

JOSLIN DIABETES CENTER & JOSLIN CLINIC
CLINICAL GUIDELINE FOR ADULTS WITH DIABETES
Rev. 05/17/17

The *Joslin Clinical Guideline for Adults with Diabetes* is designed to assist primary care physicians and specialists individualize the care of and set goals for non-pregnant adults with diabetes. This Guideline focuses on the unique needs of the patient with diabetes. It is not intended to replace sound medical judgment or clinical decision-making and may need to be adapted for certain patient care situations where more or less stringent interventions may be necessary.

The objectives of the Joslin Clinical Diabetes Guidelines are to support clinical practice and to influence clinical behaviors in order to improve clinical outcomes and assure that patient expectations are reasonable and informed. Guidelines are developed and approved through the Clinical Oversight Committee that reports to the Medical Director of Joslin Diabetes Center. The Clinical Guidelines are established after careful review of current evidence, medical literature and sound clinical practice. This Guideline will be reviewed periodically and modified as clinical practice evolves and medical evidence suggests. **(Additional Guidelines are available <https://www.joslin.org/info/joslin-clinical-guidelines.html>)**

Joslin’s Guidelines are evidence-based. A modification of the GRADE system has been adopted to enable the user an evaluation of the evidence used to support each standard of care. The table provided on page 13 describes the categories in which methodological quality and strength of recommendations have been classified. Evidence levels are graded 1A through 2C, as indicated in brackets. Where evidence is not graded, recommendations are made based on the expertise of the Clinical Oversight Committee

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(1.0) APPROACH TO CARE

(1.1) Individualizing patient care:

The needs and goals of each patient are unique. Developing a treatment plan based on a thorough assessment which includes an understanding of not only

the patient’s medical needs, but also other factors that may influence the treatment plan such as social history, race, cultural issues, ethnicity, education needs (including literacy and numeracy), comorbidities and barriers to care is essential. The patient’s treatment plan should identify medical treatment, educational

interventions, follow-up, and ongoing support. Use of the electronic medical record may help to facilitate care, by enabling the team to track progress, ensuring goals are met, and communication flows between team members and the patient.

(1.2) The patient-centered approach:

Diabetes is a condition that requires self-management. A collaborative counseling model (where the patient is involved in decisions and goal setting) helps promote behavior change. Whenever appropriate, with the patient's consent, involve family members and caregivers in medical visits and education.

(1.3) Working in a team: Diabetes is best managed by a team which may include clinicians, diabetes educators, and dietitians. The patient should be informed, and fully aware of the roles the various team members play. If access to a team is not possible within the office practice, it is useful to identify community resources. Clear communication between all providers is critical to ensure patients' needs are being met.

(1.4) Frequency of medical visits: While the frequency of visits for ongoing care should be individualized, monitoring progress of the patient through medical visits is recommended at least 2-4 visits/year. Special attention should be given to patients who do not keep scheduled appointments, have frequent hospitalizations or miss days of work/school. Since many factors contribute to the patient's ability to manage their care, the provider should:

- Engage patients in identifying and resolving contributing factors or barriers to under-utilization or over-utilization of healthcare services. Patients with challenging care may benefit from consultation with Endocrinologists focused on diabetes care. For further information on when to refer patients, please refer to: Guidelines for specialty consultation/referral ([http://www.joslin.org/docs/Referral_Guidelines_8_6_13\(1\).pdf](http://www.joslin.org/docs/Referral_Guidelines_8_6_13(1).pdf))
- Refer to a diabetes educator (DE), registered dietician, social service or behavioral health professional to address possible barriers and/or psychosocial issues
- Establish a process of follow-up communication regarding adherence to the treatment plan and sustaining behaviors.

(2.0) DIAGNOSIS OF DIABETES MELLITUS

(2.1) General criteria for diagnosis:

The diagnosis of diabetes mellitus can be made based upon:

- Random plasma glucose ≥ 200 mg/dl (11.1 mmol/L) and symptoms of diabetes (polyuria, polydipsia, ketoacidosis, or unexplained weight loss) *OR*
- Fasting plasma glucose (FPG)* ≥ 126 mg/dl (6.9 mmol/L) *OR*
- Results of a 2-hour 75-g Oral Glucose Tolerance Test (OGTT)* ≥ 200 mg/dl (11.1 mmol/L) *OR*
- Glycated hemoglobin* (A1C) $\geq 6.5\%$ (46 mmol/mol)
**

** These tests should be confirmed by a repeat test, on a different day, unless unequivocally high*

***A glycated hemoglobin (A1C) level of 6.5% or higher on 2 separate days is acceptable for diagnosis of diabetes. [1B]. However, some individuals may have an A1C < 6.5% with diabetes diagnosed by previously established blood glucose criteria. Therefore, presence of either criterion is acceptable for diagnosis. Those with an A1C of 5.7-6.4% (39-46 mmol/mol) are considered to have pre-diabetes have a high risk for developing diabetes. These patients should be treated with lifestyle changes and followed more frequently.*

The A1C should be performed in a laboratory using a method that is National Glycohemoglobin Standardization Program (NGSP) certified and standardized to the Diabetes Control and Complications Trial (DCCT) assay.
A point of care (POC) A1C is not acceptable for diagnosis of diabetes.

(2.2) Hemoglobin A1C

Diagnosis:

See above section on Diagnosis of Diabetes Mellitus

(2.2a) Goals:

A1C target goal should be individualized for each patient. A goal of < 7%(53 mmol/mol) is chosen as a practical level for most patients to reduce the risk of long term complications of diabetes. Achieving this goal is recommended if it can be done safely, and practically. **[1B]**

Alternative A1C goals may be set based upon presence or absence of microvascular and/or cardiovascular complications, hypoglycemic unawareness, cognitive status and life expectancy. **[1A]** For patients with longstanding type 2 diabetes with pre-existing CVD, or

high CAD risk (diabetes plus two or more additional risk factors), consider revising A1C goals to avoid adverse consequences of tight glycemic control e.g. hypoglycemia. [1A]

Some clinicians may translate patients' A1C level into their estimated average glucose level (eAG), based upon the work of the A1C Derived Average Glucose Study (ADAG)). This metric is also a valid tool that may be used to assess the response of patients to their diabetes treatment plan. [1C]

Joslin's A1C target goal for most patients is consistent with that of the ADA. Other expert panels such as AACE suggest that the A1C target goal should be $\leq 6.5\%$ in those newly diagnosed with diabetes and without comorbidities.

(2.2b) Caveats: The A1C may not reflect glycemic control in special patient populations, including pediatric and geriatric populations, patients with anemia or other blood disorders resulting in rapid turnover of red blood cells, in chronic liver and renal disease, following recent transfusions, or in the hospital setting. It is therefore important to interpret A1C results accordingly when determining treatment plans and goals.

