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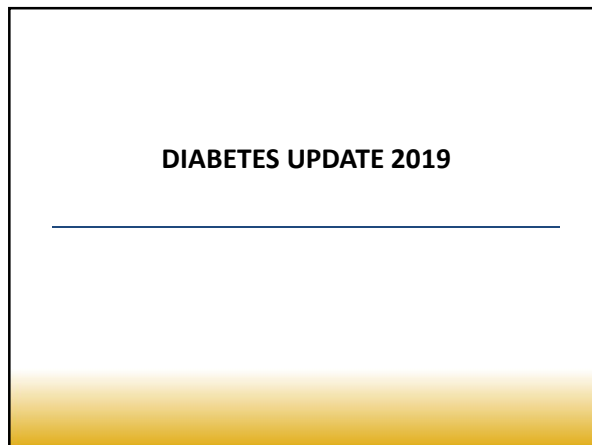
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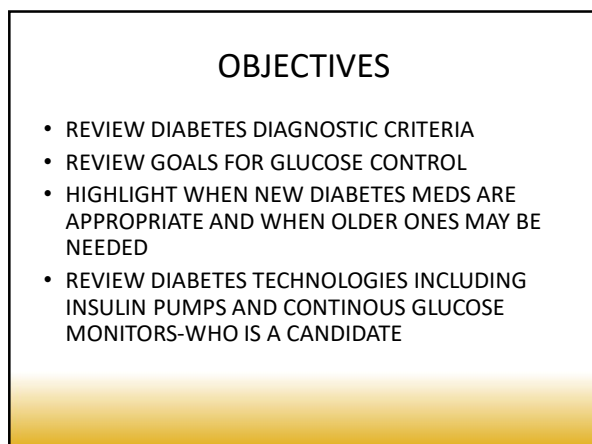
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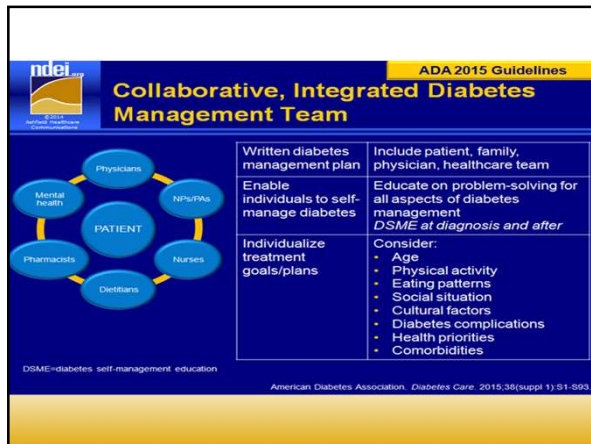
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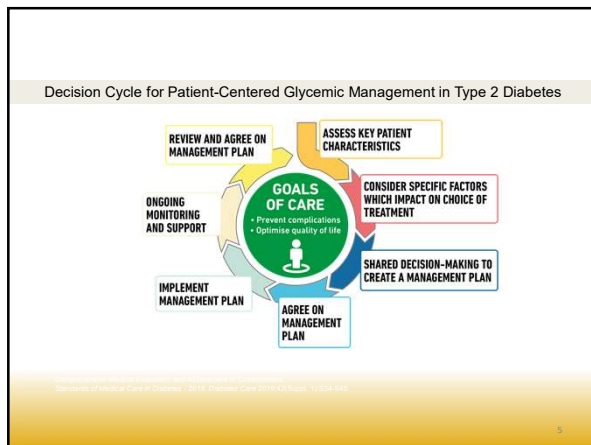
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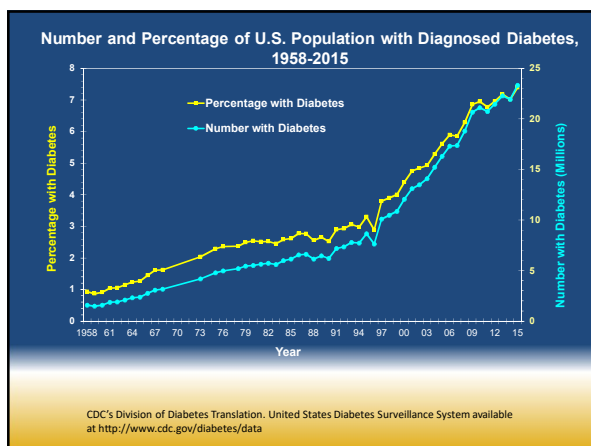
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## DIABETES NUMBERS

- **Prevalence:** In 2015, 30.3 million Americans, or 9.4% of the population, had diabetes.
  - Approximately 1.25 million American children and adults have type 1 diabetes.
- **Undiagnosed:** Of the 30.3 million adults with diabetes, 23.1 million were diagnosed, and 7.2 million were undiagnosed.
- **Prevalence in Seniors:** The percentage of Americans age 65 and older remains high, at 25.2%, or 12.0 million seniors (diagnosed and undiagnosed).
- **New Cases:** 1.5 million Americans are diagnosed with diabetes every year.
- **Deaths:** Diabetes remains the 7th leading cause of death in the United States in 2015, with 79,535 death certificates listing it as the underlying cause of death, and a total of 252,806 death certificates listing diabetes as an underlying or contributing cause of death.

