



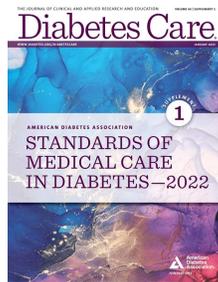
Advancing Your Career in Diabetes Education

ADA/EASD Meds Update, Solving Glucose Mysteries, 3 Steps to DeFEET Amputations

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Coach Bev has no Conflict of Interest

- ▶ She's not on any speaker's bureau
- ▶ Does not invest or have any financial relationships with diabetes related companies.
- ▶ Gathers information from reading package inserts, research and articles
- ▶ The ADA Standards of Medical Care is main resource for course content



Diabetes Overview and Glycemic Goals

Objectives:

1. Discuss current state of diabetes in the U.S.
2. Describe goals of care.
3. Summarize the new ADA/ EASD Management Guidelines.
4. Apply new guidelines to individual case studies.



Global Epidemic

Diabetes is spiralling out of control
1 in 10 adults are living with diabetes. Almost half are undiagnosed.

537 million adults are living with diabetes

3 in 4 adults with diabetes live in low- and middle-income countries

Diabetes around the world in 2021

International Diabetes Federation

www.DiabetesAtlas.org

World Diabetes Day is November 14

CDC Announces

35% of Americans will have Diabetes by 2050

Boyle, Thompson, Barker, Williamson
2010, Oct 22:8(1)29
www.pophealthmetrics.com

Diabetes in America 2022 - CDC

- ▶ 11% of adults have diabetes (37.3 mil)
- ▶ 23% of those don't know they have diabetes
- ▶ 38% of adults have prediabetes (96 mil)
- ▶ 19% reported being told they have prediabetes.

Figure 1. Trends in age-adjusted prevalence of diagnosed diabetes, undiagnosed diabetes, and total diabetes among adults aged 18 years or older, United States, 2001-2020.

CDC 2022 Report
<https://www.cdc.gov/diabetes/data/statistics-report/diagnosed-diabetes.html>

www.DiabetesEd.net

Geography of Diabetes, Poverty, Weight – Diabetes Belt

Americans Living in Poverty

<https://stacks.cdc.gov/view/cdc>

CDC Identifies diabetes belt

Search for:

What states are in the diabetes belt?

The 15 states that have counties in the diabetes belt are **Alabama, Arkansas, Florida, Georgia, Kentucky, Louisiana, Mississippi, North Carolina, Ohio, Pennsylvania, South Carolina, Tennessee, Texas, Virginia and West Virginia.** Mar 8, 2011

Diagnosed Diabetes by Ethnic Group

Highest prevalence among

- Indigenous people
- Mexican and Puerto Ricans
- Asian Indians and Filipinos

<https://www.cdc.gov/diabetes/data/statistics-report/diagnosed-diabetes.html>

Figure 2. Age-adjusted estimated prevalence of diagnosed diabetes by race/ethnicity group and sex for adults aged 18 years or older, United States, 2018-2019

Race/Ethnicity Group	Men (%)	Women (%)
American Indian/Alaska Native	~14.5	~15.0
Asian, Non-Hispanic	~11.0	~9.0
Black, Non-Hispanic	~12.5	~12.0
Hispanic	~11.5	~12.0
White, Non-Hispanic	~8.0	~7.0

Socioeconomics – Diabetes Diagnosis

Prevalence varied significantly by education level, an indicator of SES status

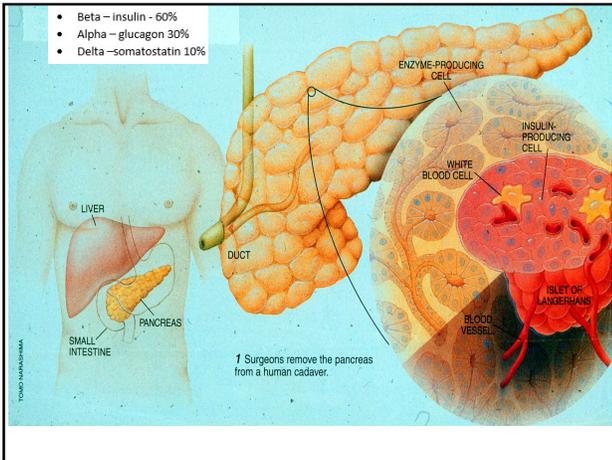
- 13.4% - Less than high school education
- 9.2% - High school education
- 7.1% - More than high school education

CDC 2022

1. Improving Care and Promoting Health in Populations

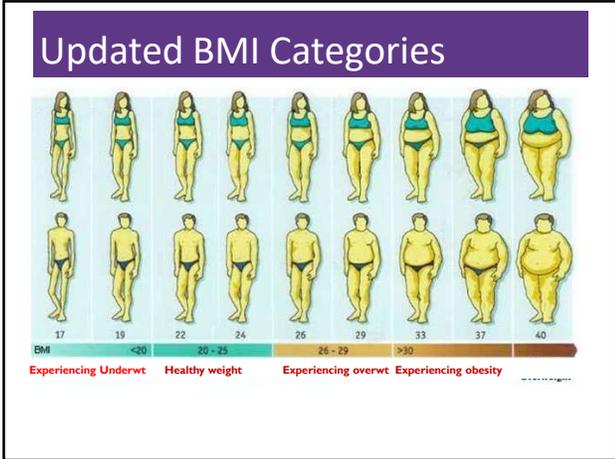
- ▶ Population Health includes:
 - ▶ Outcomes (mortality, morbidity)
 - ▶ Disease burden (incidence and prevalence)
 - ▶ Behavioral and metabolic factors (A1c, MNT, exercise)
- ▶ Diabetes annual cost 2017 - \$327 bil
- ▶ Targets
 - ▶ 64% of ind's met A1c targets
 - ▶ 70% achieved BP targets
 - ▶ 57% met LDL target
 - ▶ In total, 23% met all targets

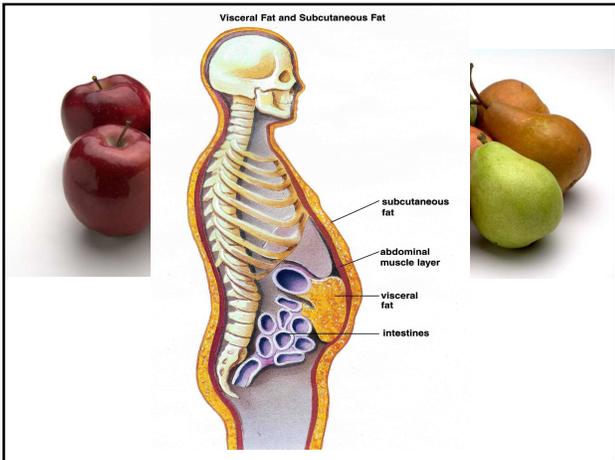




Hormones Effect on Glucose

Hormone	Effect
▶ Glucagon (pancreas)	⬆️
▶ Stress hormones (kidney)	⬆️
▶ Epinephrine (kidney)	⬆️
▶ Insulin (pancreas)	⬇️
▶ Amylin (pancreas)	⬇️
▶ Gut hormones - incretins (GLP-1) released by L cells of intestinal mucosa, beta cell has receptors)	⬇️

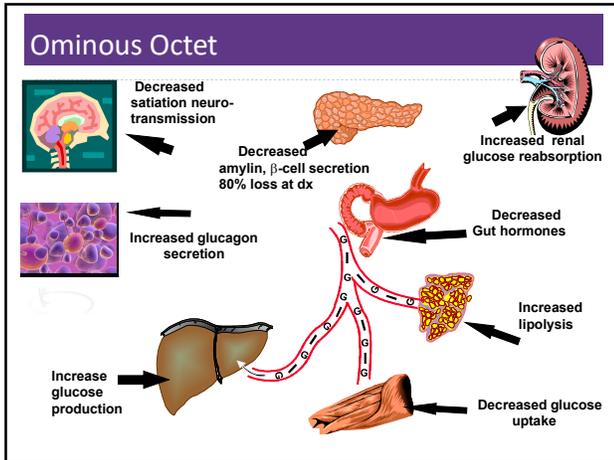




What is Type 2 Diabetes?

► Complex metabolic disorder
 (Insulin resistance and deficiency)
 with social, behavioral and environmental risk factors unmasking the effects of genetic susceptibility.

New Diagnosis?
 Call 800 – DIABETES to request "Getting Started Kit"
www.Diabetes.org



ABCs of Diabetes

- ▶ **A**1c less than 7% (avg 3 month BG)
 - ▶ Pre-meal BG 80-130
 - ▶ Post meal BG <180
- ▶ **B**lood Pressure < 140/90
 - ▶ BP target <130/80
 - ▶ With CVD or if 10 year CVD Risk > 15%
- ▶ **C**holesterol
 - ▶ Statin therapy

A1c and Estimated Avg Glucose (eAG)

A1c (%)	eAG
5	97 (76-120)
6	126 (100-152)
7	154 (123-185)
8	183 (147-217)
9	212 (170 -249)
10	240 (193-282)
11	269 (217-314)
12	298 (240-347)

eAG = 28.7 x A1c - 46.7 ~ 29 pts per 1%
Translating the A1c Assay Into eAG - ADAG Study

6. Glycemic Targets: *Standards of Medical Care in Diabetes—2020*
American Diabetes Association
Diabetes Care 2020; Jan; 43(Supplement 1): S66-S76.
<https://doi.org/10.2337/14C00006>

Ambulatory Glucose Profile

► Standardized report with visual cues for those on CGM devices

- For most with type 1 or type 2 diabetes
 - > 70% of readings within BG range of 70-180mg/dL
 - < 4% of readings < 70 mg/dL
 - < 1% of readings < 54 mg/dL
 - < 25% of readings > 180 mg/dL
 - < 5% of readings > 250 mg/dL



- For under 25 years, with A_{1c} goal is < 7.5%, time-in-range target is set to about 60%.

NEW! ADA & European Association for the Study of Diabetes (EASD) Consensus Management of Hyperglycemia in Type 2



Sept 23, 2022

Davies MJ, Arora VR, Collins BS, Gabbay RA, Green J, Maruthur NM, Rosas SE, Del Prato S, Mathieu C, Mingrone G, Rossing P, Tankova T, Tsapas A, Buse JB. Diabetes Care 2022; <https://doi.org/10.2337/DC22-0034>. Diabetologia 2022; <https://doi.org/10.1007/s00125-022-05787-2>.

ADA & European Association for the Study of Diabetes (EASD) Consensus Management of Hyperglycemia Type 2



Management of Hyperglycemia in Type 2 Diabetes, 2022. A Consensus Report by the

Davies MJ, Arora VR, Collins BS, Gabbay RA, Green J, Maruthur NM, Rosas SE, Del Prato S, Mathieu C, Mingrone G, Rossing P, Tankova T, Tsapas A, Buse JB. *Diabetes Care* 2022; <https://doi.org/10.2337/DC22-0034>.