(2.2c) Monitoring:

Monitor the A1C 2-4 times a year as part of the scheduled medical visit [1C] to evaluate efficacy of treatment plan. The A1C may be checked more frequently if the treatment program requires revision, or the advice regarding behavior changes need reinforcement.

Having the A1C result at the time of the visit can be useful in making timely treatment decisions. [1C] Alternatively the A1C may be performed prior to the medical visit using a point of care (POC) method.

(2.2d) Treatment:

If A1C is $\geq 7\%$ and $<8\%$, or above the individualized goal for 6 or more months:

- Review and clarify the management plan with the patient with special attention given to address:
 - nutrition and meal planning
 - physical activity
 - medication administration, schedule and technique
 - self-monitoring blood glucose (SMBG) schedule and technique
 - treatment of hypoglycemia and hyperglycemia
 - sick day management practices
- Reassess goals and adjust medication as needed [1A]
- Establish and reinforce individualized glycemic goals with patient

- Refer patient to a certified diabetes educator (DE) for evaluation, diabetes self-management education (DSME) and support for ongoing consultation. [1C]
- Consider referral to registered dietitian (RD) for medical nutrition therapy (MNT). [1B]
- Schedule follow-up appointment within 3-4 months or more frequently as the situation may dictate

If A1C is $\geq 8\%$

- Review and clarify the plan as previously noted
- Assess for psychosocial stress as a potential barrier to adequate response to treatment [1C]
- Establish and reinforce individualized glycemic goals with the patient
- Intensify therapy
- Refer patient to DE for evaluation, DSME and support for ongoing consultation. Document reason if no referral initiated
 - Refer patient to RD for MNT [1C]

If the patient has a history of severe recurrent hypoglycemia or hypoglycemia unawareness (a condition in which the patient is unable to recognize symptoms of hypoglycemia):

- Assess for changes in daily routine such as decreased food intake or increased activity [1C]
- Refer to DE for evaluation, DSME and hypoglycemia prevention; encourage family/friend attendance
- Review use of glucagon
- Consider revising A1C goal
- Discuss and reinforce goals with patient
- Adjust medications accordingly [1B]
- If insulin-treated, consider use of a more physiologic insulin replacement program such as basal/bolus therapy
- Consider and screen for other medical causes
- Consider referral for blood glucose awareness training, if available
- Consider use of continuous glucose monitoring (CGM)
- Schedule follow-up appointment within 1-2 months. If history of recent, severe hypoglycemia or change in pattern of hypoglycemia, recommend increase in frequency of communicating blood glucose levels to provider or diabetes educator.

(3.0) SELF-MONITORING OF BLOOD GLUCOSE

Self-monitoring of blood glucose (SMBG) is an important component of the treatment program for all people with diabetes. Its use is to gauge treatment efficacy, help in treatment design, provide feedback on the impact of nutritional intake and activity, provide patterns that assist in medication selection, and for those on insulin, assist in daily dose adjustments. [1B]

(3.1) Goals:

Goals for glycemic control for most people with diabetes are:

- Fasting glucose: 80-130 mg/dl (4.4 – 7.2 mmol/L)
- 2-hour postprandial glucose: <180 mg/dl (9.9 mmol/L)
- Bedtime glucose: 90-150 mg/dl (4.9- 8.3 mmol/L)

(3.2) Frequency:

The frequency of SMBG should be individualized based on factors such as glucose goals, medication changes and patient motivation. Most patients with type 1 diabetes should monitor 4-6 times per day. Some patients may need to monitor even more frequently.

Most patients using intensive insulin therapy should ideally monitor before meals and bedtime, prior to exercise, when they suspect hypoglycemia, after treating hypoglycemia as well as prior to driving. In patients with Type 1 diabetes, there is a correlation between increased SMBG frequency and lower A1C. For patients with type 2 diabetes, the frequency of monitoring is dependent upon such factors as mode of treatment and level of glycemic control. **[1C]**

(3.3) Postprandial monitoring:

To obtain meaningful data for treatment decisions, it is helpful for the patient to monitor for several consecutive days (e.g., 2-4 days). In addition to obtaining fasting and preprandial glucose levels, consider obtaining glucose readings 2-3 hours postprandial, as postprandial hyperglycemia has been implicated as an additional cardiovascular risk factor. **[1B]**

Postprandial monitoring is particularly recommended for patients who:

- Have an elevated A1C but fasting glucose is at target
- Are initiating intensive (physiologic) insulin treatment programs
- Are experiencing problems with glycemic control
- Are using glucose-lowering agents targeted at postprandial glucose levels
- Are making meal planning or activity adjustments

One -hour postprandial glucose monitoring should be considered:

- During pregnancy**[1A]**
- For those patients using alpha-glucosidase inhibitors

Encourage the patient to bring SMBG results (written records or meter for downloading) to each visit for review with provider/educator.

(3.4) Using alternate sites to monitor:

Blood glucose levels from sites such as the upper arm, forearm, and thigh may lag those taken from the

fingertips, particularly when glucose levels are changing rapidly. Glucose levels may change rapidly with exercise, eating, after insulin administration or with hypoglycemia. For this reason, alternate site monitoring is not recommended in the following situations:

- When the blood glucose may be changing rapidly
- For patients using intensive insulin treatment programs
- If hypoglycemia is suspected
- In patients with hypoglycemia unawareness

(3.5) Continuous glucose monitoring:

Real time continuous glucose monitoring (CGM) measures interstitial glucose levels and correlates with plasma glucose levels. CGM requires calibration with SMBG at least twice daily. Use of CGM technology, has been shown to decrease A1C in adults 25 years old and older using intensive insulin therapy along with CGM, compared with those using intensive insulin therapy with SMBG. The best predictor of A1C lowering was increased frequency of sensor use. CGM can be helpful in insulin-treated patients with hypoglycemia unawareness and/or frequent severe hypoglycemic episodes. The FDA recently approved the use of properly calibrated CGM devices (i.e. - Medtronic 670G pump/sensor and DexCom G5 sensor) to help make treatment decisions. Patients with insulin-treated diabetes over 65 years old who would benefit from CGM should have access with insurance coverage. Intensive diabetes education and support are essential for optimal CGM implementation and ongoing use.

