation are 200 calories in addition to the pregnancy meal plan. However, an energy intake of 1800 kcal/day usually meets the nutritional requirements for lactation and this may allow for a gradual weight loss in overweight type 2 individuals or those who had GDM (Djelmis *et al.*, 2005).

3.4 Dietary Recommendations for Infant and Young Children

Dietary recommendations for children with diabetes are based on healthy eating recommendations suitable for all children and adults (National Institute for health and Care Excellence, 2004) and therefore the whole family. Nutritional advice must be adapted to cultural, ethnic and family traditions and the psychosocial needs of the individual child. A specialist pediatric dietician with experience in childhood diabetes should be available as part of a pediatric interdisciplinary diabetes care team to provide education, monitoring and support to the child, parents, careers, extended family, nursery, school teachers, and babysitters.

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There is international agreement that carbohydrate should not be restricted in type 1 diabetes as it may have deleterious effects on growth. Carbohydrate should account for 50–55% of total daily energy intake. Sucrose and sucrose-containing food should be eaten in the context. For children above 1 year an amount of 2.8–3.4 gms per megajoule is recommended (Smart *et al.*, 2009). Alternatively, for children above 2 years; age in years + 5 = grams of fiber per day (Williams, 2006) is recommended. Intake of a variety of fiber containing foods such as legumes, fruit, vegetables and wholegrain cereals should be encouraged. Soluble fiber in vegetables, legumes and fruit may be particularly useful in helping to reduce lipid levels (Smart *et al.*, 2009).

Over recent decades surveys have shown children and young people with diabetes consume fat and saturated fat above dietary recommendations (Virtanen *et al.*, 2001) and this situation has not changed (Helgeson *et al.*, 2006; Mayer-Davies *et al.*, 2006; Overby *et al.*, 2007). The intake of total fat, saturated fat, and trans-fatty acids should be decreased (Franz, 2002^a) while Monounsaturated fatty acids (MUFA) and polyunsaturated fatty acids (PUFA) can be used as substitutes to keep lipid intake within recommended ranges or to improve the lipid profile (Franz, 2002^a).

Total daily energy intake of protein of 10–15% is recommended. Intake of protein decreases during childhood from approximately 2 g/kg/day in early infancy to 1 g/kg/day for a ten year old and to 0.8–0.9 g/kg/day in later adolescence (Smart *et al.*, 2009). Sources of vegetable protein such as legumes should be encouraged. Sources of animal protein also recommended include fish, lean cuts of meat and low fat dairy products (Smart *et al.*, 2009).

Children with diabetes have the same vitamin and mineral requirements as other healthy children (Smart *et al.*, 2009). Optimum vitamin, mineral and antioxidant intake should be maintained for general health and cardiovascular protection. Many fresh fruits and vegetables are naturally rich in antioxidants (tocopherols, carotenoids, vitamin C, and flavonoids) and are strongly recommended for young

people with diabetes. Supplements of vitamins, minerals or trace elements are not usually recommended unless nutritional assessment confirms a specific deficiency. Supplements such as vitamin D for young children are recommended in some countries following the national guidelines for healthy children (Franz, 2002^a).

4.0 Appraisal of the role of Medicinal plants in treatment of diabetes

Medicinal plants play a significant role in the treatment of several diseases and have been reported to be safe for use as oral remedy (Bashir *et al.*, 2015, Lawal *et al.*, 2015, Lawal *et al.*, 2016, Lawal *et al.*, 2017, Yusuf *et al.*, 2018a, 2018b, Umar *et al.*, 2019, Yusuf *et al.*, 2020, Adeshina *et al.*, 2020). Traditional plants are reported to have significant anti-diabetic properties with no harmful side effects. They are rich sources of anti-diabetic compounds such as flavonoids, alkaloids, phenolic and tannins that improve the efficiency of pancreatic tissues by increasing the insulin secretion or decreasing the intestinal absorption of glucose (Bindu & Narendhirakannan, 2019; Kooti *et al.* 2016).

Literature suggests that there are approximately 410 experimentally proven medicinal plants with anti-diabetic properties out of which the complete mechanism has been studied only for 109 plants. Several medicinal plant extracts have been shown to modulate the metabolic pathways such as glycolysis, gluconeogenesis, Krebs cycle, glycogen synthesis and their degradation, synthesis and release of insulin, cholesterol synthesis, carbohydrate metabolism and absorption (Prabhakar and Doble 2008).