Diabetologia 2022; <https://doi.org/10.1007/s00125-022-05787-2>.

Choice of Glucose Lowering Meds

- ▶ Person centered
- ▶ Co-conditions
- ▶ Values and needs
- ▶ Reduce risk
 - ▶ CV and renal
- ▶ Affordable
- ▶ Weight impact



Metformin, Sulfonylureas & DPP-IV Inhibitors



Quick Question 1

▶ KR had GDM, now has prediabetes. Started on Metformin 500mg BID. Which of the following is true?



- Metformin can cause kidney problems.
- If you forget to take metformin before the meal, hold the dose
- Metformin may cause loose stools
- Stop metformin if GFR is less than 60.

Medication Taking Behaviors

- ▶ Adequate medication taking is defined as 80%
- ▶ 23% of time, if A1c, B/P, lipids above target - due to med taking behavior
- ▶ Assess for barriers
- ▶ If taking meds 80% of time and goals not met, consider medication intensification



Barriers include:
Forgetting to fill Rx, forgetting to take, fear, depression, health beliefs, med complexity, cost, knowledge gap, system factors, etc.

Work on targeted approach for specific barrier

Sulfonylureas

- ▶ Mechanism: Stimulate beta cells to release insulin
- ▶ Dosed 1-2x daily before meals
- ▶ Adverse effects
 - ▶ Hypoglycemia, Weight gain, watch renal function
- ▶ Low cost, \$12 for 3 months supply



Sulfonylureas • Stimulates sustained insulin release	glyburide: (Diabeta) (Glynase PresTabs)	1.25 – 20 mg 0.75 – 12 mg	Can take once or twice daily before meals. Low cost generic. Side effects: hypoglycemia and weight gain. Eliminated via kidney. Caution: Glyburide most likely to cause hypoglycemia. Lowers A1c 1.0% – 2.0%.
	glipizide: (Glucotrol) (Glucotrol XL)	2.5 – 40 mg 2.5 – 20 mg	
	glimepiride (Amaryl)	1.0 – 8 mg	

Case Study KR – Poll 2

KR is a 47yoM with type 2 diabetes x 5 years. Complains of dizziness/shakiness 3x/week, especially after surfing. Last A1C=6.7%. Which of their medications is most likely causing hypoglycemia?

- Metformin
- Sitagliptin (Januvia)
- Glimepiride (Amaryl)
- Pioglitazone (Actos)



Preventing Hypoglycemia

Nocturnal Lows

- ▶ If bedtime glucose <110, **reduce meds**
- ▶ If increased daytime activity, may need extra hb snack
- ▶ Eval pre-dinner insulin/meds

Other

- ▶ Monitor kidney function / wt loss
- ▶ Monitor BG trends
- ▶ Too much meds?
- ▶ Skipped /delayed meals?
- ▶ Plan ahead
- ▶ Alcohol precautions
- ▶ Exercise planning



DPP-4 Inhibitors – “Incretin Enhancers”

Januvia (sitagliptin) Tradjenta (linagliptin)
Onglyza (saxagliptin) Nesina (alogliptin)

DPP-4 Inhibitors “Incretin Enhancers” <ul style="list-style-type: none"> • Prolongs action of gut hormones • Increases insulin secretion • Delays gastric emptying 	sitagliptin (Januvia)	25 - 100 mg daily – eliminated via kidney*	* If creat elevated, see med insert for dosing. Side effects: headache and flu-like symptoms. Can cause severe, disabling joint pain. Contact MD, stop med. Report signs of pancreatitis. † Saxagliptin and alogliptin can increase risk of heart failure. Notify MD for shortness of breath, edema, weakness, etc. No wt gain or hypoglycemia. Lowers A1c 0.6%-0.8%.
	saxagliptin (Onglyza)†	2.5 - 5 mg daily – eliminated via kidney*, feces	
	linagliptin (Tradjenta)	5 mg daily – eliminated via feces	
	alogliptin (Nesina)†	6.25 - 25 mg daily – eliminated via kidney*	

Atherosclerotic Cardiovascular Risk Reduction – High Importance

USE OF GLUCOSE-LOWERING MEDICATIONS IN THE MANAGEMENT OF TYPE 2 DIABETES

HEALTHY LIFESTYLE BEHAVIORS; DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT (DSMES); SOCIAL DETERMINANTS OF HEALTH (SDOH)

Goal: Cardiovascular Risk Reduction in High-Risk Patients with Type 2 Diabetes (in addition to comprehensive CV risk management)

Goal: Achievement and Maintenance of Glycemic and Weight Management Goals

Goal: Cardiovascular Risk Reduction in High-Risk Patients with Type 2

+ASCVD¹
 Defined differently across CVOTs but all included individuals with established CVD (e.g., MI, stroke, any revascularization procedure). Variably included: conditions such as transient ischemic attack, unstable angina, amputation, symptomatic or asymptomatic coronary artery disease.

+Indicators of high risk
 While definitions vary, most comprise ≥55 years of age with two or more additional risk factors (including obesity, hypertension, smoking, dyslipidemia, or albuminuria)

Management of Hyperglycemia in Type 2 Diabetes, 2022. A Consensus Report by the

Choosing glucose-lowering medication in people with CVD

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+ASCVD/Indicators of High Risk

GLP-1 RA^a with proven CVD benefit EITHER/OR SGLT2i^b with proven CVD benefit

↓

If HbA_{1c} above target

↓

- For patients on a GLP-1 RA consider adding SGLT2i with proven CVD benefit or vice versa
- TZD^c

Most effective meds based on Cardiovascular Outcomes Trial (CVOT)

GLP-1 RA's Preferred
 semaglutide (Ozempic), liraglutide (Victoza), dulaglutide (Trulicity)

~ Or ~

SGLT2i
 Empagliflozin (Jardiance), canagliflozin (Invokana), dapagliflozin (Farxiga)

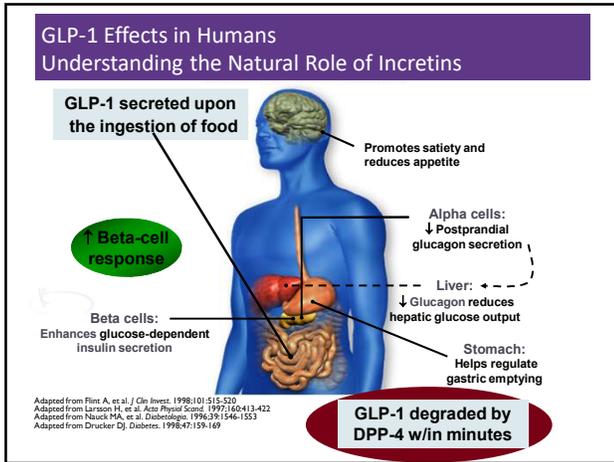
ASCVD = atherosclerotic cardiovascular disease

Management of Hyperglycemia in Type 2 Diabetes, 2022. A Consensus Report by the

GLP & GIP Receptor Agonists



Reduce Major Adverse Cardiovascular Events (MACE)



Pocket Card: GLP-1 & GIP RA

GLP-1 & GIP Receptor Agonists

Class/Main Action	Name	Dose Range	Considerations
GLP-1 Receptor Agonist (GLP-1 RA) "Incretin Mimetic" • Increases insulin release with food • Slows gastric emptying • Promotes satiety • Suppresses glucagon	exenatide (Byetta)	5 and 10 mcg BID	Side effects for all: Nausea, vomiting, weight loss, injection site reaction. Report signs of acute pancreatitis (severe abdominal pain, vomiting), stop med. Increase dose monthly to achieve targets. Black box warning: Thyroid C-cell tumor warning (avoid if family history of medullary thyroid tumor). *Significantly reduces risk of CV death, heart attack, and stroke. *Approved for pediatrics 10-17 yrs Lowers A1c 0.5 – 1.6% Weight loss of 1.6 to 6.0 kgs
	exenatide XR* (Bydureon)	2 mg 1x a week Pen injector - Bydureon BCise	
	liraglutide (Victoza)**	0.6, 1.2 and 1.8 mg daily	
	dulaglutide* (Trulicity)	0.75, 1.5, 3.0 and 4.5 mg 1x a week pen injector	
	lixisenatide (Adlyxin)	10 mcg 1x a day for 14 days 20 mcg 1x day starting day 15	
Dual Incretin Agonist Combines both GLP-1 and GIP Incretins. Same action profile as GLP-1 RA, with more intensive action profile.	semaglutide* (Ozempic)	0.5, 1.0 and 2.0 mg 1x a week pen injector	
	(Rybelsus) Oral tablet	3, 7, and 14 mg daily in a.m. Take on empty stomach w/H2O sip	
	Tirzepatide (Mounjaro)	2.5, 5.0, 7.5, 10, 12.5 and 15 mg 1x a week pre-filled single dose pen Increase dose by 2.5 mg once monthly to reach targets.	Side effects include: Nausea, diarrhea, injection site reactions. Avoid if family history medullary thyroid tumor. Report pancreatitis. Lowers A1c ~ 1.8 – 2.4% Weight loss of ~ 5.4 – 10 kgs

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GLP-1 Receptor Agonist Devices

Byetta

5mcg or 10mcg pen
1 pen/month
Requires Rx for needles

Ozempic

3 pen options: 0.5, 1, 2mg
1 pen/month
Comes with needles

Victoza

1.2mg, 2 pens/mo
- (15 doses per pen)
1.8mg, 3 pens/month
- (10 doses per pen)
Requires Rx for needles

Adlyxin

Contains 14 doses (20mcg)
2 pens/month
Requires Rx for needles

Bydureon

2mg pen
4 pens/month
Shake 15 seconds
Never see needle

Trulicity

0.75, 1.5, 3, 4.5mg pens
4 pens/month
Never see needle

Oral Semaglutide (Rybelsus)

- ▶ Dose: 3, 7 and 14 mg daily
- ▶ Take daily at least 30 mins before first food, beverage, or other oral meds
- ▶ Take with no more than 4 ounces of plain water
- ▶ Swallow tablets whole (don't cut or crush)
- ▶ Dosing:
 - ▶ Start with 3 mg once daily for 30 days
 - ▶ Then increase to 7mg once daily for 30 days
 - ▶ If A1c at target, maintain at 7mg daily
 - ▶ If A1c not at target, increase to 14 mg once daily



GLP-1 Receptor Agonist Indications

Drug	Lower BG	Reduce CV Risk?	Wt loss approved?
Exenatide IR (Byetta) Lixisenatide (Adlyxin) Semaglutide (Rybelsus)	Yes		
Exenatide ER (Bydureon)	Yes for 10 yrs and older		
Dulaglutide (Trulicity)	Yes	Yes	
Semaglutide (Ozempic)	Yes	Yes	Yes Wegovy 2.4mg
Liraglutide (Victoza)	Yes for 10 yrs and older	Yes	Yes Saxenda 3mg

Package inserts, dailymed.nlm.nih.gov

GLP-1 RA Approved for Weight Loss

- ▶ Saxenda and Victoza same active ingredient (liraglutide) at different doses
 - ▶ Saxenda 3 mg (Victoza 1.8 mg)
 - ▶ 6% wt loss, \$1619 a month
- ▶ Wegovy and Ozempic same active ingredient (semaglutide) at different doses
 - ▶ Wegovy 2.4mg (Ozempic 2mg)
 - ▶ 6% wt loss, \$1619 a month
- ▶ Both are FDA approved as a treatment option for chronic weight management in addition to a reduced calorie diet and physical activity.
- ▶ Approved for use in adults with a
 - ▶ BMI of ≥ 30 or
 - ▶ BMI of ≥ 27 or greater who have hypertension, type 2 diabetes, or dyslipidemia.