(4.0) HYPOGLYCEMIA

(4.1) Classification:

Prompt action is recommended for the treatment of hypoglycemia. When possible, the patient should confirm symptoms with SMBG to document hypoglycemia. All patients with type 1 diabetes should ensure that a family member/companion/caregiver knows how to administer a glucagon injection in the event the patient is unable or unwilling to take carbohydrate orally. **[1C]**

The International Hypoglycemia Study Group recently recommended that hypoglycemia be classified as

- Level 1(Glucose alert level) with glucose less than 70 mg/dL (3.8 mmol/L); which is considered sufficiently low for treatment with fast-acting carbs
- Level 2(clinically significant hypoglycemia) with glucose less than 54 mg/dL (2.9 mmol/L) which is considered serious and clinically important hypoglycemia;
- Level 3(severe hypoglycemia) with no specific glucose threshold but associated with cognitive impairment requiring external assistance.

(4.2) Treatment:

- Caution patient to avoid alternate site monitoring with blood glucose meter when hypoglycemic
- Treat as mild-moderate hypoglycemia if patient is symptomatic or unable to confirm hypoglycemia with SMBG, or if blood glucose levels are >54 mg/dl (2.9 mmol/L) and <70 mg/dl (3.8 mmol/L) (<90 mg/dl (4.9 mmol/L) at bedtime or overnight).
- To treat mild to moderate hypoglycemia (plasma glucose 54-70 mg/dl (3.8-2.9 mol/L) most times of the day and < 90 mg/dl (4.9 mmol/L) at bedtime or overnight), begin with 15-20 grams of carbohydrate (1/2 cup juice or regular soft drink, 3-4 glucose tabs). **[1C]**
- If glucose level is ≤54 mg/dl (2.9 mmol/L), consume 20-30 grams of carbohydrate. **[1C]**
- Recheck blood glucose after 15 minutes. **[1B]**
- Repeat hypoglycemia treatment if blood glucose does not return to normal range after 15 minutes. **[1C]**
- Follow with additional carbohydrates if next meal is more than one hour away. **[1C]**
- If hypoglycemia persists after 2-3 treatments, patient or companion should be instructed to contact their healthcare provider or seek emergency care.
- In event of severe hypoglycemia (altered consciousness, unable to take carbohydrate orally, or requiring the assistance of another person) treat with glucagon and/or intravenous glucose. **[1C]**
- For patients with hypoglycemia unawareness, the threshold for treatment of hypoglycemia needs to be individualized. **[1C]**
- For patients using real-time CGM, check 15 minutes post treatment using a finger stick and not the sensor reading. Due to the physiologic lag between blood and interstitial glucose, the sensor will yield a lower result and can lead to over-treatment. **[1B]**
- For patients with gastroparesis, treat hypoglycemia with oral glucose gel.
- The patient's treatment plan should be revised if hypoglycemic events are frequent, or if they have hypoglycemia unawareness.

(4.3) Education:

- Instruct the patient to obtain and wear or carry diabetes identification.
- Instruct patient to carry treatment for hypoglycemia at all times.
- Instruct all patients with type 1 diabetes and patients with type 2 diabetes who are at risk for hypoglycemia to check blood glucose before operating a motor vehicle or other potentially dangerous equipment. In addition, advise them to

check blood glucose regularly if driving for one or more hours. Hypoglycemia should be treated immediately, and patients should not drive until their blood glucose has reached and remained at a safe range for at least 30 minutes and/or until cognitive function is restored **[1B]**

- Identify potential causes of hypoglycemia to prevent its occurrence. **[1C]**
- Be clear in communicating modified treatment goals in individuals with hypoglycemia unawareness
- Glucagon injections should be prescribed, to all patients with severe hypoglycemia. Education on its use should be provided to the patient, and his or her caregivers/ family members if possible.

(5.0) DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT (DSME/S)

All people with diabetes should receive DSME/S according to the National Standards for Diabetes Self-Management Education and Support, to facilitate knowledge and to implement and sustain self-care skills and problem-solving (1B). Critical time points recommended for DSME/S are:

- At diagnosis
- Annually for assessment of education, nutrition and emotional needs
- When new complicating factors arise
- When transitions in care occur

Multiple visits with a diabetes educator (DE) are recommended to evaluate progress toward goals (1B)

Group education sessions are encouraged for cost effectiveness and efficiency of staff utilization. Group education is a benefit to patients as it allows them to share ideas and concerns and enables them to learn from one another. **[1B]**

(6.0) MEDICAL NUTRITION THERAPY (MNT)

There is not a one-size-fits all eating pattern for individuals with diabetes. Patients with newly diagnosed diabetes should receive either individualized or group MNT, preferably by a registered dietitian nutritionist (RDN) who is knowledgeable and skilled in providing diabetes specific MNT. MNT delivered by a registered dietitian is associated with an A1C decrease of 0.3-1% for people with type 1 diabetes and 0.5-2% for people with type 2 diabetes. (1A) Goals of MNT are to promote healthy eating patterns while addressing the unique nutrition needs of each patient based on their personal preferences, cultural background, health literacy, barriers to change and each individual's ability to make changes in their eating habits.

Weight management is important for overweight and obese people living with type 1 and type 2 diabetes. There is strong evidence that modest and persistent weight loss is beneficial to the management of type 2 diabetes and can delay the progression from pre-diabetes to type 2 diabetes.

For further details please refer to Joslin's Clinical Nutrition Guideline for Overweight and Obese Adults with Type 2 Diabetes;

http://www.joslin.org/docs/Nutrition_Guidelines_101916.pdf

(7.0) PHYSICAL ACTIVITY

All adults should consult their healthcare provider and/or see an exercise physiologist to discuss a safe exercise program that is appropriate to their abilities. [1C]

(7.1) Physical activity for healthy adults

- Physical activity should be an integral component of the diabetes care plan to optimize glucose control, decrease cardiovascular risk factors, and achieve or maintain optimal body weight. [1A]
- A moderate-intensity aerobic (endurance) physical activity minimum of 30 minutes 5 days per week or vigorous-intensity aerobic physical activity for a minimum of 20 minutes 3 days per week should be achieved unless contraindicated. Activity can be accumulated toward the 30-minute minimum by performing bouts, each lasting 10 or more minutes. [1A]
- All adults, and particularly those with type 2 diabetes, should **decrease the amount of time spent in daily sedentary behavior**. Prolonged sitting should be interrupted every 30 min for blood glucose benefits, particularly in adults with type 2 diabetes.
-
- A target of 60-90 minutes, 6-7 days per week is encouraged for weight loss if overweight or obese [1A]
- To increase lean body mass, full body resistance training should be incorporated into the activity plan 3-4 days per week, and include upper, core and lower body strengthening exercises using free weights, resistance machines or resistance bands. [1B]
Beginning training intensity should be moderate, involving 10–15 repetitions per set, with increases in weight or resistance undertaken with a lower number of repetitions (8–10) only after the target number of repetitions per set can consistently be exceeded; increase in resistance can be followed by

a greater number of sets and lastly by increased training frequency.