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## PREDIABETES NUMBERS

- Eighty-six million people aged 20 years and older
- 1 in 3 American adults
- The percentage of U.S. adults with prediabetes is similar for non-Hispanic whites (35%), non-Hispanic blacks (39%), and Hispanics (38%)
- Without weight loss and moderate physical activity, 15-30% of people with prediabetes will develop type 2 diabetes within 5 years

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## COST OF DIABETES

- *Updated March 22, 2018*
- \$327 billion: Total costs of diagnosed diabetes in the United States in 2017
- \$237 billion for direct medical costs
- \$90 billion in reduced productivity
- After adjusting for population age and sex differences, average medical expenditures among people with diagnosed diabetes were 2.3 times higher than what expenditures would be in the absence of diabetes.
- Difficult to measure cost in relation to quality of life

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### CLASSIFICATION.

Diabetes can be classified into the following general categories:

1. Type 1 diabetes (due to autoimmune  $\beta$ -cell destruction, usually leading to absolute insulin deficiency)
2. Type 2 diabetes (due to a progressive loss of  $\beta$ -cell insulin secretion frequently on the background of insulin resistance)
3. Gestational diabetes mellitus (GDM) (diabetes diagnosed in the second or third trimester of pregnancy that was not clearly overt diabetes prior to gestation)
4. Specific types of diabetes due to other causes, e.g., monogenic diabetes syndromes (such as neonatal diabetes and maturity-onset diabetes of the young [MODY]), diseases of the exocrine pancreas (such as cystic fibrosis and pancreatitis), and drug- or chemical-induced diabetes (such as with glucocorticoid use, in the treatment of HIV/AIDS, or after organ transplantation)

Classification and Diagnosis of Diabetes:  
Standards of Medical Care in Diabetes - 2019. Diabetes Care 2019;42(Suppl. 1):S13-S28

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### Staging of Type 1 Diabetes

Table 2.1—Staging of type 1 diabetes (4,5)

	Stage 1	Stage 2	Stage 3
Stage	<ul style="list-style-type: none"><li>• Autoimmunity</li><li>• Normoglycemia</li><li>• Presymptomatic</li></ul>	<ul style="list-style-type: none"><li>• Autoimmunity</li><li>• Dysglycemia</li><li>• Presymptomatic</li></ul>	<ul style="list-style-type: none"><li>• New-onset hyperglycemia</li><li>• Symptomatic</li></ul>
Diagnostic criteria	<ul style="list-style-type: none"><li>• Multiple autoantibodies</li><li>• No IGT or IFG</li></ul>	<ul style="list-style-type: none"><li>• Multiple autoantibodies</li><li>• Dysglycemia: IFG and/or IGT</li><li>• FPG 100–125 mg/dL (5.6–6.9 mmol/L)</li><li>• 2-h PG 140–199 mg/dL (7.8–11.0 mmol/L)</li><li>• A1C 5.7–6.4% (39–47 mmol/mol) or <math>\geq 10\%</math> increase in A1C</li></ul>	<ul style="list-style-type: none"><li>• Clinical symptoms</li><li>• Diabetes by standard criteria</li></ul>

American Diabetes Association Standards of Medical Care in Diabetes.  
Classification and diagnosis of diabetes. Diabetes Care 2017; 40 (Suppl. 1): S11–S24

American Diabetes Association

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### DIAGNOSTIC CRITERIA FOR DM AND PREDIABETES

	Fasting Glucose	Random BG	A1c	GTT
Normal	< 99 mg/dl		3.5–5.6%	
Prediabetes	100–125 mg/dl		5.7–6.4%	
Diabetes	> 126 mg/dl	$\geq 200$ mg/dl	>6.5%	
Gestational		50 gm non-fasting 1 hour $\geq 140$ mg/dl		100 gm OGTT Fasting $\geq 95$ mg/dl 1 hr $\geq 180$ mg/dl 2 hr $\geq 155$ mg/dl 3 hr $\geq 140$ mg/dl

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## SCREENING RECOMMENDED

- All adults at 45, earlier if
  - BMI  $\geq 25$
  - First degree relative
  - Physical inactivity
  - High risk race/ethnicity
  - Women delivered > 9# baby or PCOS
  - HDL < 35
  - A1c > 5.7%
  - Hypertension
  - CVD
- Gestational Diabetes
  - Screen for undiagnosed DM at first prenatal visit based on DM risk criteria
  - Screen all women at 24-28 weeks for those not previously known to have diabetes
  - 6 to 12 weeks postpartum, rescreen for diabetes

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## CHILDREN AT RISK

- Screening to begin age 10 or puberty onset if
  - Weight is >120% ideal body weight
  - Plus any two risk factors
    - Family history of type 2 dm
    - Race/ethnicity (Native American, African American, Latino, Asian American, Pacific Islander)
    - Signs of insulin resistance: acanthosis nigricans, hypertension, dyslipidemia, PCOS, small for gestational age
    - Maternal history of DM or GDM during gestation