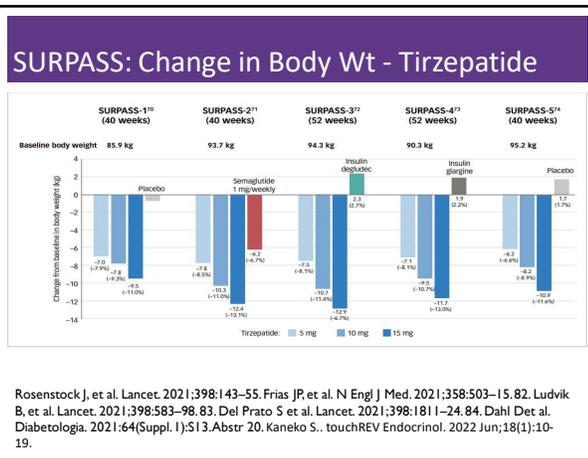
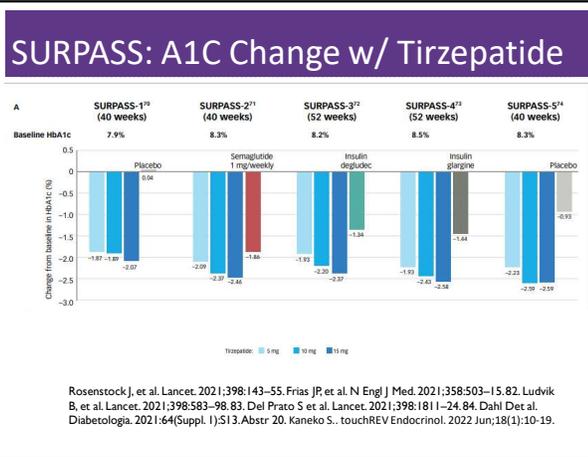


SURPASS Clinical Program

► Evaluated A1C & Weight reductions with Tirzepatide

Study acronym	Study type	Number of participants	Eligibility	Comparator	Study duration (weeks)	Primary outcome
SURPASS-1 ¹⁶	Randomized double-blind	478	Drug-naïve	Placebo	40	HbA1c
SURPASS-2 ¹⁷	Randomized open-label	1,879	Metformin	Semaglutide	40	HbA1c
SURPASS-3 ¹⁷	Randomized open-label	1,947	Metformin w/wo SGLT2	Insulin degludec	52	HbA1c
SURPASS-4 ¹⁷	Randomized open-label	2,002	1-3 antidiabetic medicines (metformin, SGLT2 or sulfonylurea) with cardiovascular risk	Insulin glargine	52	HbA1c
SURPASS-5 ¹⁸	Randomized double-blind	475	Insulin glargine (U100) w/wo metformin	Placebo	40	HbA1c

Rosenstock J, et al. Lancet. 2021;398:143–55. Frias JP, et al. N Engl J Med. 2021;358:503–15.82. Ludvik B, et al. Lancet. 2021;398:583–98.83. Del Prato S et al. Lancet. 2021;398:1811–24.84. Dahl Det al. Diabetologia. 2021;64(Suppl. 1):S13. Abstr 20. Kaneko S., touchREV Endocrinol. 2022 Jun;18(1):10-19.



Counseling Points: GLP-1 RA & GLP-1/GIP

- ▶ Avoid if personal or family history of medullary thyroid cancer
- ▶ Start at lower dose and titrate
- ▶ Eat smaller meals to reduce nausea
- ▶ Avoid high fat meals
- ▶ Rotate sites
- ▶ Store extra pens in fridge
- ▶ Avoid in combo with DPP-4 inhibitors
- ▶ Caution with pancreatitis
- ▶ Ask about recent eye exam
 - ▶ Potential increase in diabetes retinopathy



Case Study Question 3

KZ is on max dose of metformin and is started on tirzepatide to help with BMI of 39 and decrease CV risk. What outcomes can KR expect at highest dose?

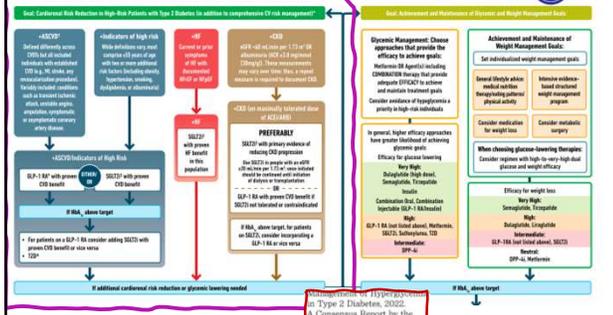
- a. A1C drop of 3-4%, Wt loss of 3-5kg
- b. Increased risk of UTI, A1C drop of 1-2%
- c. Potential for ~10% body wt loss
- d. Doubles risk of pancreatic cancer



Let's Break it Down / ADA & EASD

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HEALTHY LIFESTYLE BEHAVIORS, DIABETES SELF-MANAGEMENT EDUCATION AND SUPPORT (DSMES), SOCIAL DETERMINANTS OF HEALTH (SDOH)



Evaluating Kidney Function - Albumin

Urinary Albumin Creatinine Ratio (UACR)

UACR can be assessed with a urinary spot collection.

Results are viewed by lab short description	
Collection Date & Time	01/13/2022 07:59
ALBUMIN, RANDOM	2.9
ALBUMIN/CREATININ	32
CREATININE, RANDO	91

$2.9 / 91 = 0.0318$ mg/mg or 31.8 (32) in mg/g

- Evaluates ratio of urine albumin /creatinine in mg/g
- Target range < 30mg/g
- If elevated, repeat test to verify
- Check at diagnosis in T2D and within 5 years in T1D

Albuminuria Categories	Urinary Albumin Creatinine Ratio (UACR)
Normal to mildly increased – A1	< 30 mg/g
Moderately increased – A2	30 – 299 mg/g
Severely increased – A3	300 mg/g +

Evaluating Kidney Function - GFR

Glomerular Filtration Rate (GFR)– target is 60 or greater

Stage 3 indicates progressive renal failure

- GFR 30 to 59
- Stage 4 and 5 indicates severe loss and failure
- GFR 29 or less

Kidney Disease Stage	GFR
Stage 1 – Normal	90+
Stage 2 – Mild loss	89 - 60
Stage 3a – Mild to Mod	59 - 45
Stage 3b – Mod to Severe	44 - 30
Stage 4 – Severe loss	29 - 15
Stage 5 – Kidney failure	14 - 0

www.DiabetesEd.net



From: 11. Chronic Kidney Disease and Risk Management: Standards of Medical Care in Diabetes—2022

Diabetes Care. 2021;45(Supplement 1):S175-S184. doi:10.2337/dc22-S011

CKD is classified based on:			Albuminuria categories			
			A1	A2	A3	
			Normal to mildly increased	Moderately increased	Severely increased	
			<30 mg/g <3 mg/mmol	30-299 mg/g 3-29 mg/mmol	≥300 mg/g ≥30 mg/mmol	
GFR categories (estimated, 75 mL/min/1.73 m ²) Description and range	G1	Normal to high	≥90	1 if CKD	Treat 1	Refer* 2
	G2	Mildly decreased	60-89	1 if CKD	Treat 1	Refer* 2
	G3a	Mildly to moderately decreased	45-59	Treat 1	Treat 2	Refer 3
	G3b	Moderately to severely decreased	30-44	Treat 2	Treat 3	Refer 3
	G4	Severely decreased	15-29	Refer* 3	Refer* 3	Refer 4+
G5	Kidney failure	<15	Refer 4+	Refer 4+	Refer 4+	

Date of Download: 10/11/2022

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SGLT-2 Inhibitors Comparison

Common Oral Diabetes Meds

Class/Main Action	Name(s)	Daily Dose Range	Considerations
SGLT2 Inhibitors "Glucoretic" • Decreases glucose reabsorption in kidneys	Canagliflozin* (Invokana)	100 - 300 mg 1x daily	Side effects: hypotension, UTIs, genital infections, increased urination, weight loss, ketoacidosis. Heart Failure, CV & Kidney Protection: 1st line therapy for Heart Failure (HF), Kidney Disease (CKD), Cardiovascular Disease, before or with metformin. Considerations: See Package Insert (PI) for GFR cut-offs, dosing. Limited BG lowering effect if GFR < 45, still benefits kidneys & heart at lower GFR. For renal protection, use SGLT-2 therapy if eGFR ≥ 25 & UACR ≥ 300 (ADA). Benefits: SGLT-2s* reduce BG, CV death & HF, slow CKD. Lowers A1c 0.6% -1.5%.
	Dapagliflozin* (Farxiga)	5 - 10 mg 1x daily	
	Empagliflozin* (Jardiance)	10 - 25 mg 1x daily	
	Ertugliflozin (Steglatro)	5 - 15 mg 1x daily	

Benefits of SGLT-2 Inhibitors

A1C lowering

Weight loss

Cardiovascular benefits

Renal benefits

Heart failure benefits

Blood pressure lowering

Side Effects of SGLT-2 Inhibitors

Genitourinary infections

Volume depletion

Increased urination

Hypotension

Hyperkalemia

Diabetes ketoacidosis (DKA)

SGLT-2i Indications

Drug	Lower BG	Reduce CV Risk?	Use to treat Heart Failure?	Slow renal disease?
Ertugliflozin (Steglatro)	Yes			Yes
Canagliflozin (Invokana)	Yes	Yes	Yes w/ Diabetes	Yes
Dapagliflozin (Farxiga)	Yes	Yes	Yes +/- Diabetes	Yes
Empagliflozin (Jardiance)	Yes	Yes	Yes +/- Diabetes	Yes

Package inserts, dailymed.nlm.nih.gov

SGLT2i: Managing Adverse Effects

- ▶ Maintain good hygiene to reduce risk of genital mycotic infections
 - ▶ Higher risk with higher glucose
- ▶ DKA risk
 - ▶ Use caution with reducing insulin dose
- ▶ Monitor BP
 - ▶ May need to reduce antihypertensive meds
- ▶ UTI risk greater with hyperglycemia
- ▶ Amputations observed with canagliflozin
 - ▶ Good foot care, check feet daily
- ▶ Monitor renal function/potassium



Case Study LS

LS is 69 with type 2 diabetes for over a decade. Takes metformin 1000mg twice daily and dulaglutide 3mg weekly.