- Stretching exercises should be done when muscles are warm or at the end of the activity plan to loosen muscles and prevent soreness. [1B]

(7.2) Physical activity for adults with medical or physical limitations

- A moderate-intensity aerobic (endurance) physical activity minimum of 30 minutes 5 days per week or vigorous-intensity aerobic physical activity for a minimum of 20 minutes 3 days per week should be achieved, as feasible, unless contraindicated. Activity can be accumulated toward the 30-minute minimum by performing bouts, each lasting 10 or more minutes. [1A]
- To increase lean body mass, resistance training should be incorporated into the activity plan 3-4 days per week, as feasible, and include upper, core and lower body strengthening exercises using free weights, resistance machines or resistance bands. [1B]
- Incorporate balance exercises to prevent falling and injury.
- Functional Fitness Testing is useful to assess patients' functionality and track their progress. Testing such as 6-Minute Walk Test, 2-Minute Step Test, Balance Assessment and Hand strength should be included at baseline and post intervention [1C]
- **See section on EYES**

(8.0) CARDIOVASCULAR HEALTH

(Also see sections on *Lipids, Blood Pressure, Physical Activity and Smoking*)

(8.1) Anti-platelet therapy

A daily enteric-coated ASA (75-162 mg) unless contraindicated * as a **primary** prevention strategy for men ≥ 50 years of age [1C] and for women ≥ 60 years of age [1C] with ONE or more of the following risk factors:

- Family history of premature** CAD or stroke
- HTN
- Current cigarette smoker
- Albuminuria
- Hyperlipidemia

Recommend a daily enteric-coated ASA (75-162 mg) or clopidogrel (75 mg, if aspirin intolerant) or another agent of the class, as a **secondary** prevention strategy for anyone with ONE or more of the following: [1A]

- History of MI, angina, or documented CAD
- Vascular revascularization
- Non-hemorrhagic stroke
- TIA
- PAD

**Possible contraindications for antiplatelet therapy may include allergy, bleeding tendency, anticoagulant therapy, recent gastrointestinal bleeding and clinically active hepatic disease. Eye disease is usually not a contraindication for ASA therapy.*

***Premature – 1st degree male relatives younger than 55 years of age; 1st degree female relatives younger than 65 years of age*

(8.2) Other therapeutic considerations:

Consider using beta-blocker in all patients with a history of MI or with documented CAD unless contraindicated. [1A]

Consider using ACE inhibitors (or ARBs if ACE inhibitors not tolerated) in patients with known CAD or cardiovascular risk factors and age 55 yrs. or greater. [1B]

Thiazolidinediones (pioglitazone, rosiglitazone) are contraindicated in patients with NYHA classes III and IV and conditions of fluid overload (i.e. CHF). (See Pharmacological Guideline for additional caveats on TZDs) [1A]

Consider recommending aerobic activity if not clinically contraindicated, and a weight-loss program if patient is overweight or obese. [1A]

(8.3) When to conduct a stress test:

Based on current research and understanding of coronary artery disease in diabetes, it is reasonable to screen patients with diabetes who: [1C]

- Complain of typical or atypical chest pain
- Have an abnormal ECG
- Have a diagnosis of peripheral artery disease (PAD) or carotid disease
- Are >35 years of age with sedentary lifestyle about to start a rigorous exercise program.

There is currently no strong evidence to support screening asymptomatic patients with type 2 diabetes for silent myocardial ischemia. [1C]

Patients with autonomic neuropathy may have increased risk of asymptomatic ischemia and therefore warrant careful attention. [1B]

If stress testing is performed, either nuclear imaging or echocardiography with ECG monitoring is recommended. An exercise stress test is preferred, if resting ECG is normal and patient is able to exercise, as the response to exercise is an important prognostic factor. If the patient cannot adequately exercise, pharmacologic stress testing is warranted.

(8.4) Lipid management

(8.4a) Screening for lipid disorders:

Adults should be screened annually for lipid disorders with measurements of serum cholesterol, triglycerides, and LDL and HDL cholesterol, preferably fasting. [1B]

(8.4b) Treatment

All patients should receive information about a meal plan designed to improve glycemic and lipid control, physical activity recommendations, and cardiovascular risk reduction strategies (with an emphasis on smoking cessation and blood pressure control.) Consultation with appropriate education discipline is preferred. [1A].

Institute therapy after abnormal values are confirmed.

- All patients with any form of clinical diagnosis of atherosclerotic cardiovascular disease (ASCVD), or if LDL-C \geq 190 mg/dl: treat with statin to reduce LDL-C \geq 50% [1A]
- Patients, ages 40-75 years old, without clinical evidence of ASCVD and LDL-C 100-190 mg/dl: treat with statin to reduce LDL-C by 30-50%. Consider reduction of > 50% if one or more additional major risk factors* are present:
 - Calculated 10-year risk of ASCVD > 7.5 % using the ACC/AHA risk equation calculator (<http://my.americanheart.org/cvriskcalculator>)[1B]
 - Family history of premature ASCVD
 - High blood pressure
 - Tobacco use
 - Albuminuria
- Patients, ages 40-75 years old without clinical evidence of ASCVD and LDL-C 70-100mg/dl: treat with statin to reduce LDL-C by > 30% [1C]
- In patients < 40 yrs. of age, consider statin if LDL-C > 100 mg/dl and multiple CVD risk factors [2B]
- In patients > 75 yrs. of age, there is no clear evidence for benefits of initiating statin therapy in the absence of ASCVD or multiple CV risk factors. [2C]
- Re-check lipids after drug initiation or dose escalation in 6-12 week. Thereafter, check lipids every 3-12 months to monitor adherence. May down-titrate statin dose if LDL- C < 40 mg/dl.
- No evidence for benefits of statin therapy in patients on hemodialysis, or those with heart failure (NYHA class II-IV). [1B]
- If adequate reduction in LDL-C as described above has been achieved, a specific LDL-C goal (< 70 and < 100 mg/dl) or non-HDL-C goal (< 100 and < 130 respectively) for those with or without ASCVD, respectively, is not recommended unless baseline lipid levels not known.