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## Acanthosis Nigricans



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## ADA/AACE GLUCOSE GOALS

Glycemic control & A1c Target		
	ADA	AACE
A1c (%)	<7	<6.5
Preprandial (mg/dl)	80-120	<110
Postprandial (mg/dl)	140-180	<140
Bedtime (mg/dl)	100-140	100-140

ADA: American Diabetes Association  
 AACE: American Association of Clinical Endocrinologists

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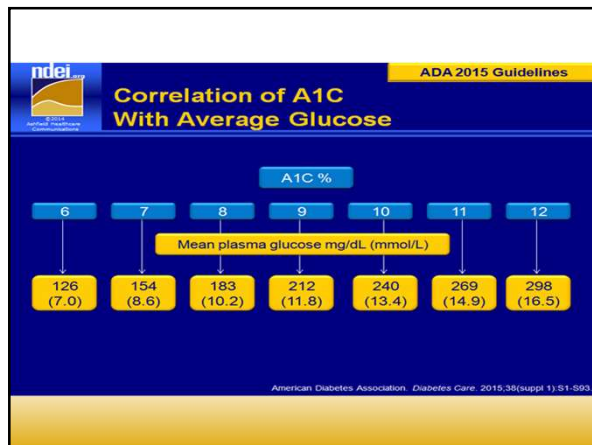
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## INDIVIDUALIZATION OF GOALS

- More stringent targets if able to achieve without significant hypoglycemia
- Less stringent targets for those with history of severe hypoglycemia, limited life expectancy, advanced micro or macrovascular disease, limited life expectancy

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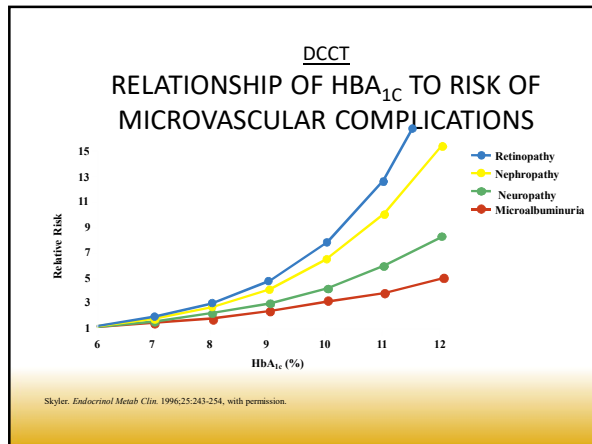
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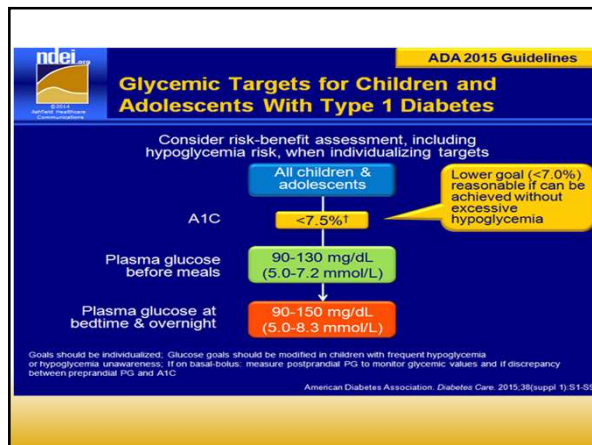
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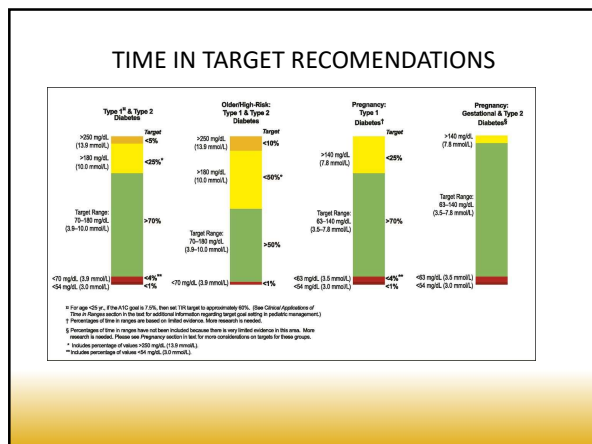
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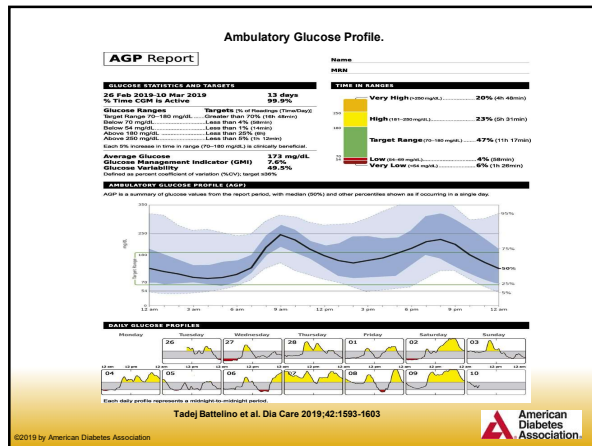
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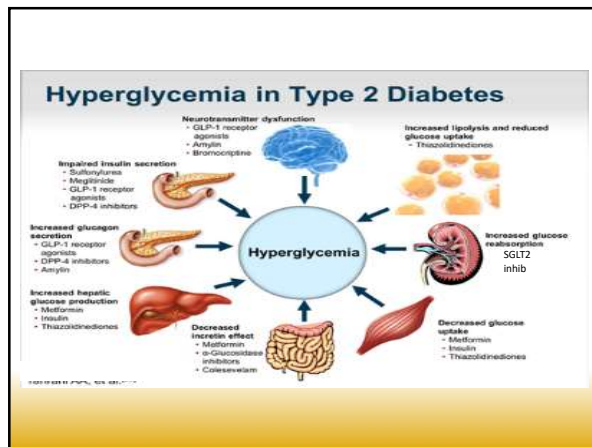
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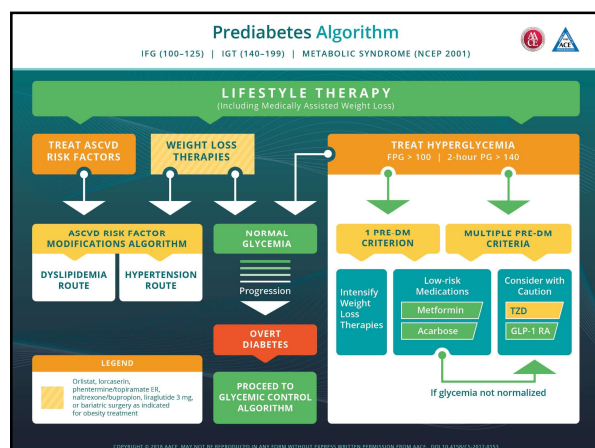