We presented the list of medicinal plants that are known to possess anti-diabetic properties. These plants are traditionally used by various tribal people for the treatment of several ailments. Pharmacological studies have reported their hypoglycemic, anti-hyperglycemic, insulin mimicking, anti-lipidemic properties and hence when administered in proper dosages can be beneficial for ameliorating the various complications associated with diabetes mellitus (Bindu & Narendhirakannan, 2019).

Table 2: Medicinal plants with anti-diabetic properties

Adapted from Bindu & Narendhirakannan (2019).

S. no.	Botanical name	Common name(s)	Family	Part(s) used for the study	Activities
1	Abelmoschus esculentus (L.) Moench	Lady's finger	Malvaceae	Seeds, fruit	Antidiabetic, antioxidant and antihyperlipidemic
2	Acacia nilotica (L.) Delile	Gum arabic tree	Leguminosae	Leaves	Antioxidant, antidiabetic, antihyperlipidemic
3	Allium cepa L.	Onion	Amaryllidaceae	Bulb	Hypoglycemic, anti-diabetic, anti-hyperlipidemic
4	Aloe vera (L.) Burm.f.	Aloe	Asphodelaceae	Leaves	Anti-hyperglycemic, anti-hypercholesterolemic, hypoglycemic, hypolipidemic
5	Annona squamosa L.	Sugar apple	Annonaceae	Leaves, fruit pulp	Hypoglycemic, anti-diabetic, anti-lipidemic
6	Artemisia pallens Wall. ex DC.	Artemisia	Asteraceae	Aerial parts	Hypoglycemic
7	Averrhoa bilimbi L.	Bilimbi	Oxalidaceae	Leaves, roots, fruit	Anti-hyperglycemic, anti-hyperlipidemic, hypoglycemic, anti-diabetic
8	Azadirachta indica A. Juss	Neem tree	Meliaceae	Leaves, seeds	Hypoglycemic, β -cell regeneration
9	Berberis vulgaris L.	Barberry	Berberidaceae	Root, fruit	Hypoglycemic, anti-diabetic
10	Beta vulgaris L.	Beet	Amaranthaceae	Leaves, fruit	Anti-hyperglycemic, hypoglycemic
11	Biophytum sensitivum (L.) DC.	Little tree plant	Oxalidaceae	Leaves	Hypoglycemic
12	Brassica juncea (L.) Czern.	Brown mustard	Brassicaceae	Seeds	Hypoglycemic
13	Bryophyllum pinnatum (Lam.) Oken	Life plant, miracle leaf	Crassulaceae	Leaves	Antidiabetic, hypoglycemic
14	Boerhavia diffusa L.	Spreading hogweed	Nyctaginaceae	Leaves	Anti-diabetic
15	Bombax ceiba L	Cotton tree	Bombacaceae	Leaves	Antihyperglycemic, antihyperlipidemic
16	Caesalpinia bonduc (L.) Roxb	Fever nut	Caesalpiniaceae	Seeds, root bark	Hypoglycemic, antihyperglycemic, antihypercholesterolemic, antihypertriglyceridemic
17	Cajanus cajan (L.) Millsp.	Pigeon pea	Fabaceae	Leaves, seeds	hypolipidemic
18	Camellia sinensis (L.) Kuntze	Tea plant	Theaceae	Leaves	Hypoglycemic
					Anti-hyperglycemic, antihyperglycemic and