- In patients with ASCVD or those with familial hypercholesterolemia (FH) who are unable to achieve LDL-C goal with maximum tolerated statin therapy, add ezetimibe or a bile acid sequestrant. Also consider a PCSK9 inhibitor in such cases.
- For primary prevention of cardiovascular disease, consider a use of a bile acid sequestrant or niacin (alone, or in combination therapy) for patients intolerant to multiple statins, or who have unacceptable adverse events. [2B]
- Statins are contraindicated during pregnancy or if contemplating pregnancy.
-

Patients with LDL-C at goal and fasting triglycerides

≥ 150 mg/dl or HDL-C ≤ 40 mg/dl

- Optimize glycemic control [1A]
- Refer to RD for dietary modification and therapeutic lifestyle changes (TLC) [1A]
- Consider referral to an exercise specialist for an appropriate exercise regimen
- Recheck lipids within 6-12 weeks
- In patients with fasting triglyceride levels 200-499 mg/dl and/or HDL-C ≤ 35 mg/dl after optimal statin therapy, consider adding a fibrate, [2B]
- If triglycerides persistently ≥ 500 mg/dl, secondary causes of hypertriglyceridemia should be considered and managed appropriately. Initiate treatment with very low fat meal plan and with a fibrate for prophylaxis against acute pancreatitis; reassess lipid status when triglycerides < 500 mg/dl [1A]
- If fasting triglycerides remain > 500 mg/dl after initiation of fibrate, consider the addition of fish oil (to provide 2-4 gm omega-3 fatty acids daily), or niacin [2B]

(8.5) Blood pressure management

(8.5a) Blood pressure measurement:

- Check BP at all routine visits after patient has been seated for at least 5 minutes. Use proper-size cuff and arm position. Postural BP (sitting then standing) should be checked initially, and as clinically indicated:
 - In cases of known or suspected orthostatic hypotension (defined as a fall in systolic BP (SBP) of > 20 mmHg or diastolic BP (DBP) of ≥ 10 mmHg or an increase in heart rate by more than 20 beats per minute after 3 minutes of standing)
 - In cases where standing upright is associated with lightheadedness, syncope or signs of brain hypoperfusion. [1C]
- Initiate lifestyle changes if BP $> 130/80$ mmHg
- Consider initiating pharmacologic therapy if the average of 3 blood pressure measurements is $> 140/90$ mmHg.

- Schedule for follow up blood pressure check within 1 month [1B]

(8.5b) Blood pressure targets:

- BP goal for each patient > 18 years of age is $\leq 140/90$ mmHg. [1B]
- SBP ≤ 130 mmHg may be appropriate for individuals without CVD or without multiple risk factors. [1B]
- No clear evidence exists for significant benefits to be gained by lowering SBP to < 120 mmHg in those with CHD or multiple risk factors. [1B]
- BP goal for patients with albuminuria > 300 mcg/mg is $< 130/80$ mmHg, if tolerated. [1C]
- Initial goal for patients with isolated systolic HTN (SBP > 180 mmHg and DBP < 80 mmHg) is a SBP < 160 mmHg. [2B]
- Initial goal for patients with SBP 160-179 mmHg is to lower SBP by 20 mmHg. If well tolerated, lower BP goals may be indicated. [1B]

(8.5c) Treatment

If SBP ≥ 140 mmHg or DBP ≥ 90 mmHg, a 3-month trial of lifestyle modification is warranted as follows: [1C]

- Counsel about meal plan, use of DASH and DASH-sodium diet, activity, weight loss, sodium reduction, alcohol and stress reduction
- Consider referral to RD for medical nutrition therapy (MNT)
- Encourage home BP self-monitoring and documenting it in a log to bring to clinic appointments.
- Instruct patient to have BP checked two times a week prior to the next appointment
- Follow-up with healthcare provider within 2-4 weeks
- Initiate or adjust therapy with antihypertensive agents as clinically indicated if BP remains above goals

Studies have shown that aggressive management and control of blood pressure may result in long-term benefits

• Pharmacotherapy:

Efficaciousness is the most important consideration in choosing an initial anti-hypertensive drug. In that sense, any available antihypertensive drug can be an appropriate choice however, other considerations (presence of albuminuria, co-existing CAD, or cost) dictate a preference for ACE inhibitors, ARBs, calcium channel blocker (CCB), and thiazide-type diuretics. [1A] In general, ACEI and ARB should not be used in combination

Consider ACE inhibitors or ARBs for patients with persistent urine albumin/creatinine ratio > 30 mcg/mg. These drugs require monitoring of serum creatinine and

K⁺ within 1 week of starting therapy and periodically thereafter. [1A] (See section on *Renal Disease and Albuminuria*)

ACEI/ARBs are contraindicated during pregnancy or if contemplating pregnancy

(9.0) RENAL HEALTH

(9.1) Screening for Renal Health

Measure serum creatinine at least annually to estimate glomerular filtration rate (GFR) regardless of degree of urine albumin excretion. (See *Joslin's Guideline for Specialty Consultation/Referral* for guidance as to when to refer to a renal specialist.) [1C]

Estimate GFR (eGFR) using CKD-EPI calculation. If eGFR is <60 ml/min, evaluate for complications of kidney disease (anemia, hyperparathyroidism, and vitamin D deficiency).

Screen for albuminuria by checking urine albumin/creatinine (A/C) ratio as follows:

- Type 1 patients within 5 years after diagnosis and then yearly [1C]
- Type 2 patients at diagnosis (after glucose has been stabilized) and then yearly [1C]
- Annually in all patients up to age 70 years [2C]
- As clinically indicated in patients >70 years of age

Albuminuria is recognized as a major independent risk factor for CAD in patients with diabetes. Albuminuria may be measured with a spot or timed urine collection. Spot urine is preferred for simplicity.

Continue use of routine urinalysis as clinically indicated. [2C]

Patients should be advised that BP control, glycemic control and management of albuminuria may slow the progression of CKD.

(9.2) Treatment

If A/C ratio < 30 mcg/mg or timed urine albumin < 30 mg/24 hr:

- recheck in 1 year

If A/C ratio 30-300 mcg/mg or timed urine albumin 30-300 mg/24 hr:

- Confirm presence of albuminuria with at least 2 of 3 positive collections done within 3-6 months. In the process, rule out confounding factors that cause a false-positive such as UTI, pregnancy, excessive exercise, menses or severe hypoglycemic event. [1C]
- Consider testing first morning urine
- Consider consult with nephrologist for blood pressure control, successive increases in albumin and other issues (i.e., GFR < 60 ml/min) [2C]

Once confirmed:

- Evaluate BP and initiate/modify aggressive blood pressure treatment to achieve a BP of < 130/80 mmHg [2B]
- Recommend patient self-monitor BP with portable cuff and maintain a record/log. The monitoring schedule should be determined with the healthcare provider and is based on patient circumstance.
- Strive to improve glycemic control with an optimal goal A1C of < 7% or as otherwise clinically indicated [1A]
- Refer to diabetes educator for glucose management
- Initiate/ modify ACE inhibitor or ARB treatment if albuminuria persists. Check K⁺ and creatinine about 1 week after making these medication changes. [1A]
- Repeat A/C ratio at least every 6 months. Consider increase in frequency when changes in medication are made. [2C]