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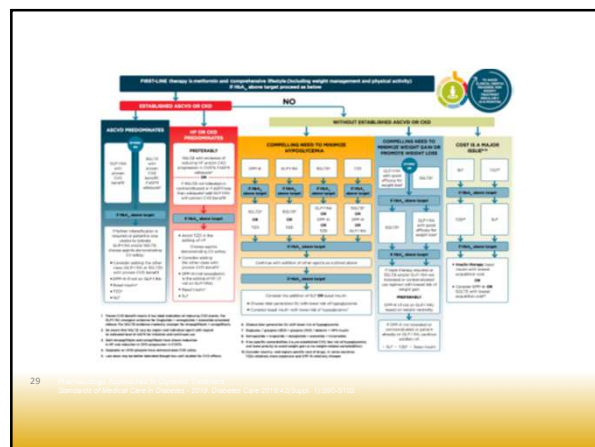
Table 1.3—Drug-specific and patient factors to consider when selecting antihyperglycemic treatment in adults with type 2 diabetes									
	Indication	Contraindication	Relative efficacy	Relative safety	Relative cost	Relative convenience	Relative overall benefit	Additional considerations	
Insulin	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
GLP-1 receptor agonists	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
SGLT2 inhibitors	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
Thiazolidinediones	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
Metformin	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
DPP-4 inhibitors	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
Sulfonylureas	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
Meglitinides	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain
	Yes	No	High	Low	Low	High	High	1. Associated with hypoglycemia	2. Associated with weight gain

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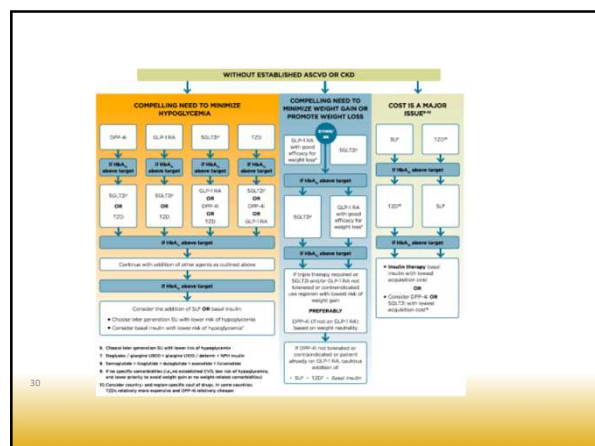