A1C=7.3%.

UACR 212mg/gm

eGFR=56

B/P 146/82

Weight: 205lbs, 5'7,"

BMI=32kg/m²

Lost 10lbs in the last year



LS continued Poll 4

► What is the best drug to add to LS regimen?

- A. Glipizide
- B. Dapagliflozin (Farxiga)
- C. Pioglitazone (Actos)
- D. Linagliptin (Tradjenta)
- E. More than 1 correct answer



What teaching would you include? Could we increase the dulaglutide dose?

How would we manage other risk factors:

- Use of ACE-inhibitor/ARB, BP management
- Statin for CV risk

What the Rx?



Now on to other diabetes medications we haven't yet covered

How Many Drug Options for Diabetes?

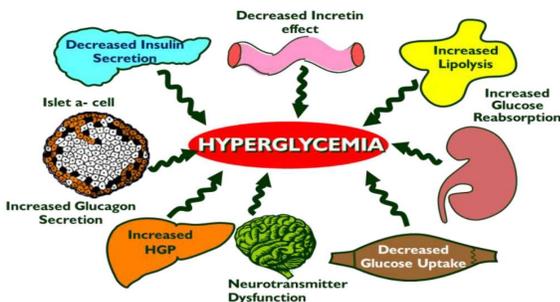
- ▶ Biguanide
- ▶ Sulfonylureas
- ▶ Meglitinides
- ▶ Thiazolidinediones (TZD's)
- ▶ Dipeptidylpeptidase-4 (DPP-4) inhibitors
- ▶ Glucagon-like-peptide-1 (GLP-1) receptor agonists
- ▶ GLP/GIP receptor agonists
- ▶ Sodium glucose cotransporter-2 (SGLT-2) inhibitors
- ▶ Bile acid sequestrant
- ▶ Dopamine-2-agonist
- ▶ Amylin mimetic
- ▶ Alpha-glucosidase inhibitors
- ▶ Insulin
- ▶ Glucagon

Other Oral Diabetes Medications

Class/Main Action	Name(s)	Daily Dose Range	Considerations
Thiazolidinediones "TZDs" • Increases insulin sensitivity	pioglitazone (Actos) rosiglitazone (Avandia)	15 – 45 mg daily 4 – 8 mg daily	Black Box Warning: TZDs may cause or worsen CHF. Monitor for edema and weight gain. Increased peripheral fracture risk. Actos may increase risk of bladder cancer. Lowers A1c 0.5% – 1.0%
Glucosidase Inhibitors • Delays carb absorption	acarbose (Precose) miglitol (Glyset)	25 – 100 mg w/meals; 300 mg max daily dose	Start low dose, increase at 4-8 wk intervals to decrease GI effects. Caution with liver or kidney problems. In case of hypo, treat w/ glucose tabs. Lowers A1c 0.5 – 1.0%
Meglitinides • Stimulates rapid insulin burst	repaglinide (Prandin) nateglinide (Starlix)	0.5 – 4 mg w/meals (metabolized in liver) 60 – 120 mg w/meals (eliminated via kidney)	Take before meals. Side effects may include hypoglycemia and weight gain. Lowers A1c 1.0% – 2.0%
Dopamine Receptor Agonists • Resets circadian rhythm	bromocriptine mesylate—Quick Release "QR" (Cycloset)	1.6 to 4.8 mg a day (each tab 0.8 mg)	Take within 2 hrs of waking. Side effects: nausea, headache, fatigue, hypotension, syncope, somnolence. Lowers A1c 0.6% – 0.9%
Bile Acid Sequestrants • Decreases cholesterol / BG levels.	colesevelam HCL (Welchol)	Up to six (6) 625 mg pills (3 tabs am, 3 tabs pm) 3.75gm packet in 4-8 ounces of fluid	Do not use if history of bowel obstruction, triglycerides >500, or pancreatitis. Can decrease absorption of certain meds, soluble vitamins. Lowers LDL by 15-30%. Side effects GI in nature. Lowers A1c 0.5%

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Drug Targets in Diabetes



DeFronzo et al. Diabetes Spectrum Volume 27, Number 2, 2014

Poll Question 6

- ▶ A potential side effect of SGLT-2 Inhibitors is:
 - a. ketoacidosis
 - b. Hypertension
 - c. Kidney tenderness
 - d. Increased uric acid



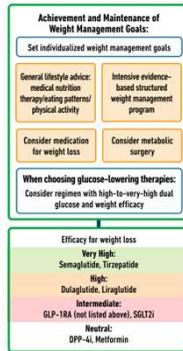
PR wants to lose more weight

- ▶ After 3 months, PR's A1C has decreased to 7.1%. After meeting with diabetes care and education specialist, has increased physical activity to 3 days/week of walking. Lost 6lbs and would like to lose more.
- ▶ Current DM Meds: empagliflozin(Jardiance) 25mg daily.
- ▶ BMI=34kg/m²
- ▶ What drug would be best to start next?



When Goal is to Avoid Weight Gain

- ▶ These meds associated with wt loss
 - ▶ GLP-1 and GIP agonists
 - ▶ SGLT-2 Inhibitors
- ▶ These meds are weight neutral
 - ▶ Metformin
 - ▶ DPP-4 Inhibitor



ADA /EASD
Management of Hyperglycemia in Type 2 Diabetes, 2022
A Consensus Report by the

7. Best medication to lose weight?

▶ Lost 6lbs and would like to lose more.

Current DM Meds: empagliflozin (Jardiance) 25mg

daily. BMI=34.

▶ What drug would be best to start next?



- A. Tirzepatide
- B. Metformin
- C. Semaglutide
- D. Degludec
- E. A or C

Of course, lifestyle coaching and RD referral included in plan!

Poll 8 - Check Your Knowledge

Which of the following medications is **least** affordable?

- A. Pioglitazone (Actos)
- B. Metformin (Glucophage)
- C. Glimepiride (Amaryl)
- D. Sitagliptin (Januvia)



How much do they cost?

▶ Which of the following groups of meds for a month supply are cheapest? (multip^l

- a. Actos and Avandia \$5 & \$324
- b. Glipizide, Glyburide, Glimepiride \$10 for 3 mo's
- c. Metformin and Metformin XR \$10 for 3 mo's
- d. Januvia and Onglyza \$596 & \$549
- e. Exenatide and Semaglutide \$909, \$1022
- f. Empagliflozin and SGLT-2s \$658
- g. Tirzepatide (Mounjaro) \$974

See Table 9.3 in ADA Standards on Median Monthly Average Wholesale Price (AWP) 2022



Hypoglycemia (Glucose) Alert Values

▶ BG <70mg/dl – Level 1

- ▶ Follow 15/15 rule and contact provider to make needed changes



▶ BG < 54mg/dl – Level 2

- ▶ Indicates serious hypo. Contact provider for med change. Glucagon Emergency Kit

▶ Severe Hypoglycemia – Level 3

- ▶ Requires external assistance – no threshold

Hypoglycemia: Identify, Treat, & Prevent



Step 1

Identify your signs of hypoglycemia or low blood sugar:

- Sweaty
- Shaky
- Hungry
- Can't think straight
- Headache
- Irritated, grouchy
- Other

Step 2

If have signs of hypo, treat with carbs until glucose reaches 70+, then eat usual meal.

- Sugary drink, 4–8oz
- Piece of fruit
- Raisins, handful
- Glucose tabs, 4+
- Honey or glucose gel
- Skittles candy, 15+

Step 3

Have glucagon rescue meds available.

In case of severe hypo, identify someone (ahead of time) who can get medical help & give a glucagon rescue medication.

Notify your provider of low blood sugar events.

Hypoglycemia Levels:

- Level 1 – Glucose less than 70
- Level 2 – Glucose less than 54
- Level 3 - Severe, needs assistance

Identify Causes of Hypo & Problem Solve to Prevent Future Episodes

- » Low carb meal
- » Extra activity
- » Drinking alcohol
- » Delayed, missed meal
- » Too much insulin/meds
- » Insulin timing

www.DiabetesEd.net

For educational purposes DiabetesEd Services

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Glucagon-Rescue Medications for Diabetes-Related Hypoglycemia



Name/Delivery	Supplied	Dose Range		Age / Route / Storage
		Adult	Peds / Age WT Dosing	
Glucagon Emergency Kit Injection requires mixing glucagon powder	1mg / 1mL vial + syringe	1mg	0.03mg/kg or < 6yrs or < 25 kgs 0.5mg ≥ 6yrs or > 25kgs 1mg	All ages approved SubQ or IM admin Expires in 2 years at room temp.
Baqsimi Nasal glucagon powder	3 mg intranasal device	3 mg	< 4 yrs: not recommended 4 yrs or older 3mg dose	Approved Age 4+ Nasal admin Expires ~ 2 years at room temp (keep in shrink-wrapped tube).
Gvoke Injectable liquid stable glucagon solution	0.5mg/1.0mg prefilled syringe or 0.5mg/1.0mg HypoPen auto-injector	1 mg	< 2yrs: not recommended 2- 12 yrs < 45kg 0.5mg ≥ 45kg 1mg 12 yrs or older 1mg	Approved Age 2+ SubQ admin in arm, thigh, abdomen Expires in 2 years at room temp (keep in foil pouch).
Dasiglucagon (Zegalogue) Stable liquid glucagon analog	0.6mg/0.6mL Prefilled syringe Autoinjector	0.6mg	< 6yrs: not recommended 6yrs or older 0.6mg	Approved Age 6+ SubQ in abdomen, buttocks, thigh outer upper arm Expires in 1 year at room temp. (store in red protective case).