If A/C ratio > 300 mcg/mg (> 300 mg/24 hr) or persistent albuminuria (positive dipstick for protein or ≥ 30 mg/dl):

- Follow all guidelines as stated for A/C ratio 30-300 mcg/mg
- Consider BP goal of < 130/80 mmHg [2B]
- Evaluate for patient adherence, with emphasis on avoidance of high sodium and very high protein intake
- Consider referral to RD for MNT

Consider referral to nephrologist to:

- Assess cause(s) of impaired kidney function including assessing for non-diabetes kidney disease
- Maximize therapies aimed at slowing progression of kidney disease (e.g., blood pressure control and reduction of urine protein level)
- Treat complications of kidney disease (hyperphosphatemia, anemia, etc.)
- Evaluate a rapid rise in serum creatinine, abnormal sediment, concomitant hematuria or sudden increase in albuminuria.
- Assess problems with ACE inhibitors use, difficulties in management of high BP or hyperkalemia
- Manage resistant hypertension (blood pressure that remains above goal despite concurrent use of three antihypertensive agents of different classes (one of which should be a diuretic, and all should be at maximum dose tolerated)

(10.0) OCULAR HEALTH

(10.1) Screening for eye disease

Refer patient for comprehensive dilated eye exam or validated retinal imaging to determine level of retinopathy.

- Type 1 diabetes: initial eye exam at start of puberty or once patient is 10 years of age or older, whichever is earlier, within 3-5 years of diagnosis. Annual eye exam thereafter **[1A]**
- Type 2 diabetes: at diagnosis and annually thereafter **[1A]**
- Pregnancy in pre-existing diabetes: prior to conception and during first trimester with follow-up during pregnancy as determined by first trimester exam and 6-12 weeks post-partum. **[1B]**
- For physiologic insulin therapy (pump therapy or multiple daily injections): consult with patient's eye care provider or evaluate retinal status with validated retinal imaging to determine level of retinopathy and appropriate follow-up care prior to initiating physiologic insulin therapy. **[1A]**

(10.2) Treatment:

Aggressively treat known medical risk factors for onset and progression of retinopathy:

- Strive to improve glycemic control with optimal A1C goal of < 7%. **[1A]**
- Monitor eye disease carefully when intensifying glycemic control. **[1A]**
- Strive for BP <130/80 mmHg. **[1B]**
- Treat albuminuria. **[1B]**
- Strive to maintain total cholesterol, LDL, HDL and triglyceride levels as per the recommendations outlined in the *Lipids* Section of this Guideline. **[1A]**
- Treat anemia. **[1B]**

Activity programs that involve strenuous lifting, harsh, high-impact components, or activities that place the head in an inverted position for extended periods of time may need to be revised depending on the level of retinopathy.

Reinforce follow-up with eye care provider for any level of retinopathy including no apparent retinopathy. The frequency of follow-up is dependent upon the level of retinopathy and presence of risk factors for onset and progression of retinopathy and is determined by the eye care provider.

- For high-risk proliferative diabetic retinopathy, scatter (panretinal) laser photocoagulation and/or intravitreal anti-vascular endothelial growth factor (VEGF) injection is indicated promptly. **[1A]**
- For clinically significant macular edema (CSME), or center-involved macular edema, focal laser and/or

intravitreal anti-vascular endothelial growth factor (VEGF) injection is generally indicated regardless of level of retinopathy. **[1A]**

- The level of diabetic retinopathy and diabetic macular edema (DME) generally determines follow-up. * **[1A]** See suggested follow-up below:

If No Diabetic Retinopathy:

12 months

If Mild Nonproliferative Diabetic Retinopathy:

Without DME, 12 months

With DME, ** monthly if undergoing (anti-VEGF) treatment, otherwise 3-4 months

If Moderate Nonproliferative Diabetic Retinopathy:

Without DME, 6-9 months

With DME, ** monthly if undergoing anti-VEGF treatment, otherwise 3-4 months

If Severe - Very Severe Nonproliferative Diabetic Retinopathy:

Without DME, *** 3-4 months

With DME, ** monthly if undergoing anti-VEGF treatment, otherwise 3-4 months

If Proliferative Diabetic Retinopathy less than High-Risk:

Without DME, ** * 1 week – 3-4 months

With DME, ** 1 week – monthly if undergoing anti-VEGF treatment, otherwise 3-4 months

If High-Risk Proliferative Diabetic Retinopathy

With or without DME – scatter (panretinal) laser photocoagulation and/or intravitreal anti-vascular endothelial growth factor (VEGF) injection with follow-up in 3 months, monthly if undergoing anti-VEGF treatment

*The presence of known risk factors for onset and progression of retinopathy may suggest follow-up at shorter intervals for all levels of retinopathy

** Focal laser surgery and/or intravitreal anti-VEGF injection is generally indicated for CSME or center-involved macular edema. If receiving anti-VEGF treatment, follow-up is generally monthly

*** Scatter laser surgery may be indicated, especially for type 2 diabetes or type 1 diabetes of long duration

(11.0) NERVOUS SYSTEM HEALTH

(11.1) Screening for neuropathy

(11.1a) Methods:

- Ask patient about loss of sensation in the limbs, symptoms of pain, tingling, paresthesia, weakness or gait instability.
- Evaluate feet for sensation using a 128 Hz tuning fork and Semmes-Weinstein 5.07 monofilament. **[1B]**
- Evaluate reflexes
- Laboratory screening with complete blood count, lipid panel, thyroid panel, B12 level (methylmalonic acid and/or homocysteine if low normal B12), serum and urine protein electrophoresis, as clinically indicated.
- Neurophysiologic testing (EMG, nerve conduction studies or skin biopsy analysis of intra-epidermal nerve fiber density) should be considered in atypical cases.
- Assess for symptoms of autonomic neuropathy such as erectile dysfunction, gastroparesis, or postural hypotension. If symptoms of autonomic neuropathy are present, refer for evaluation by formal autonomic testing (including heart rate variability testing), blood maneuver and the blood pressure response to upright tilt table testing or standing) **[1B]**

(11.1b) Frequency:

- For patients with type 1 and 2 diabetes without complications, conduct symptom and examination screen at time of diagnosis and at least annually. **[1C]**
- For the “at-risk patients,” * conduct symptom and examination screen at all routine interval visits. **[1C]**
- Laboratory screening at the time of diagnosis of diabetes or with change in symptoms or examination. **[1C]**
- Screen for cardiovascular autonomic neuropathy at the time of diagnosis of type 2 diabetes, or 5 years after diagnosis of type 1 diabetes. Screening should be repeated yearly or with development of symptoms. **[1C]** If symptoms of autonomic neuropathy are present, refer for evaluation by formal autonomic testing (including heart rate variability testing, blood pressure and heart rate response to a Valsalva maneuver and the blood pressure response to upright tilt table testing or standing.) **[1B]**
- Neurophysiologic testing only for atypical cases. **[1C]**

“At-Risk Patients” include patients who smoke, have vascular insufficiency, neuropathy, retinopathy, nephropathy, history of ulcers or amputations, structural deformities, infections, skin/nail abnormality, are on anticoagulation therapy or who cannot see, feel or reach their feet.