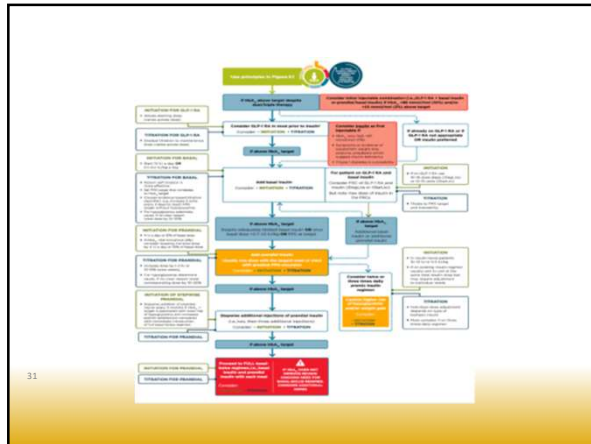
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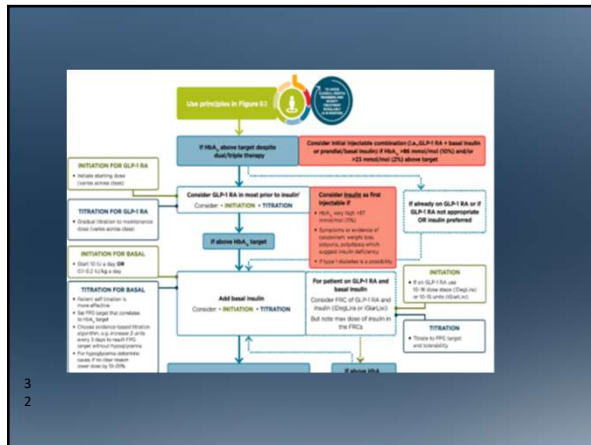
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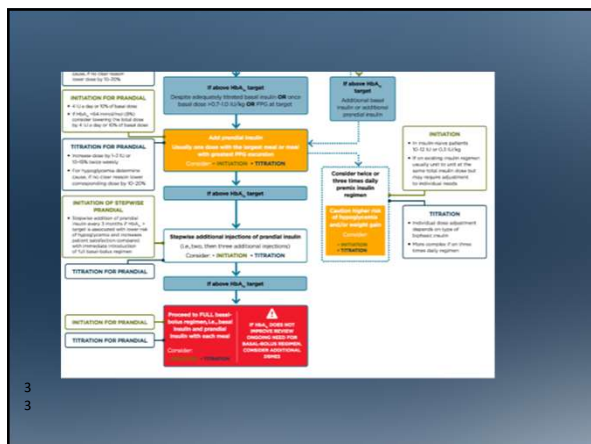
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# Insulin Therapy in T2DM

- The progressive nature of T2DM should be regularly & objectively explained to T2DM patients.
- Avoid using insulin as a threat, describing it as a failure or punishment.
- Give patients a self-titration algorithm.

American Diabetes Association Standards of Medical Care in Diabetes.  
Approaches to glycemic treatment. Diabetes Care 2017; 40 (Suppl. 1): S64-S74

 American Diabetes Association

# Combination Injectable Therapy in T2DM

### New Recommendation: Pharmacologic Therapy For T2DM

- In patients with long-standing suboptimally controlled type 2 diabetes and established atherosclerotic cardiovascular disease, empagliflozin or liraglutide should be considered as they have been shown to reduce cardiovascular and all-cause mortality when added to standard care. Ongoing studies are investigating the cardiovascular benefits of other agents in these drug classes. **B**

American Diabetes Association Standards of Medical Care in Diabetes.  
Approaches to glycemic treatment. Diabetes Care 2017; 40 (Suppl. 1): S64-S74

American Diabetes Association

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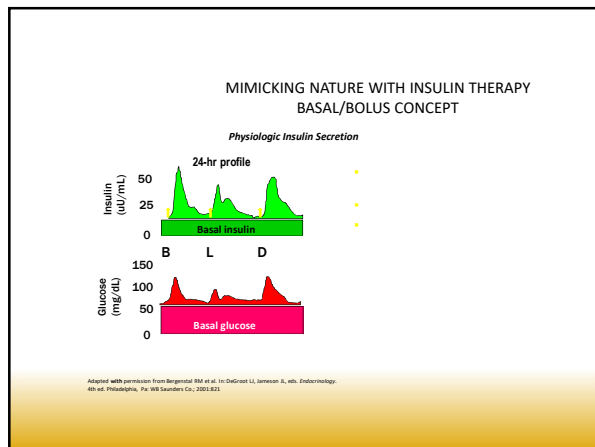
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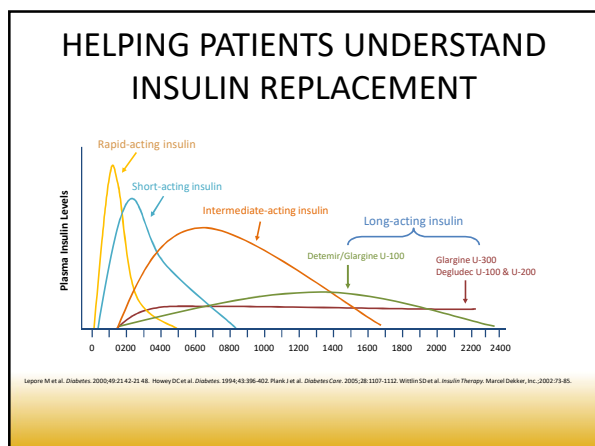
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CURRENTLY AVAILABLE BASAL INSULINS

	NPH Insulin	Insulin Glargine U-100 & U-300	Insulin Detemir
Insulin Type	Human; intermediate-acting	Analogue; long-acting	Analogue; long-acting
Onset	1-4 hours	N/A	N/A
Peak	4-10 hours	No pronounced peak	Relatively flat
Effective Duration	10-16 hours	U-100: Up to 24 hrs U-300: Beyond 24 hrs (pen only)	Up to 24 hours

Lilly US00 regular similar in duration of NPH

Lucifora D, et al. Diabetes Care. 2011;34:1312-1314.  
Nawander K, et al. Clin Diabetes. 2009;27:68-69.  
Tibber J, et al. EASD September 23-27 2013; Barcelona, Spain, (Abstract 1023).  
Jee T, et al. EASD September 23-27 2013; Barcelona, Spain, (Abstract 1029).