*All raise BG 20+ points. Can cause nausea, vomiting. After admin, roll person on side. Seek medical help. If no response after 1st dose, give 2nd dose in 15 mins. When awake, give oral carbs ASAP when safe to swallow. Please consult package insert for detailed info.
All PocketCard content is for educational purposes only. Please consult prescribing information for detailed guidelines.
DiabetesEd.net Copyright Diabetes Education Services 2021 ©2021

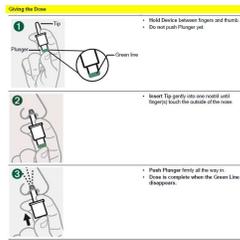
Glucagon Emergency Kit



Store 68-77 degrees prior to reconstitution, single use only

Nasal Glucagon - Baqsimi

- ▶ Approved for ages 4 +
- ▶ Absorbed nasally
- ▶ No reconstitution or refrigeration needed
- ▶ Kept in temps up to 86
- ▶ Raises BG 67-73 mg/dl
- ▶ Don't use in those with
 - ▶ Pheochromocytoma
 - ▶ Insulinoma
 - ▶ See package insert



Gvoke HypoPen – Single dose injector For children ages 2+. Peds dose up to 45kg

Gvoke HypoPen™ (glucagon injection) 1 mg per 0.2 mL NDC 72066-121-11 BB Only
Contains 1 Single-Dose Auto-Injector

FOR LOW BLOOD SUGAR EMERGENCY

- 1. Prepare**: Tear Open Pouch at Dotted Line. Remove Auto-Injector. Pull off Red Cap.
- 2. Inject**: Choose Injection Site and Expose Skin. Push Down on Skin to Start. Hold Down for 5 Seconds. Wait for Window to Turn Red. Hold Down for 5 Sec.
- 3. Assist**: Turn Patient on Side. Call Emergency Medical Help.

After the Injection, Put the Used Pen in a Safe Place Until It Can be Disposed of into a FDA Cleared Sharps Container.

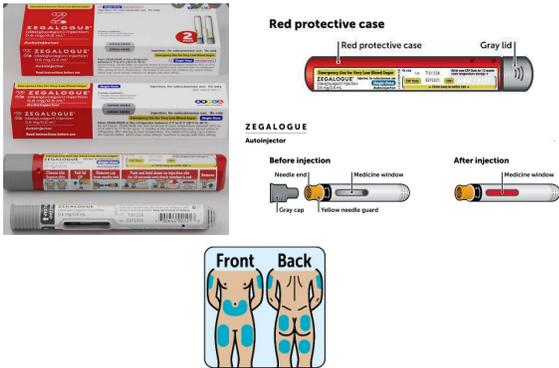
Gvoke HypoPen™ (glucagon injection) 0.5 mg per 0.1 mL NDC 72066-120-11 BB Only
Contains 1 Single-Dose Auto-Injector

FOR LOW BLOOD SUGAR EMERGENCY

- 1. Prepare**: Tear Open Pouch at Dotted Line. Remove Auto-Injector. Pull off Red Cap.
- 2. Inject**: Choose Injection Site and Expose Skin. Push Down on Skin to Start. Hold Down for 5 Seconds. Wait for Window to Turn Red. Hold Down for 5 Sec.
- 3. Assist**: Turn Patient on Side. Call Emergency Medical Help.

After the Injection, Put the Used Pen in a Safe Place Until It Can be Disposed of into a FDA Cleared Sharps Container.

Dasiglucagon (Zegalogue)



EV Arrives and Requests Help

▶ 58 yr old complains of 4 lb wt gain for past month. BMI 31, wt 90 kg. B/P 142/96. A1C 8.3%

What story do these meds tell?
Any med(s) missing?
Any med needs to be stopped?

▶ Meds include:

- ▶ Sitagliptin, Metformin
- ▶ Actos 15mg ac breakfast
- ▶ Basaglar 58 units
- ▶ Semaglutide 0.5mg weekly
- ▶ Levothyroxine (ran out)
- ▶ Lisinopril 10mg
- ▶ Gabapentin 100 mg TID



EV Arrives and Requests Help

▶ 58 yr old complains of 4 lb wt gain for past month. BMI 31, wt 90 kg. B/P 142/96. Checks BG in morning; 150ish. A1C 8.3%

What does this tell us about EV?

▶ Meds include:

- ▶ Sitagliptin (DPP-IV), Metformin
- ▶ Basaglar 58 units (Basal)
- ▶ Semaglutide 0.5mg wk (GLP-1)
- ▶ Levothyroxine (ran out)
- ▶ Lisinopril 10mg (ACE)
- ▶ Lovastatin 20mg (Statin)
- ▶ Gabapentin 100 mg TID (leg pain)

- Struggling with weight
- B/P & A1C above target
- Overbasalized (max dose 0.5 units/kg a day)
- Why not taking thyroid med?
- Lower extremity pain contributing to distress?
- Elevated CV risk?

EV is Gaining Weight and is Tired

- ▶ 58 yr old complains of 4 lb wt gain for past month. BMI 31, wt 90 kg. B/P 142/96. Checks BG in morning; 150ish. A1C 8.3%
- ▶ Meds include:
 - ▶ Sitagliptin, Metformin
 - ▶ Actos 15mg ac breakfast
 - ▶ Basaglar 58 units
 - ▶ Semaglutide 0.5mg weekly
 - ▶ Levothyroxine – ran out
 - ▶ Lisinopril 10mg
 - ▶ Gabapentin 100 mg TID



Labs

A1C – 8.3%
 UACR 26 GFR >60
 TSH 10.6
 LDL 98 mg/dl, Trig 158
 ALT 85 IU/L, AST 90 IU/L
 (normal range 25-50)

Life situation

Takes care of dad with dementia
 Gums inflamed
 No eye doctor for year
 Both feet hurt at night

ABCs of Diabetes

- ▶ A1c less than 7%
 - ▶ Pre-meal BG 80-130
 - ▶ Post meal BG <180
- ▶ Blood Pressure < 140/90
 - ▶ BP target <130/80
 - ▶ If CVD or 10-year CVD Risk > 15%
- ▶ Cholesterol
 - ▶ Statin therapy indicated if 40+



Advocating for Best Health for people with Diabetes

- ▶ Modifiable
 - ▶ Sleep
 - ▶ Activity
 - ▶ Smoking
 - ▶ Dietary Habits
 - ▶ Glucose
 - ▶ Blood Pressure
 - ▶ Lipids
 - ▶ Oral Care
 - ▶ Immunizations
 - ▶ Psychosocial care



▶ Make small, achievable goals. We are in this for the long run.

Diabetes is a long path



Get at least 7 hours of sleep a night – Check for sleep apnea

Obstructive Sleep Apnea - OSA

- ▶ OSA affects ~25% of people with type 2
- ▶ Up to 60% of those with type 2 have disordered sleep
- ▶ Associated with increased CVD risk
- ▶ 4-10 increased risk if BMI 30+ with visceral adiposity
- ▶ Treatment:
 - ▶ Lifestyle modification
 - ▶ Continuous positive airway pressure and devices
 - ▶ Surgery



Where are we on this continuum?



Only about 50% of us are meeting activity goals

Good Exercise Info / Quotes

▶ “Passagiata” – take an after meal stroll

- ▶ Exercise decreases A1c 0.7%
- ▶ No change in body wt, but 48% loss in visceral fat



“Every minute of activity lowers blood sugar one point.”

“I don’t have time to exercise, I MAKE time.”
Mike Huckabee

Best Shake For People with Diabetes



“The only diet shake I recommend is the shake your booty makes when you exercise.”

From Debbie Nagata's slide collection

Smoking and Diabetes

Smoking increases risk of diabetes 30%



- Ask at every visit
- Assess
- Advise
- Assist with stop smoking
- Arrange for referrals
- Organize your clinic

USDA www.myplate.gov

Balancing Calories

- ▶ Enjoy your food, but eat less.
- ▶ Avoid oversized portions.

Foods to Increase

- ▶ Make half your plate fruits and vegetables.
- ▶ Make at least half your grains whole grains.
- ▶ Switch to fat-free or low-fat (1%) milk.

Foods to Reduce

- ▶ Compare sodium in foods like soup, bread, and frozen meals — and choose the foods with lower numbers.
- Drink water instead of sugary drinks.



Plan Your Portions



Use a smaller plate. This is a 9-inch plate to help guide you.

Diabetes Toolkit

Meter

- Strips that aren't expired?

List of Meds

Plan for Lows

Emergency Plan

Power back-up

- ▶ BG Checks and logging results
- ▶ Diabetes ID
- ▶ Phone, medic alert, on person
- ▶ Carbohydrate source
- ▶ Granola bar, glucose tabs, GU, gummy bears
- ▶ Rescue Meds

Collaborative Action Plan

- ▶ Increase semaglutide to 1.0mg
- ▶ Decrease basaglar by 10 units
- ▶ Stop sitagliptin, pioglitazone (Actos)
- ▶ Walk after lunch during work week
- ▶ Restart levothyroxine, Re-Check TSH
- Re-evaluate in 4 weeks.
- ▶ Eat one serving of veggie a day and decrease meat intake to 4 nights a week.
- ▶ Meet with RD/RDN
- ▶ Check BG a few times a week before bed (in addition to am)



What about alcohol intake?
Are these goals realistic?

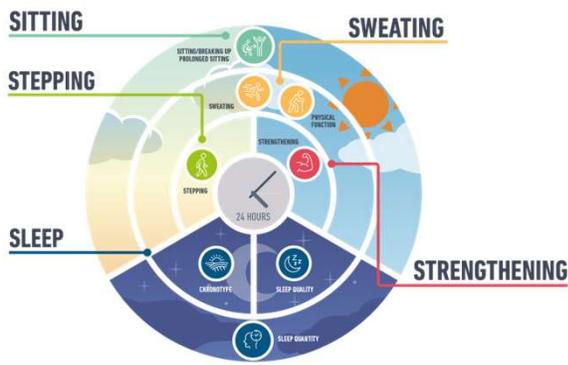
Exercise Standards

- ▶ Adults – 150 min/wk moderate intensity
▶ over 3 days a week.
- ▶ Don't miss > 2 consecutive days w/out exercise
- ▶ Get up every 30 mins - Reduce sedentary time
- ▶ Flexibility and balance training 2-3 xs a week (Yoga and Tai Chi)
- ▶ T1 and T2 – resistance training 2 -3 xs a week



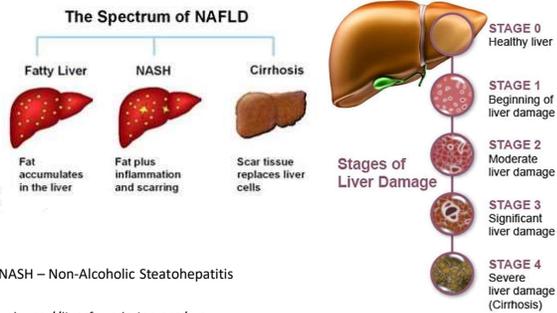
ADA Standards 2022

FIGURE 2: IMPORTANCE OF 24-HOUR PHYSICAL BEHAVIOURS FOR TYPE 2 DIABETES



Management of Hypoglycemia in Type 2 Diabetes, 2022. A Consensus Document by the

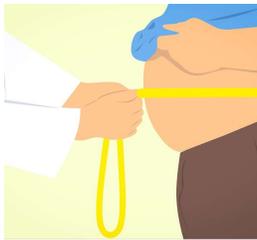
Natural History of NAFLD to NASH



NASH – Non-Alcoholic Steatohepatitis

<https://liverfoundation.org/wp-content/uploads/2020/11/StagesFibrosis.jpg>

Fatty liver disease and diabetes



Associated with :

- Increased BMI (30+)
 - Larger waist circumference,
 - Elevated triglycerides
 - Lower HDL cholesterol levels.
- ADA 2022

First indicators may include elevated alanine transaminase (ALT) and aspartate transaminase (AST).