(11.2) Treatment:

For patients with acute problems or who are “at risk”:

- Consider referral to neurologist for:
 - atypical neuropathy
 - rapidly progressive symptoms
 - severe pain unresponsive to first line therapy
 - weakness suggestive of diabetic amyotrophy

For patients with symptoms related to diabetic peripheral or autonomic neuropathy:

- Consider medications as they improve quality of life **[1A]**

(12.0) FOOT HEALTH

(12.1) Screening

(12.1a) Methods:

Screening should include:

- Questions about loss of sensation in the limbs, or symptoms of pain, including claudication, tingling or other paresthesia
- Foot evaluation for sensorimotor (Semmes-Weinstein 5.07 monofilament and 128 Hz tunic fork.) **[1B]**
- Evaluate reflexes, skin and soft tissues integrity, nail condition, callous formation, vascular sufficiency (pedal pulses) and biomechanical integrity
- Examination of shoes for wear and appropriateness.

(12.1b) Frequency:

- For patients with type 1 and 2 diabetes without complications and significant risk factors, conduct foot screen at time of diagnosis and at least annually thereafter. **[1C]**
- For the “at-risk patients,” * check feet at all routine interval visits. **[1C]**

“At-Risk Patients” include patients who smoke, have vascular insufficiency, neuropathy, retinopathy, nephropathy, history of ulcers or amputations, structural deformities, infections, skin/nail abnormality, are on anticoagulation therapy or who cannot see, feel or reach their feet.

(12.2) Treatment:

For patients with acute problems or who are “at risk”:

- Refer to podiatric physician for routine care and evaluation [1B]
- Refer to DE for foot care training** [1C]
- Consider referral to neurologist for:
 - atypical neuropathy
 - rapidly progressive symptoms
 - severe pain unresponsive to first line therapy
 - weakness suggestive of diabetic amyotrophy

For current ulcer or infection: mild* [1C]**

*** Mild Infection or Ulcer

Superficial (no foul odor) No significant ischemia
No bone or joint involvement No systemic toxicity
Minimal or no cellulitis (< 2 cm)

- Instruct patient in non-weight bearing, if appropriate
- Apply local dressings with topical antiseptic
- Consider baseline x-ray to evaluate for bone integrity and possible osteomyelitis
- Consider systemic antibiotic therapy
- Refer to podiatric physician for evaluation and treatment
- Refer for foot care training
- Ensure follow-up appointments are kept

For limb-threatening** ulcer or infection: [1C]**

****Limb-threatening:

Deep ulcer involvement	Bone or joint Gangrene
Lymphangitis	
Cellulitis (>2cm)	Systemic toxicity
Significant ischemia	No social support system
Immunocompromised	Foul odor in ulcer

Osteomyelitis, is presumed to be present if able to probe through the ulcer to the bone.

- Urgent hospitalization
- Consult a podiatric physician and vascular surgeon for immediate evaluation and treatment

(12.3) **Foot care training: [1C]

- Foot care training should address:
 - Avoidance of foot trauma
 - Daily foot inspection
 - Nail care
 - Callous formation
 - Proper footwear
 - Impact of loss of protective sensation on morbidity
 - Need for smoking cessation
 - Action to take when problems arise
 - Importance of glucose control on disease progression

(13.0) ORAL HEALTH

- Periodontal disease is associated with suboptimal diabetes control and may be a risk factor for cardiovascular disease. There is mixed evidence on the impact of treatment of periodontal disease on glycemic control.
- Referral to a dentist should be considered an essential component of a comprehensive diabetes care plan,
- At initial visit and annually, discuss need for dental cleaning at *least* every six months. [1C]
- Refer to dental specialist for oral symptoms and findings such as sore, swollen, or bleeding gums, loose teeth or persistent mouth ulcers. [1C]
- If edentulous, refer to dental specialist for restoration of functional dentition.

(14.0) BEHAVIORAL HEALTH

A psychosocial evaluation should be an integrated component of the initial assessment and the ongoing care of all patients with diabetes and should be strongly considered in the following situations:

Newly diagnosed diabetes:

Assess at least the following: [1C]

- Ability to cope with the emotional impact and lifestyle changes of diabetes
- Level of social support
- Barriers to treatment and self-management
- Type and degree of non-diabetes related life stress

During hospitalizations or any intensification in treatment, significant life change, problems with self-management, or metabolic stability. Key areas to assess:

- Diabetes distress: consider using PAID as a screening tool.
- Depression: consider using PHQ-9 or PHQ-2 as a screening tool
- Anxiety (e.g., compulsive SMBG fear of injections).
- Exaggerated fear of hypoglycemia: consider referral for blood glucose awareness training.
- Disordered eating: consider inquiry about insulin omission or bingeing if A1C >9% or recurrent DKA
- Family conflict related to diabetes
- Substance abuse: consider use of CAGE (alcohol screening tool)

Newly diagnosed complications from diabetes:

Assess at least the following:

- Emotional impact (diabetes distress, depression, anxiety) and lifestyle changes for patient and family.

- Barriers to treatment and self-management.
- Level of social support
- Type and quantity of non-diabetes related life- stress

Patients using second generation or atypical antipsychotic medications should be monitored for weight gain with resulting increases in glucose, lipid and blood pressure levels.

(15.0) WOMEN'S HEALTH

(Refer to Joslin's *Guideline for Detection and Management of Diabetes in Pregnancy* for more details) http://www.joslin.org/docs/Pregnancy-Guidelines_11-13-2016_corrected_1-11-2017.pdf

- *All women of reproductive age, should be assessed for the possibility of pregnancy prior to initiating new medications, and counseled on their potential risks on the developing fetus.*
- Counsel women with the potential for conception about contraception use and relationship of blood glucose control to fetal development and pregnancy outcomes. **[1C]**
- At initial and annual visit, discuss sexual function.
 - Assess for infectious, hormonal, psychological, or structural etiologies if dysfunction exists.
 - Refer to specialist as indicated. **[1C]**
- Follow appropriate guidelines for pap/pelvic and mammography screening for primary care patients. **[1B]**
- Individualize approach to bone health for women with risk factors for osteoporosis, including surgical and natural menopause. **[1B]**
 - Ensure adequate intake of calcium and vitamin D.