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CURRENTLY AVAILABLE SHORT-ACTING PRANDIAL INSULINS

	Regular Insulin	Insulin Lispro U100 U200 (pen only)	Insulin Aspart	Insulin Glulisine
Insulin type	Human; short-acting	Analogue; rapid acting	Analogue; rapid acting	Analogue; rapid acting
Onset, hr	0.5-1	< 0.3-0.5	< 0.25	< 0.25
Peak, hr	2-3	0.5-2.5	0.5-1.0	1-1.5
Effective duration, hr	3-6	3-6.5	3-5	3-5
Injection: meal timing, min	-30 to +45	-15 to immediately after	-5 to -10	-15 to +20

American Diabetes Association. Practical Insulin: A Handbook for Prescribing Providers. 3rd ed. 2011:1-69.

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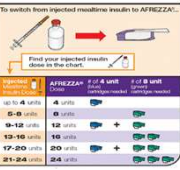
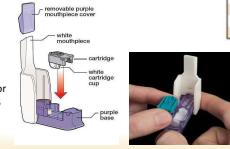

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INHALED INSULIN - AFREZZA



Spirometry prior to starting, at 6 months, and then annually

AFREZZA<sup>®</sup> (prescribing information) Bridgewater, NJ: Sanofi-Aventis U.S. LLC, 2015.

Now have 12 unit dosing available

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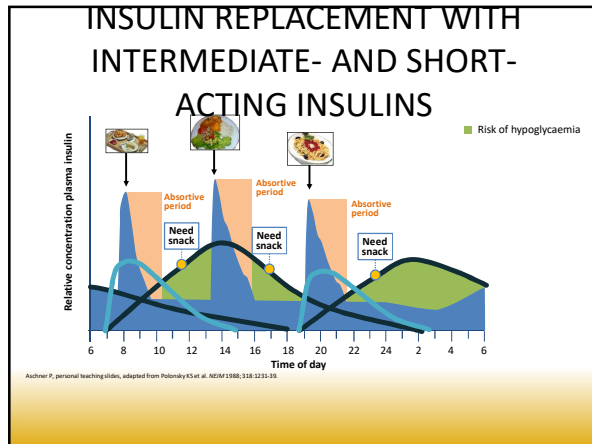
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**ADA 2015 Guidelines**

## Pharmacologic Therapy for Type 1 Diabetes Management

<b>Insulin treatment for most patients with type 1 diabetes</b> <ul style="list-style-type: none"> <li>Treat with multiple-dose insulin injections (3-4 injections/day of basal and prandial insulin) or continuous subcutaneous insulin infusion</li> <li>Match prandial insulin dose to carbohydrate intake, premeal blood glucose, and anticipated activity</li> <li>Use insulin analogs to reduce risk of hypoglycemia</li> <li>Consider using sensor-augmented low glucose suspend threshold pump in patients with frequent nocturnal hypoglycemia and/or hypoglycemia unawareness</li> </ul>	<table border="1" style="width: 100%; border-collapse: collapse;"> <tr> <th style="background-color: #0070c0; color: white;">Other agents</th> <th style="background-color: #0070c0; color: white;">Investigational agents</th> </tr> <tr> <td style="padding: 5px;"> <b>Pramlintide (amylin analog)</b> <ul style="list-style-type: none"> <li>Delays gastric emptying</li> <li>Blunts pancreatic secretion of glucagon</li> <li>Enhances satiety</li> <li>Induces weight loss</li> <li>Lowers insulin dose</li> <li>Use only in adults</li> </ul> </td> <td style="padding: 5px;"> <b>Metformin + insulin</b> <ul style="list-style-type: none"> <li>Reduces insulin requirements &amp; improves metabolic control in obese/overweight with poor glycemic control</li> </ul> <b>Incretins</b> <ul style="list-style-type: none"> <li>GLP-1 receptor agonists</li> <li>DPP-4 inhibitors</li> <li>SGLT2 inhibitors</li> </ul> </td> </tr> </table>	Other agents	Investigational agents	<b>Pramlintide (amylin analog)</b> <ul style="list-style-type: none"> <li>Delays gastric emptying</li> <li>Blunts pancreatic secretion of glucagon</li> <li>Enhances satiety</li> <li>Induces weight loss</li> <li>Lowers insulin dose</li> <li>Use only in adults</li> </ul>	<b>Metformin + insulin</b> <ul style="list-style-type: none"> <li>Reduces insulin requirements &amp; improves metabolic control in obese/overweight with poor glycemic control</li> </ul> <b>Incretins</b> <ul style="list-style-type: none"> <li>GLP-1 receptor agonists</li> <li>DPP-4 inhibitors</li> <li>SGLT2 inhibitors</li> </ul>
Other agents	Investigational agents				
<b>Pramlintide (amylin analog)</b> <ul style="list-style-type: none"> <li>Delays gastric emptying</li> <li>Blunts pancreatic secretion of glucagon</li> <li>Enhances satiety</li> <li>Induces weight loss</li> <li>Lowers insulin dose</li> <li>Use only in adults</li> </ul>	<b>Metformin + insulin</b> <ul style="list-style-type: none"> <li>Reduces insulin requirements &amp; improves metabolic control in obese/overweight with poor glycemic control</li> </ul> <b>Incretins</b> <ul style="list-style-type: none"> <li>GLP-1 receptor agonists</li> <li>DPP-4 inhibitors</li> <li>SGLT2 inhibitors</li> </ul>				

Slide covers investigational agents not yet FDA approved for the treatment of type 1 diabetes in the United States  
American Diabetes Association. Diabetes Care. 2015;38(suppl 1):S1-S93.