The growing epidemic of NAFLD in western societies:

- 20 to 30% of overall population - 45 to 75% of ind's with type 2 diabetes

Review article | Open Access | Published: 05 June 2020
Nonalcoholic fatty liver disease and type 2 diabetes: where do Diabetologists stand?
Shahana Samad | Nisha Akhthar | N. Gaurav Hande
Clinical Diabetes and Endocrinology • Article number 9 (2020) | <https://clindiaabetesendo.biomedcentral.com/articles/10.1186/s40842-020-00097-1>

Symptoms of Fatty Liver

If symptoms do appear, they may include:

- ▶ A feeling of fullness in the middle or upper right side of the abdomen
- ▶ Abdominal pain, nausea
- ▶ Loss of appetite or weight loss
- ▶ Weakness
- ▶ Jaundice
- ▶ Swelling of the abdomen and legs
- ▶ Mental confusion
- ▶ Extreme fatigue or tiredness
- ▶ Signs of advanced disease include:
 - ▶ Portal hypertension, spider angiomas, reddening of palms, declining platelet counts

Mayo Clinic

Finding Liver Disease

- ▶ Imaging procedures used to diagnose NAFLD include:
 - ▶ **Abdominal ultrasound**, which is often the initial test when liver disease is suspected.
 - ▶ **Transient elastography**, an enhanced form of ultrasound that measures the stiffness of liver. Liver stiffness indicates fibrosis or scarring.
 - ▶ **Magnetic resonance elastography**, works by combining MRI imaging with sound waves to create a visual map (elastogram) showing the stiffness of body tissues
 - ▶ **Biopsy** by liver specialist confirms definitive diagnosis



Mayo Clinic

Treatment for NAFLD and NASH

Interventions that improve metabolic abnormalities include: weight loss, glycemic improvement and meds that treat hyperglycemia, dyslipidemia

Table 4.6—Management of patients with nonalcoholic fatty liver disease and nonalcoholic steatohepatitis

Variable	Lifestyle intervention ^a	Liver-directed pharmacotherapy	Diabetes care (in individuals with diabetes)	Cardiovascular risk reduction
NAFLD	Yes	No	Standard of care	Yes
NASH with fibrosis stage 0 or 1 (F0, F1)	Yes	No	Standard of care	Yes
NASH with fibrosis stage 2 or 3 (F2, F3)	Yes	Yes	Pioglitazone, GLP-1 receptor agonists ^b	Yes
NASH cirrhosis (F4)	Yes	Yes	Individualize ^c	Yes

NAFLD, nonalcoholic fatty liver disease; NASH, nonalcoholic steatohepatitis. ^aAll patients require regular physical activity and healthy diet and to avoid excess alcohol intake; weight loss recommended. ^bAmong glucagon-like peptide 1 (GLP-1) receptor agonists, semaglutide has the best evidence of benefit in patients with NASH and fibrosis. ^cEvidence for efficacy of pharmacotherapy in patients with NASH cirrhosis is very limited and should be individualized and used with caution. Adapted from "Preparing for the NASH Epidemic: A Call to Action" (92).

For biopsy proven NAFLD – these treatments improve liver histology but need long term studies (ADA 2022):
 Pioglitazone (Actos) Vitamin E GLP-1 Receptor Agonists

Collaborative Action Plan and F/U

- ▶ Make appointment with dentist and eye doctor.
- ▶ Brush twice daily and floss daily.
- ▶ Need some relief from nerve pain.
- ▶ Feet calluses



Meds for Neuropathy – Cheat Sheet

Neuropathy Medication for Diabetes

Prevention – Maintain glycemic control; quit smoking, alcohol reduction, exercise.

Pathogenetically Oriented Therapy

- Alpha lipoic acid 600 – 1,800 mg a day

Prescription Therapy:

1st line – Tricyclic Antidepressants (Amitriptyline, Nortriptyline, Desipramine)

- Calcium Channel Modulators (Gabapentin, Pregabalin)
 - Serotonin Norepinephrine Reuptake Inhibitors (SNRI – Venlafaxine, Duloxetine)
- 2nd Line** - Topical Capsaicin Cream for localized pain – Apply 2-4 x daily for up to 8 wks
- Opioids (Tramadol, Oxycodone)

Reasons for Treatment Failure

- Dose too low
- Inadequate trial – requires 2-8 weeks of treatment to observe symptom reduction
- Pt expecting elimination of symptoms – only reduces symptoms by about 50%
- Incorrect diagnosis: If in doubt, refer to neurologist
- If patient does not respond or has adverse effects, change medication class
- In patient has some but inadequate relief, raise the dose and consider adding or changing meds.

References: Ziegler, D. Painful diabetic neuropathy. Diabetes Care 2009; 32 (Supp 2): S414-S419

Meds for Neuropathy – Cheat Sheet

Class	Generic / Trade Name	Usual Daily Dose Range	Comments	Side Effects/ Caution
1st Line Agents Tricyclic Antidepressants TCA Improves neuropathy and depression	Amitriptyline / Elavil	25 – 100 mg* Avg dose 75mg	Usually 1 st choice	Take 1 hour before sleep. Side effects: dry mouth, tiredness, orthostatic hypotension.
	Nortriptyline / Pamelor	25 – 150 mg* (for burning mouth)	Less sedating and anticholinergic	
	Desipramine / Norpramine	25 – 150 mg* *Increase by 25mg weekly till pain relieved		Caution: not for pts w/ unstable angina (<6 mo), MI, heart failure, conduction system disorder.
Calcium Channel Modulators	Gabapentin/ Neurontin	100 - 1,200mg TID	Improves neuropathy treatment	Sedation, dizziness, peripheral edema, wt gain
	Pregabalin / Lyrica	50 - 200mg TID	*FDA approved for neuropathy treatment fewer drug interactions	Caution: CHF, suicide risk, seizure disorder.
Serotonin Norepinephrine Reuptake Inhibitor SNRI	Duloxetine / Cymbalta	60 mg daily Start at 30 mg	Improves neuropathy treatment	Nausea, sedation, HTN, constipation, dizziness, dry mouth, blurred vision.
	Venlafaxine/ Effexor	75 - 225 mg daily	Improves depression, insomnia	Caution: adjust dose for renal insufficiency, do not stop abruptly, taper dose.
2nd Line Agents Opioids	Weak opioids Tramadol / Ultram	50 – 400 mg	Sedation, nausea, constipation (always prescribe stool softener) Caution: abuse, suicide risk, short acting opioids not recommended for long term tx, can develop tolerance	
	Strong opioids Oxycodone	10 – 100 mg		
Local Treatment	Capsaicin Cream (0.025%)	Apply 2-4 x daily for up to 8 wks		
Other choices	If above medications not effective, contraindicated or intolerable consider: Bupropion/Welbutrin Paroxetine / Paxil Clonazepam / Celexa Topiramate / Topamax Topical Lidocaine (for localized pain).			



Other strategies to help ease the pain

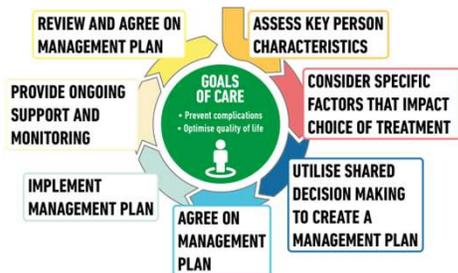
- ▶ Music
 - ▶ Podcasts
 - ▶ Movies
 - ▶ Pet's
 - ▶ Massage
 - ▶ Touch
 - ▶ Topical creams
 - ▶ Lidocaine patches
 - ▶ Mineral salts baths
 - ▶ Neurostimulators
-
- ▶ Tylenol / Ibuprofen
 - ▶ Earthing
 - ▶ Sleep
 - ▶ Hobbies
 - ▶ Aromatherapy
 - ▶ Time with special people
 - ▶ Work / volunteering

EV is feeling Empowered

- ▶ Her A1c has dropped, she feels better about herself with healthier eating and increased activity.
- ▶ She is back on her thyroid medication and has more energy.
- ▶ Appointment with Podiatrist
- ▶ The pain in her feet is better and she is more hopeful overall!



FIGURE 1: DECISION CYCLE FOR PERSON-CENTRED GLYCAEMIC MANAGEMENT IN TYPE 2 DIABETES



Diwan M, Anand V, Collins M, Gabbay RA, Green L, Mochtar MM, Roscoe M, Sirtori CR, Del Prato S, Madsen C, Mergener G, Ronghi P, Terzian T, Tsapas A, Vague P. *Diabetes Care* 2022; <https://doi.org/10.2337/2022-0334>. Under review 2022. <https://doi.org/10.2337/2022-0334>.

No DE-FEET



Preventing Agony of “DeFeet”

1. Describe risk factors for lower extremity complications.
2. Discuss prevention strategies.
3. Demonstrate steps involved in lower extremity assessment.

STANDARDS OF CARE | DECEMBER 16, 2021

12. Retinopathy, Neuropathy, and Foot Care: Standards of Medical Care in Diabetes—2022

American Diabetes Association Professional Practice Committee

Check for updates

Diabetes Care 2022;45(Supplement_1):S185–S194
https://doi.org/10.2337/acc2-2022



Notes from Beverly



All HCP can help save feet!



Some of these images may be difficult to view.

Lower Extremity Complications

- ▶ Combination of vascular, neurological, and musculoskeletal dysfunction
- ▶ Foot ulcers and amputations (from neuropathy and peripheral arterial disease) are associated with higher morbidity and mortality rates.
- ▶ Early recognition and treatment makes a difference



Diabetes and Amputations

- ▶ Rate declined 43% - 2000 – 2009
- ▶ **Increased 50% from 2009-2015**
 - ▶ 2.1 per 1000 then up to 4.2 per 1000
 - ▶ Driven by a 62% increase in minor amputations (toes)
 - ▶ Highest rates in young and middle age adults (18- 64 years).
- ▶ 130,000 adults annually with diabetes have lower extremity amputations [NIDDK / NIH](#)
- ▶ This number equates to **five out of every 1,000** people with diabetes.