(16.0) MEN'S HEALTH

- At initial and annual visit, discuss sexual function and any fertility concerns.
 - assess for hormonal, psychological, or structural etiologies if dysfunction exists. **[1C]**
- For men with type 2 diabetes, consider screening for low testosterone: **[1B]**
 - screen with total testosterone and sex hormone binding globulin
- Refer to specialist as indicated.

(17.0) ADDITIONAL CONSIDERATIONS:

(17.1) Tobacco dependence

Screen:

Assess patient's use of tobacco and e-cigarettes at initial and follow-up visits.

Treatment: (*If patient smokes*)

- Discuss rationale for and strongly recommend smoking cessation. **[1A]**
- Review options available to assist in smoking cessation, including medications and cessation programs. **[1B]**

(17.2) Identifying sleep disorders:

- At initial visit and annually, inquire about sleep quality, level of fatigue and symptoms such as snoring and restless sleep **[1C]**
- Obstructive sleep apnea is more frequent in the setting of central obesity and is a risk factor for CVD
- Refer for sleep study if indicated
- The evidence surrounding the impact of sleep apnea treatment on diabetes control has been so far inconclusive.
- Special attention with regards to shift workers should be undertaken. An individualized care plan should be tailored to their schedules, and the effect of shift work on glycemic control should be assessed at each visit.

(17.3) Immunizations:

Recommend the following vaccines:

- Influenza vaccine: yearly for all adult patients with diabetes **[1B]**
- Pneumococcal vaccine with PPSV23 (pneumococcal polysaccharide vaccine): once for all patients with diabetes. **[1B]**
 - Patients ≥ 65 years of age should receive PCV13 (pneumococcal conjugate vaccine) at least one year after vaccination with PPSV23, followed by a one-time revaccination if they received the previous dose ≥ 5 years earlier **[1C]**
 - Repeat vaccination should be considered for those with nephrotic syndrome, chronic renal disease and other immunocompromised states
 - Hepatitis B Vaccine 3-dose series: for unvaccinated adult patients with diabetes (age 19-59 years) **[1C]**. May also consider for unvaccinated adults ≥ 60 years. **[2 C]**

List of abbreviations

AACE: American Association of Clinical Endocrinologists
A1C: Glycohemoglobin (hemoglobin A1C)
A/C Ratio: albumin/creatinine ratio
ACE inhibitor: angiotensin-converting enzyme inhibitor
ADA: American Diabetes Association
ADAG: A1c-Derived Average Glucose study
ARBs: angiotensin receptor blockers
ASA: aspirin
ASCVD: arteriosclerotic cardiovascular disease
BP: blood pressure
CAD: coronary artery disease
CAGE: Alcohol screening questionnaire
CGM: Continuous glucose monitoring
CHF: Congestive heart failure
CKD: chronic kidney disease
CSME: clinically significant macular edema
CVD: cardiovascular disease
CVD: cardiovascular disease, including coronary heart disease, peripheral vascular disease, and cerebrovascular disease
DASH: Dietary approaches to stop hypertension
DBP: diastolic blood pressure
DCCT: Diabetes Control and Complication Trial
DE: diabetes educator
DKA: diabetic ketoacidosis
DME: diabetic macular edema
DSME: diabetes self-management education
eAG: estimated average blood glucose
ECG: electrocardiogram
eGFR: estimated glomerular filtration rate
EMG: electromyogram
GFR: glomerular filtration rate
GRADE: Grading of Recommendations, Assessment, Development and Evaluation
HDL-C: high-density lipoprotein cholesterol
HTN: hypertension
IDF: International Diabetes Federation
K+: potassium
LDL-C: low-density lipoprotein cholesterol
MDRD: Modification of diet in renal disease study equation http://nkdep.nih.gov/professionals/gfr_calculators/orig_con.htm
MI: myocardial infarction
min: minutes
MNT: medical nutrition therapy
NGSP: National Glycohemoglobin Standardization Program
NYHA: New York Heart Association
PAD: peripheral artery disease
PAD: peripheral Arterial Disease
PAID: Problem Areas in Diabetes
PHQ-2 : Patient Health Questionnaire 2 questions
PHQ-9 : Patient Health Questionnaire, 9 questions
POC: point of care
PVD: peripheral vascular disease
RD: registered dietitian
RECORD study: Rosiglitazone Evaluated for Cardiac Outcomes and Regulation of Glycaemia in Diabetes
rMPI: radionuclide myocardial perfusion imaging
SBP: systolic blood pressure
SMBG: self-monitoring of blood glucose
TIA: transient ischemic attack
TLC: therapeutic lifestyle changes
UTI: urinary tract infection
VEGF: vascular endothelial growth factor

Approved by the Joslin Clinical Oversight Committee on June 6, 2017

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Grading System Used in Guidelines

Grade of Recommendation	Clarity of risk/benefit	Quality of supporting evidence
1A Strong recommendation High quality of evidence	Benefits clearly outweigh risk and vice versa.	Consistent evidence from well performed randomized, controlled trials or overwhelming evidence of some other form. Further research is unlikely to change our confidence in the estimate of benefit and risk.
1B Strong recommendation Moderate quality of evidence	Benefits clearly outweigh risk and burdens, or vice versa.	Evidence from randomized, controlled trials with important limitations (inconsistent results, methodological flaws, indirect or imprecise), or very strong evidence of some other research design. Further research is likely to have an impact on our confidence in the estimate of the benefit and risk and may change the estimate.
1C Strong recommendation Low quality of evidence	Benefits outweigh risk and burdens, or vice versa.	Evidence from observational studies, unsystematic clinical experience, or from randomized controlled trials with serious flaws. Any estimate of effect is uncertain.
2A Weak recommendation High quality of evidence	Benefits closely balanced with risks and burdens.	Consistent evidence from well performed randomized controlled trials or overwhelming evidence of some other form. Further research is unlikely to change our confidence in the estimate of benefit and risk.
2B Weak recommendation Moderate quality of evidence	Benefits closely balanced with risks and burdens; some uncertainty in the estimates of benefits, risks and burdens.	Evidence from randomized controlled trials with important limitations (inconsistent results, methodological flaws, indirect or imprecise), or very strong evidence of some other research design. Further research is likely to have an impact on our confidence in the estimate of benefit and risk and may change the estimate.
2C Weak recommendation Low quality of evidence	Uncertainty in the estimates of benefits, risks and burdens; benefits may be closely balanced with risks and burdens.	Evidence from observational studies, unsystematic clinical experience, or from randomized controlled trials with serious flaws. Any estimate of effect is uncertain.

Evidence graded less than “A” is acceptable to support clinical recommendations in a guideline. It is also assumed that for many important clinical recommendations, it would be unlikely that level A evidence be obtained because appropriate studies may never be performed.

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