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## HYPOGLYCEMIA

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## CLASSIFICATION OF HYPOGLYCEMIA

Table 6.3—Classification of hypoglycemia (61)

Level	Glycemic criteria	Description
Glucose alert value (level 1)	$\leq 70$ mg/dL (3.9 mmol/L)	Sufficiently low for treatment with fast-acting carbohydrate and dose adjustment of glucose-lowering therapy
Clinically significant hypoglycemia (level 2)	$< 54$ mg/dL (3.0 mmol/L)	Sufficiently low to indicate serious, clinically important hypoglycemia
Severe hypoglycemia (level 3)	No specific glucose threshold	Hypoglycemia associated with severe cognitive impairment requiring external assistance for recovery

Glycemic targets. Diabetes Care 2017; 40 (Suppl. 1): S48-S56

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## RISK FACTORS FOR HYPOGLYCAEMIA IN THE OLDER ADULT

### HYPOGLYCAEMIA IN THE OLDER ADULT

Use of insulin or insulin secretagogues

Erratic meals

Duration of diabetes

Antecedent hypoglycaemia

Hospital discharge within the preceding 30 days

Comorbidities such as renal insufficiency

Poly-pharmacy ( $\geq 5$  concurrent medications)

Cognitive decline, depression

Shorr RI et al. Arch Intern Med. 1997;157:1682-86. Zammittini N, Frer BM. Diabetes Care. 2005; 28:2948-61.

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## NOCTURNAL HYPOGLYCAEMIA

- During sleep, the neuroendocrine response against hypoglycaemia is markedly blunted (the response threshold is shifted to lower glucose levels)
- While symptoms of hypoglycaemia trigger awakening in healthy subjects, individuals with type 1 diabetes frequently fail to respond to symptoms during sleep

Jauch-Chara V, et al. Best Pract Res Clin Endocrinol Metab 2010;24:805-815. Jones TW et al. N Engl J Med 1998;338:1857. Banerjee S, Cryer PE. Diabetes 2003;52:1195.

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## RISK FACTORS FOR NOCTURNAL HYPOGLYCEMIA

NOCTURNAL  
HYPOGLYCAEMIA

Intensive Insulin Management
Exercise
Bedtime blood glucose level
Daytime hypoglycaemia
Premix insulin twice daily/NPH insulin
Alcohol
Impaired sympathoadrenal response during sleep
Children who go to bed early
Use of sulfonylureas (type 2 diabetes)

Wilson DM et al. Factors Associated with Nocturnal Hypoglycemia in At-Risk Adolescents and Young Adults with Type 1 Diabetes. *Diabet Technol Ther* 2015; 17:385-95.  
Bae JP et al. Risk factors for nocturnal hypoglycemia in insulin-treated patients with type 2 diabetes. *Clin Ther*. 2017; 39:1790-95.

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## HYPOGLYCEMIA

- Individuals at risk for hypoglycemia should be asked about symptomatic and asymptomatic hypoglycemia at each visit
  - At what number do you feel your low blood glucose
  - Have you had any hypoglycemia that required the assistance of another person or glucagon
  - Not limited to only those on insulin

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ADA 2015 Guidelines

### Management of Hypoglycemia

At-risk patients	Ask about symptomatic and asymptomatic hypoglycemia at each encounter
Preferred treatment: glucose (15-20 g)*	
<ul style="list-style-type: none"> <li>After 15 mins of treatment, repeat if SMBG shows continued hypoglycemia</li> <li>When SMBG normal: patient should consume meal or snack to prevent recurrence</li> </ul>	
Prescribe glucagon if significant risk of severe hypoglycemia	
Hypoglycemia unawareness or episode of severe hypoglycemia	<ul style="list-style-type: none"> <li>Reevaluate treatment regimen</li> <li>Insulin-treated patients: raise glycemic targets for several weeks to partially reverse hypoglycemia unawareness and reduce recurrence</li> </ul>
Low or declining cognition	Continually assess cognitive function with increased vigilance for hypoglycemia

\*Any form of glucose-containing carbohydrate can be used; SMBG= self-monitoring of blood glucose  
American Diabetes Association. *Diabetes Care*. 2015;38(suppl 1):S1-S99.