Diabetes Care 2018
Resurgence of Diabetes-Related Nontraumatic Lower Extremity Amputation in the Young and Middle-Aged Adult U.S. Population
[https://doi.org/10.2337/DC180128](#)



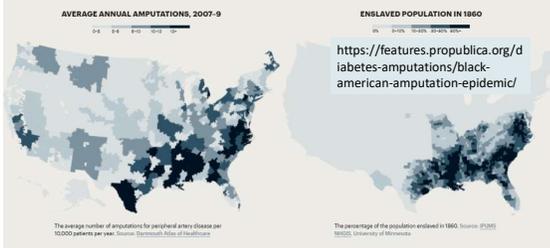
Health Disparities and Lower Extremity Amputations

- ▶ African Americans and people of color have 3-4 times the rate of amputation, compared to White Americans
- ▶ 60% of amputations in 7% of population
- ▶ Amputations cost \$30,000 – 60,000
- ▶ Associated w/ earlier death compared to revascularization



The Black American Amputation Epidemic

Despite the great scientific strides in diabetes care, the rate of amputations across the country grew by 50% between 2009 and 2015. Diabetics undergo 130,000 amputations each year, often in low-income and underinsured neighborhoods. Black patients lose limbs at a rate triple that of others. It is the cardinal sin of the American health system in a single surgery: save on preventive care, pay big on the backend, and let the chronically sick and underprivileged feel the extreme consequences.



Foot Ulcer usually doesn't lead to amputation – but it can.

- ▶ Foot ulcers occur in 4–10% of people with diabetes.
- ▶ Outcomes include:
 - 60–80 percent of foot ulcers will heal
 - 10–15 percent will remain active
 - 5–24 percent will eventually lead to limb amputation within 6–18 months of the initial evaluation

Diabetes Ther. 2012 Dec; 3(1): 4
Published online 2012 Apr 20. doi: 10.1007/s13300-012-0004-9

<https://www.ncbi.nlm.nih.gov/pmc/articles/PMC3508111/>

Management of Diabetic Foot Ulcers
Klaseytra Alexiadou¹ and John Dargatzis²

Poll Question 1

▶ Which of the following factor(s) increase risk for amputation in diabetes?

- A. Socioeconomic status
- B. Cigarette smoking
- C. Previous amputation
- D. Age and ethnicity
- E. All of the above



Racial Disparities and Amputations

NIH National Institute of Diabetes and Digestive and Kidney Diseases

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Reducing Disparities in Diabetic Amputations

April 21, 2021 0 Comments

Recent compilation of Diabetes Social Determinants of Health

Risk for amputation? Consider these factors:

- ▶ **region.** People who live in the southern United States have the highest rates of amputation. They also have the lowest rates of revascularization.
- ▶ **race.** Most people receiving amputations are minorities: Black Americans, Hispanics/Latinos, and American Indians.
- ▶ **age.** Many people who receive amputations are older. PAD may be missed in older adults because the symptoms are attributed to arthritis or gout. Also, primary care doctors may not know about PAD and may not screen patients for PAD early. Patients undergo an amputation when they are older because PAD was missed.
- ▶ **socioeconomic status.** Poorer patients and those living in poorer regions of the country have less access to quality health care and have the highest amputation rates. Unfortunately, many of these patients are minorities with low incomes.
- ▶ **hospital volume of vascular procedures.** Hospitals are better at preventing amputation if they can assemble a team of specialists proficient in aggressive limb salvage, wound care, nutritional care, and diabetes management and treatment. Rural areas, such as those in the southern United States, don't have a significant number of these specialists.

Learn about how diagnosing and treating peripheral arterial disease in people with diabetes can help prevent amputations.

Felicia A. Fakhoury, MD, a cardiologist in Bolivar County, MS, has used prevention, screening, and treatment strategies to reduce amputations by 80% in the Mississippi Delta and reduce the number of amputations in patients with diabetes. Dr. Fakhoury discusses risk factors for peripheral arterial disease (PAD) and amputation in patients with diabetes, and how to reduce disparities in diabetic amputations.

https://www.niddk.nih.gov/health-information/professionals/diabetes-discoveries-practice/reducing-disparities-in-diabetic-amputations?utm_source=diabetes%20discoveries%20%26%20practice%20blog&utm_medium=se-mail

Poll Question 2

▶ Which of the following is true about diabetes and lower extremities?



- A. Over 30% of people with diabetes experience amputation.
- B. Over 50% of amputations could have been avoided.
- C. Most amputations happen before the age of 70
- D. The rate of amputations continues to decrease.

Foot Care Standards - ADA

- ▶ Perform a comprehensive foot evaluation at least **annually** to identify risk factors for ulcers and amputations.
- ▶ Provide general preventive foot self-care education to **all people** living with diabetes.
- ▶ Sensory loss or prior ulceration or amputation?
 - ▶ inspect feet at **every visit**.
- ▶ **High-risk** may need specialized therapeutic footwear:
 - ▶ If severe neuropathy, foot deformities, ulcers, callous formation, poor peripheral circulation, or history of amputation.



High Risk of Ulcers Amputation

Poor glycemic control

Peripheral neuropathy with LOPS

Cigarette smoking

Foot deformities

Preulcerative callus or corn

- Peripheral Arterial Disease
- History of foot ulcer
- Amputation
- Visual impairment
- Chronic kidney disease (especially if on dialysis)



ADA Stds – Exam components

- ▶ Inspection of the skin
- ▶ Assessment of foot deformities
- ▶ Neurological assessment (10-g monofilament testing with at least one other assessment: pinprick, temperature, vibration), and
- ▶ Vascular assessment including pulses in the legs and feet.



Lower Extremities

- ▶ Lift the Sheets and Look at the Feet



By Alton Johnson Jr., DPM, CWSP
<https://www.woundsource.com/blog/ampulation-crisis-african-american-patients>

12 Steps to Evaluate Lower Extremity Risk



**Lift the
Sheets and
Look at
the Feet**

Question 1 and 2

▶ **Question 1: Is there a history of foot ulcers?**

▶ **Question 2: Is there a foot ulcer now?**

▶ History of a foot ulcer increases the risk of developing another foot ulcer and increases the potential of future amputation.

▶ A person with a past or present foot ulcer is considered permanently in Risk Category 3.

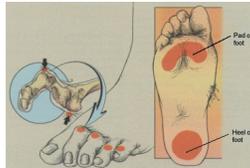


Question 3 – Deformity?

Question 3: Is there toe deformity?

Question 4: Is there an abnormal shape of the foot?

- ▶ Look for prominent bony areas,
- ▶ Partial or complete amputations of the foot or toes
- ▶ Clawed or hammer toes
- ▶ Bunions, or "Charcot Foot".



Question 3 and 4 – Charcot Foot

Question 3: Is there toe deformity?

Question 4: Is there an abnormal shape of the foot?

A Charcot Foot is a neuropathic foot that may present with:

- ▶ swelling,
- ▶ increased temperature,
- ▶ and little or no pain.
- ▶ Advanced cases show progressive signs of deformity into what is referred to as a "rocker bottom" or "boat-shaped" foot.
- ▶ A person with a Charcot Foot is permanently in Risk Category 3.



Q5 - Toenails

Question 5: Are the toenails thick or ingrown?

- ▶ Identify Mycotic, significantly hypertrophic, or ingrown nails.
- ▶ Ask how they are cutting their nails and identify problem areas.

Consider Podiatry Referral and Treatment



Q6: Callus Buildup

Question 6: Is there callus buildup?

- ▶ Identify focal and/or heavy callous.
- ▶ Determine cause and provide coaching.



Assess if the person is self-treating calluses (with a razor or other tools) and encourage them to see a foot specialist to prevent complications.



Poll Question 3

JR has dry skin cracks in the back of their heel. What is the best action?

- A. Gently scrape the dead skin with a razor
- B. Apply petroleum jelly or other lotion nightly to affected area.**
- C. Walk barefoot to promote healing
- D. Wear white cotton socks



Q7: Assess for Swelling

Question 7: Is there swelling?

Swelling may stem from a variety of causes such as a Charcot fracture, infection, or “venous stasis”.

Assess for potential causes and encourage the person to elevate extremities and receive treatment.



8- Check for Elevated Skin Temp

Question 8: Is there elevated skin temperature?

Elevated, localized skin temperature can indicate

- ▶ excessive mechanical stress,
- ▶ bone fracture
- ▶ or infection and requires further evaluation.



A temperature elevation of greater than 2 degrees centigrade or a noticeable difference by touch when compared with the contralateral foot is considered clinically significant and requires follow-up.

Q9 – Muscle Weakness

Question 9: Is there muscle weakness?

A manual muscle test of foot and great toe dorsi and plantar flexion. Weakness or inflexibility is associated with diabetes neuropathy and increases the risk of injury.



Prayer Sign

Q10 - See Bottom of Feet?

Question 10: Can the person see the bottom of his/her feet?

- ▶ Extra weight and/or lack of flexibility can make it difficult for people to visually assess their feet.
- ▶ Self-inspection and foot care are also difficult.



Q11 & 12 – How do the Shoes Fit?

Question 11: Are they wearing improperly fitted shoes?

- ▶ Can create foot pressures that lead to further complications.
- ▶ Sensory loss often results in wearing shoes that are too short and/or narrow resulting in ischemic ulcers on the medial or lateral metatarsal heads or the toes of a foot with claw toe deformity.
- ▶ Properly sized added depth shoes with soft custom molded insoles are usually indicated for those with loss of sensation and deformity to prevent ulceration.
- ▶ **Question 12: Is the footwear appropriate for their category?**



Poll question #4

▶ What is the most common cause of ulcers?

- A. Dr. Scholl's corn pads
- B. Minor trauma
- C. Trimming calluses
- D. Burns from hot water

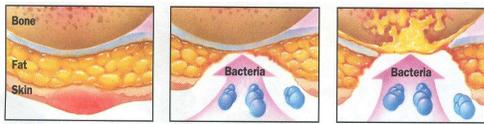


Common Causes of Ulcers

- ▶ Tight shoe and minor trauma
- ▶ Neuropathy and peripheral vascular disease
 - ▶ Autonomic: blood pooling, swelling
 - ▶ Motor: atrophic musculature, deformity, joint stiffness
 - ▶ Resulting increased plantar pressure, trauma



Foot Wounds



Blisters
Calluses

Ulcers

Bone infection

Risk Factors for Peripheral Arterial Disease

- ▶ Risk factors include:
 - ▶ diabetes
 - ▶ over the age of 60
 - ▶ hypertension,
 - ▶ hyperlipidemia,
 - ▶ who smoke, are at higher risk for PAD.

African Americans have 3-4 times increased risk of PAD
careful screening and appropriate intervention for these higher risk groups is imperative.