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19	Capparis decidua (Forssk.) Edgew.	Karira	Capparidaceae	Fruit, flowers, bark	antidiabetic Anti-diabetic, hypolipidemic
20	Capsicum annuum L.	Capsicum	Solanaceae	Fruit	Insulinotropic
21	Centaurium erythraea Rafn	Common centaury	Gentianaceae	Leaves	Antihyperglycemic
22	Catharanthus roseus (L.) G.Don	Madagascar periwinkle	Apocynaceae	Leaves	Anti-diabetic, hyperlipidemic, anti-hyperglycemic
23	Citrullus colocynthis (L.) Schrad.	Bitter apple, wild gourd	Cucurbitaceae	Fruit, seeds, root	Hypoglycemic, antihyperglycemic
24	Clausena anisata (Willd.) Hook.f. ex Benth.	Horsewood	Rutaceae	Root	Hypoglycemic
25	Cocos nucifera L.	Coconut	Arecaceae	Spadix, drupe, coconut water	Anti-hyperglycemic, antiglycation, hypoglycemic
26	Cheilocostus speciosus (J.Koenig) C.D.Specht	Cane-reed	Costaceae	Root, leaves	Antihyperglycemic, hypoglycemic, hypolipidemic
27	Curcuma longa L.	Turmeric	Zingiberaceae	Rhizome	Antioxidant, antidiabetic, hepatoprotective
28	Cyclocarya paliurus (Batalin) Iljinsk.	Wheel wingnut	Juglandaceae	Bark	Hypoglycemic, anti-ti-hyperglycemic, anti-hyperlipidemic
29	Dillenia indica L.	Elephant apple	Dilleniaceae	Leaves	Antidiabetic, hypolipidemic
30	Eucalyptus globulus Labill.	Blue-gum	Myrtaceae	Leaves	Anti-hyperglycemic
31	Embelia ribes Burm.f.	False black pepper	Myrsinaceae	Berries	Anti-hyperglycemic, anti-diabetic
32	Eugenia uniflora L.	Brazilian cherry	Myrtaceae	Fruit	Antihyperglycemic, antidyslipidemic
33	Ficus benghalensis L.	Banyan tree	Moraceae	Bark, aerial roots	Anti-diabetic
34	Ficus religiosa L.	Sacred fig	Moraceae	Bark	Anti-diabetic
35	Helicteres isora L.	Indian screw tree	Sterculiaceae	Root	Antidiabetic, hypolipidemic, antihyperglycemic
36	Hibiscus rosa-sinensis L.	Chinese hibiscus, shoe black plant	Malvaceae	Flower, leaves	Hypoglycemic, antidyslipidemic
37	Ipomoea batatas (L.) Lam	Sweet potato	Convolvulaceae	Leaves, root	Hypoglycemic
38	Juniperus communis L.	Common juniper	Cupressaceae	Berries	Hypoglycemic
39	Lantana camara L.	Big-sage, wild-sage	Verbenaceae	Fruits	Anti-hyperglycemic, anti-diabetic
40	Lithocarpus polystachyus (Wall. ex	Sweet tea	Fagaceae	Leaves	Hypoglycemic

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41	A.DC.) Rehder Phyla nodiflora (L.) Greene	Capeweed	Verbenaceae	Whole plant	Anti-hyperlipidemic, anti- hyperglycemic
42	Mangifera indica L.	Mango	Anacardiaceae	Leaves, fruit	Anti-diabetic, hypoglycemic
43	Memecylon umbellatum Burm. f.	Iron wood	Melastomataceae	Leaves	Anti-diabetic
44	Morus alba L.	White mulberry	Moraceae	Fruit, leaves	Anti-hyperglycemic, anti- hyperlipidemic, anti- glycemic
45	Mucuna pruriens (L.) DC.	Velvet bean	Fabaceae	Seeds	Anti-diabetic
46	Musa × paradisiaca L.	Plantain	Musaceae	Stem, leaves, infructescence stalks, fruit	Antidiabetic, antihyperlipidemic, anti- hyperglycemic
47	Nelumbo nucifera Gaertn.	Indian lotus	Nelumbonacea	Seeds, leaves	Hypoglycemic, antihyperlipidemic
48	Olea europaea L.	Wild olive	Oleaceae	Leaves	Antihyperglycemic, hypolipidemic, hypoglycemic
49	Ophiopogon japonicus (Thunb.) Ker Gawl.	Dwarf lilyturf	Asparagaceae	Root	Anti-diabetic, hypoglycemic
50	Opuntia streptacantha Lem.	Prickly pear cactus	Cactaceae	Leaves (cladode)	Anti-hyperglycemic
51	Phyllanthus niruri L.	Stonebreaker	Phyllanthaceae	Aerial parts	Anti-diabetic
52	Picrorhiza kurroa Royle ex Benth.	Yellow gentian	Plantaginaceae	Rhizome	Anti-diabetic
53	Piper nigrum L.	Black pepper	Piperaceae	Fruit	Anti-hyperglycemic
54	Prosopis glandulosa Torr.	Honey mesquite	Fabaceae	Pods	Anti-diabetic
55	Psidium guajava L.	Guava	Myrtaceae	Fruit, leaves	Anti-hyperglycemic, anti- hyperlipidemic
56	Pterocarpus marsupium Roxb.	Indian kino tree	Fabaceae	Bark	Hypoglycemic, anti- hyperglycemic, inhibition of aldose reductase activity
57	Punica granatum L.	Pomegranate	Punicaceae	Leaves, fruit	Anti-diabetic, anti- hyperlipidemic, anti- glycation
58	Salacia reticulata Wight	Marking nut tree, kotalahimbatu	Celastraceae	Root, stem, leaves	Anti-diabetic, anti- hyperlipidemic
59	Semecarpus anacardium L.f.	Marking nut	Anacardiaceae	Nut	Anti-hyperlipidemic, hypoglycemic and antihyperglycemic
60	Senna auriculata (L.) Roxb.	Avaram senna, styptic weed, Matara tea	Fabaceae	Leaves, flower	Antihyperglycemic, hypolipidemic, antihyperlipidemic
61	Solanum virginianum L.	Yellow-fruit	Solanaceae	Leaves	Anti-hyperglycemic