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## TECHNOLOGY

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## INSULIN INFUSION PUMP OPTIONS



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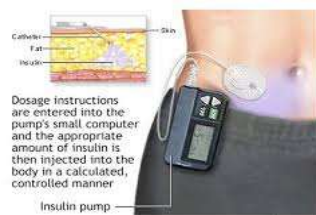
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## INSULIN PUMP-HOW IT WORKS



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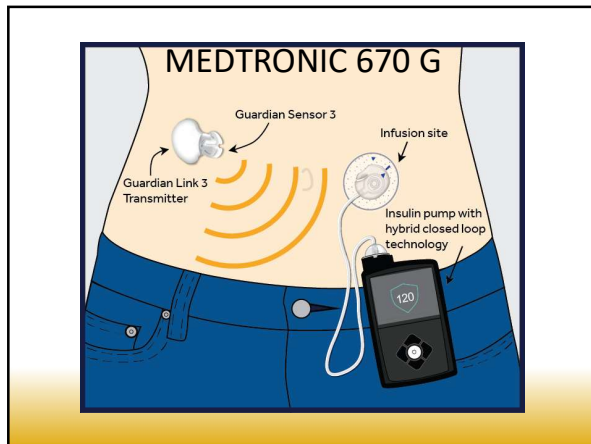
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## DEXCOM PERSONAL KR



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## DEXCOM DIAGNOSTIC HS

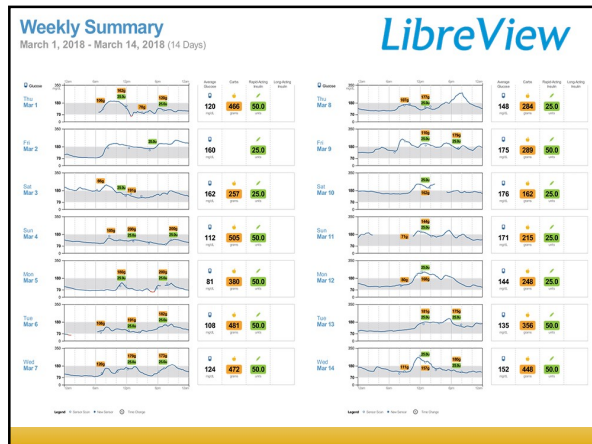


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## FREESTYLE LIBRE



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### Recommendations: Glucose Monitoring (2)

- Most patients on multiple-dose insulin (MDI) or insulin pump therapy should do SMBG **B**
  - Prior to meals and snacks
  - At bedtime
  - Prior to exercise
  - When they suspect low blood glucose
  - After treating low blood glucose until they are normoglycemic
  - Prior to critical tasks such as driving
  - Occasionally postprandially

American Diabetes Association Standards of Medical Care in Diabetes. Glycemic targets. Diabetes Care 2017, 40 (Suppl. 1): S48-S56.

American Diabetes Association

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### Recommendations: Glucose Monitoring (3)

- When used properly, CGM in conjunction with intensive insulin regimens is a useful tool to lower A1C in selected adults (aged  $\geq 25$  years) with type 1 diabetes. **A**
- Although the evidence for A1C lowering is less strong in children, teens, and younger adults, CGM may be helpful in these groups. Success correlates with adherence to ongoing use of the device. **B**
- CGM may be a supplemental tool to SMBG in those with hypoglycemia unawareness and/or frequent hypoglycemic episodes. **C**

American Diabetes Association Standards of Medical Care in Diabetes. Glycemic targets. Diabetes Care 2017, 40 (Suppl. 1): S48-S56.

American Diabetes Association

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## Recommendations: Glucose Monitoring (4)

- Given variable adherence to CGM, assess individual readiness for continuing use of CGM prior to prescribing. **E**
- When prescribing CGM, robust diabetes education, training, and support are required for optimal CGM implementation and ongoing use. **E**
- People who have been successfully using CGM should have continued access after they turn 65 years of age. **E**

American Diabetes Association Standards of Medical Care in Diabetes.  
Glycemic targets. Diabetes Care 2017; 40 (Suppl. 1): S48-S56

American Diabetes Association

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ADA 2015 Guidelines

### Continuous Glucose Monitoring (CGM)

Useful for A1C lowering in select adults (aged ≥25 yrs) with type 1 diabetes requiring intensive insulin regimens	<ul style="list-style-type: none"> <li>• May be useful among children, teens, and younger adults*</li> <li>• Success related to adherence to ongoing use</li> </ul>
May be a useful supplement to SMBG among patients with	<ul style="list-style-type: none"> <li>• Hypoglycemia unawareness and/or</li> <li>• Frequent hypoglycemic episodes</li> </ul>
Variable adherence to CGM	<ul style="list-style-type: none"> <li>• Assess individual readiness for continuing prior to prescribing</li> <li>• Robust diabetes education, training, support critical for optimal CGM implementation</li> </ul>

\*Evidence for A1C lowering less strong in these populations  
SMBG=Self-monitoring of blood glucose

American Diabetes Association. Diabetes Care. 2015;38(suppl 1):S1-S93.

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<https://professional.diabetes.org/content-page/practice-guidelines-resources>

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Childs, B, Cypress, M, Spollett G (Ed) Complete Nurses Guide to Diabetes Care, 3<sup>rd</sup> Ed, Arlington, VA: American Diabetes Association, 2017

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