Symptoms of Peripheral Arterial Disease

What are symptoms of PAD?

- ▶ The classic symptom of PAD is pain in the legs with physical activity, such as walking, that gets better after rest.
- ▶ However, up to 4 in 10 people with PAD have no leg pain.
- ▶ Symptoms of pain, aches, or cramps with walking (claudication) can happen in the buttock, hip, thigh, or calf.



American College of Cardiology

Signs of Peripheral Arterial Disease

Physical signs

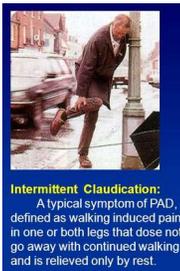
- ▶ include leg muscle atrophy (weakness);
- ▶ hair loss; smooth, shiny skin;
- ▶ skin that is cool to the touch, especially if accompanied by pain while walking (that is relieved by stopping walking);
- ▶ decreased or absent pulses in the feet;
- ▶ sores or ulcers in the legs or feet that don't heal; and cold or numb toes.



Peripheral Arterial Disease Intermittent Claudication

Physical Exam – Skin

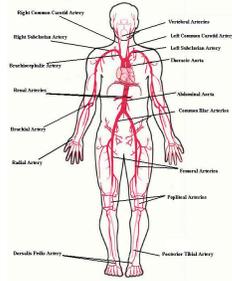
- ▶ Pale or blue, purple
- ▶ Dependent rubor, blanching when elevated
- ▶ Cool to touch, loss of hair, nonhealing wounds, gangrenous
- ▶ Diminished pulses
- ▶ Treatment = Protect feet
- ▶ Avoid constriction, increase walking, stop smoking, get ABI, medications and/or surgery



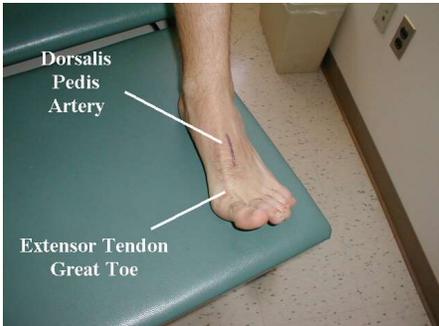
Intermittent Claudication:
A typical symptom of PAD, defined as walking induced pain in one or both legs that does not go away with continued walking and is relieved only by rest.

Vascular Status Assessment

- ▶ Posterior tibial pulse
- ▶ Dorsalis pedis pulse
- ▶ Temperature
- ▶ Appearance



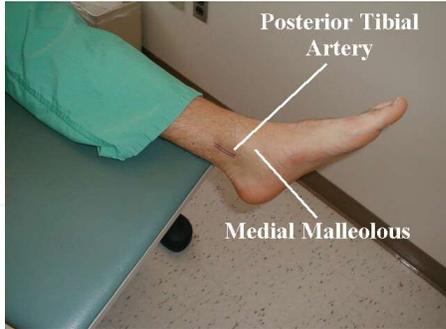
Dorsalis Pedis Pulse



Taking the Dorsalis Pedis Pulse



Posterior Tibial Pulse



Taking the Posterior Tibial Pulse

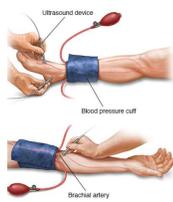


Refer. Include Multi-Disciplinary Team

- ▶ **If claudication or decreased/absent pedal pulses**
 - ▶ refer for ankle-brachial index and for further vascular assessment
- ▶ **Foot ulcers and high-risk feet**
 - ▶ Refer to multidisciplinary team (e.g., dialysis, Charcot foot, prior ulcers or amputation)
- ▶ **Foot care specialists recommended:**
 - ▶ those who smoke
 - ▶ histories of prior lower-extremity complications
 - ▶ loss of protective sensation
 - ▶ structural abnormalities
 - ▶ peripheral arterial disease
- ▶ **Ongoing preventive care lifelong surveillance.**

ADA Stds – Poor Circulation / High risk

- ▶ If have symptoms of claudication or decreased or absent pedal pulses:
 - ▶ refer for ankle-brachial index and for further vascular assessment as appropriate
- ▶ A multidisciplinary approach is recommended for individuals with foot ulcers and high-risk feet
- ▶ Use of specialized foot wear recommended



Using the ABI: An Example

Right ABI
80/160=0.50

Left ABI
120/160=0.75

ABI
(Normal >0.99)

Brachial SBP
150 mm Hg

Brachial SBP
160 mm Hg

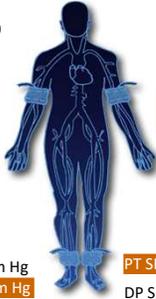
Highest brachial Systolic BP

PT SBP 40 mm Hg
DP SBP 80 mm Hg

PT SBP 120 mm Hg
DP SBP 80 mm Hg

Highest of PT or DP Systolic BP

ABI=ankle-brachial index; DP=dorsalis pedis; PT=posterior tibial; SBP=systolic blood pressure



Interpreting the Ankle-Brachial Index

ABI	Interpretation
1.00–1.29	Normal
0.91–0.99	Borderline
0.41–0.90	Mild-to-moderate disease
≤0.40	Severe disease
≥1.30	Noncompressible

from Hirsch AT, et al. J Am Coll Cardiol. 2008;47:e1-e192. Figure 6.

Loss of Protective Sensation (LOPS)

"I didn't notice"

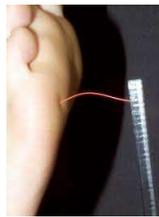
- ▶ Needle in foot
- ▶ Pebble in shoe
- ▶ Stepped on a nail
- ▶ Cut too deep
- ▶ Shoes were rubbing
- ▶ Others?



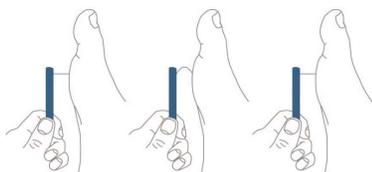
Loss of Protective Sensation

▶ Monofilament Testing

- ▶ 5.07 touched to plantar surface and top of foot
- ▶ C shape delivers 10 gms pressure
- ▶ Test four sites
 - Each great toe
 - 1st, 3rd and 5th metatarsal head



5.07 monofilament delivers 10gms linear pressure



Check Feet Daily

- ✓ Check and wash your feet daily. If you have trouble bending, use a mirror to see the bottom of your feet. Make sure to dry well and check in between toes.
- ✓ Let your provider know right away if you discover any sores, red areas, calluses, drainage, or unusual foot odor.
- ✓ Prevent dry skin and cracks by applying lotion or petroleum jelly to the top and bottom of your feet a few times a week.



Lotions – Apply to Top and Bottom



Education Points – Wear Shoes

- ✓ Avoid going barefoot, even inside, to avoid accidental injury.
- ✓ Buy new shoes at the end of the day when feet are most swollen.
- ✓ Break in new shoes gradually by wearing them for a few hours each day (1 hour the first day, 2 hours the second day, etc.).
- ✓ Inspect shoes for rough spots, torn linings, or other objects which could injure your feet. Make sure there is enough room to wiggle your toes.



ADA Standards - Shoes

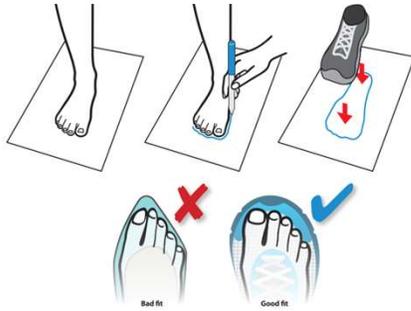
- ▶ Broad and square toe box
- ▶ Laces with 3-4 eyes per side or Velcro straps
- ▶ Padded tongue
- ▶ Quality lightweight materials
- ▶ Sufficient depth to accommodate a cushioned insole
- ▶ Custom shoes as needed
- ▶ Medicare approves 1 pair of custom shoes and 3 inserts yearly.



Dr. Comfort 6 Wide



Make Sure There is Enough Room



Feet Deserve Special Care



Medicare Pays for Therapeutic Shoes
Under the Therapeutic Shoe Bill, Medicare patients with diabetes are entitled to one pair of shoes and three pairs of orthotic inserts. This is in addition to the benefit already available under Medicare. You will need a doctor and take care of the paperwork. Our Certified Therapists come to your home to ensure a proper fit, fit style, material and ensure adherence to Medicare rules. Call today to get your shoes. Deductible or copayments may apply. Sorry, no HMO's.

- ▶ Daily inspection
- ▶ With order from MD and Loss of Protective Sensation (LOPS), Medicare Covers:
 - ▶ Annual custom shoes
 - ▶ 3 pairs of orthotic inserts

Foot Care Tips – Check Temp

- ✓ Avoid heating pads, Jacuzzis and hot water bottles. Use sunscreen to avoid sunburn.
- ✓ Since feet may not sense temperatures that are too hot or cold, you need to protect them. Wear warm socks or lined shoes if feet become cold.
- ✓ Use diabetes socks that are free of seams and not too tight around the calf.
- ✓ No bathroom surgery (this includes trimming calluses with a razor or liquid corn and callus removers). This can lead to injury.



Diabetes Socks



- Seamless
- Not too tight at calf
- Good cushion
- Cotton/poly blend
- Affordable

No Bathroom Surgery



Get Help and Prevent Injury

- ▶ Have a foot doctor trim your toenails if you cannot see or feel your feet, you cannot reach your feet, your toenails are thick or yellowed or your nails curve and grow into the skin.



Cutting Thick Toenails



<https://www.wikihow.com/Trim-Toenails-with-Fungus>

Tinea Pedis / Toenail Fungus

- ▶ Fungus that infects the areas between toes and skin of feet is called athlete's foot (tinea pedis).
- ▶ Onychomycosis, also called tinea unguium, is a fungal infection that affects either the fingernails or toenails.
 - ▶ Often subungual



For skin fungus, try an antifungal cream

For toenail fungus, topical or oral antifungals

ADA Stds – Education is Critical

► Based on risk assessment, review:

- Foot care, including nail and skin care
- Daily foot monitoring
- If have LOPS, how to evaluate feet status
- Footwear and home behaviors
- Identify resources if have trouble with cognition or physical constraints



Lower Extremities

► "If there is ANY foot problems, take off your shoes and socks and show your feet!"

- Complete foot exam annually
- More frequent checks on those at high risk
- Keep close eye if loss of protective sensation, foot deformities, or a history of foot ulcers



You Can Make A Difference

► Assess

- Nail condition, nail care, in between the toes
- Who trims your nails
- Have you ever cut your self?
- Shoes – type and how often
- Socks
- Skin/skin care and vascular health
- Ability to inspect
- Loss of protective sensation



FREE NO DeFEET Webinar – Invite your Friends and Colleagues



Thank You



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