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62	Swertia chirayita (Roxb.) Buch.-Ham. ex C.B.Clarke	nightshade Clearing nut tree, bitter stick	Gentianaceae	Whole plant	Anti-diabetic
63	Symplocos cochinchinensis (Lour.) S. Moore	Laurel sapphire berry	Symplocaceae	Leaves	Anti-diabetic, anti- lipidemic
64	Tetraena alba (L.f.) Beier & Thulin	White bean- caper Malabar	Zygophyllaceae	Whole plant	Antihyperglycemic and antihyperlipidemic
65	Tinospora sinensis (Lour.) Merr.	gulbel, Chinese tinospora	Menispermaceae	Root	Hypoglycemic and hypolipidemic
66	Trigonella foenum- graecum L.	Fenugreek	Fabaceae	Seeds	Hypoglycemic, hypcholesterolemic, anti- hyperglycemic
67	Urtica dioica L	Stinging nettle	Urticaceae	Aerial parts	Anti-hyperglycemic
68	Vitex negundo L.	Chinese chastetree, horseshoe vitex	Lamiaceae	Leaves	Anti-hyperglycemic
69	Viscum album L.	Mistletoe	Viscaceae	Leaves	Antihyperglycemic, insulinotropic

5 Conclusion and Future Perspective

The enormous economic, social and personal cost of management of diabetes makes a compelling case for prevention. In recent years, there has been much new evidence demonstrating the potentially preventable and manageable nature of diabetes, particularly by the implementation of nutritional modification. In view of this and the devastating health impact of the disease it seems prudent that primary prevention and management should be a major priority. Since the 1970s there have been numerous healths promotion projects that have attempted to encourage diabetic's patients to lead a healthier lifestyle (Steyn *et al.*, 2004). Nutrition education and lifestyle counseling should be adapted to individual needs and delivered in a patient-centered manner. Education can be delivered both to the individual child and family and in small group settings (National Institute for health and Care Excellence 2004).

In order to achieve maximum benefit from nutritional interventions in managements of diabetes changes in

government policies and legislation will be needed in addition to individual and community-based programmes. This would include mandating more nutrition education in schools, banning the advertising of unhealthy products and subsidising healthy foods at the expense of less appropriate foods. Despite much research on nutritional factors in the aetiology and management of diabetes, the risk associated with several individual nutrients is not entirely clear. Therefore, there is an urgent need for more research and rigorous evaluation of dietetic management in diabetes mellitus. In making recommendations to the public regarding the prevention of diabetes, priority should be given to the following; consumption of a nutrient-dense diet, which is low in fat, particularly saturated fat, and free sugars and high in non-starch polysaccharides (Vessby *et al.*, 2001). Also, nutritional management should be coupled with physical activities and in the case of type 1 diabetes, insulin doses need to be adjusted to balance with nutritionally adequate food intake.

Advancement in modern medicine has resulted in the development of several pharmaceutical drugs such as biguanides, thiazolidinediones, biguanides, and insulin.

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Even though these drugs show hypoglycemic activities, they are often associated with several complications such as nephrological disorders, fatigue, upset stomach, diarrhea, etc. All these reasons have stimulated the rework on herbal medicine to find suitable alternatives that will have lesser side effects and improved therapeutic effects. The present review has summarized the list of medicinal plants that are known to possess anti-diabetic properties. These plants are traditionally used by various tribal people for the treatment of several ailments. Pharmacological studies have reported their hypoglycemic, anti-hyperglycemic, insulin mimicking, anti-lipidemic properties and hence when administered in proper dosages can be beneficial for ameliorating the various complications associated with diabetes mellitus. This review provides the scope for the readers to further explore the active constituents of anti-diabetic plants and its possible mechanisms for future research.

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