

Nerve Disease

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Diabetes Education

# Diabetes - Microvascular Complications

- ▶ Microvascular Complications
  - ▶ Diabetic eye disease, nephropathy, and neuropathy disease





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# Diabetes - Microvascular Complications and Goals of Care

#### Objectives:

- ▶ Identify 3 microvascular complications
- Describe modifiable and non-modifiable risk factors for diabetes complications
- ▶ List screening guidelines





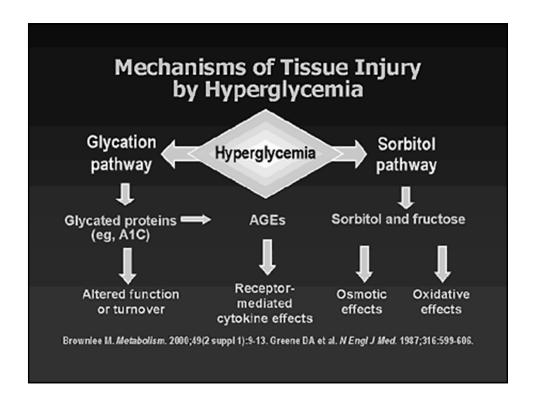


# **Quick Question**

- ▶ Does diabetes cause complications?
  - Yes
  - No
  - Uncontrolled diabetes causes complications





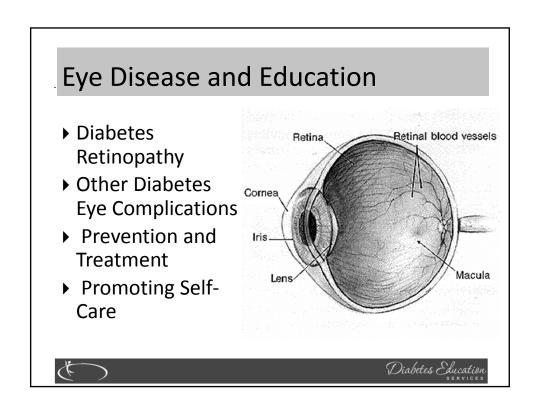


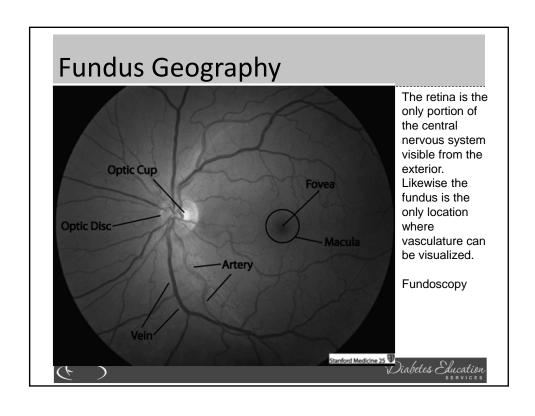
### **Quick Question 1**

- ▶ Which of the following are modifiable risk factors for microvascular disease?
  - Blood pressure, glucose levels, smoking
  - B. Age, type A personality, blood pressure
  - c. Ethnicity, blood pressure, diet
  - Blood glucose, genetics, activity level



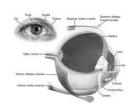




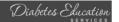


# Diabetes Eye Disease Overview

- ▶ Leading cause of adult blindness
  - ▶ Retinopathy and Diabetic Macular Edema
- ▶ Diabetes = 25x's risk of ocular complications
  - ▶ Including cataracts
- ▶ 20% of type 2 have retinopathy at diagnosis
- ▶ Only 60% of patients receive appropriate treatment







### **Retinopathy Risk Factors**

- ▶ Non- Modifiable:
  - ▶ Duration of diabetes, age at diagnosis, race other genetic factors

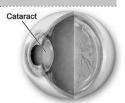


▶ Glycemic control, hypertension, smoking, hyperlipidemia, proteinuria and renal disease

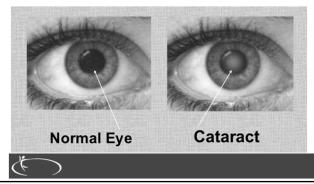




▶ Cataracts – elevated glucose levels glycosylate lens, decreasing permeability



▶ Treatment = surgery



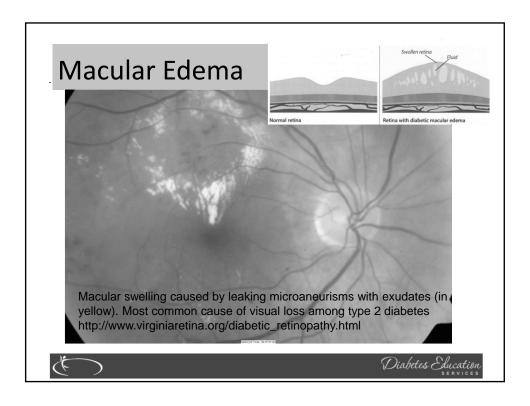




### Macular Edema

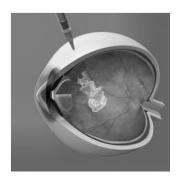
- ▶ Macular edema
  - ▶ Risk 10-15% for pt's with dm 15yrs +
  - ▶ macula responsible for central vision
  - ▶ retinal thickening w/in 3mm from the macula
  - ▶ can impair central vision causing blurring to blindness
  - ▶ Treatment
    - ▶ Focal laser treatment or
    - Monthly injections with VEGF (upcoming slide)





### New Approved Treatment for Macular Edema

- ▶ Anti-vascular endothelial growth factor (VEGF) therapy is indicated for diabetic macular edema
- ▶ VEGFs include:
  - ▶ Ranibizumab (Lucentis)
  - ▶ Avastin or
  - ▶ Eylea
- ▶ Improve vision with treatment
- ▶ Once a month injection



# Quick question 2

- ▶ Which of the following describes proliferative retinopathy?
  - A. Cotton wool spot and hemorrhages
  - B. Increased lens opacity
  - C. Stiffening of the lens
  - D. New blood vessel growth





# What is Retinopathy?

- ▶ Retina layer of nerve tissue in back of eye responsible for processing images and light
- ▶ Damage to the microvascular layer that nourishes the retina
- ▶ Leads to leakage of blood components through vessel walls and creation of unstable blood vessels secondary to hypoxia
- ▶ Disturbance in nerve layer = visual symptoms





### Natural History of Diabetic Retinopathy

- ▶ Mild nonproliferative diabetic retinopathy (NPDR)
  - ▶ Microaneurysms only
  - ▶ Reexamined annually
- ▶ Moderate NPDR
  - ▶ Microaneurysms plus other abnormalities
  - ▶ Reexamined w/in 6-12 months



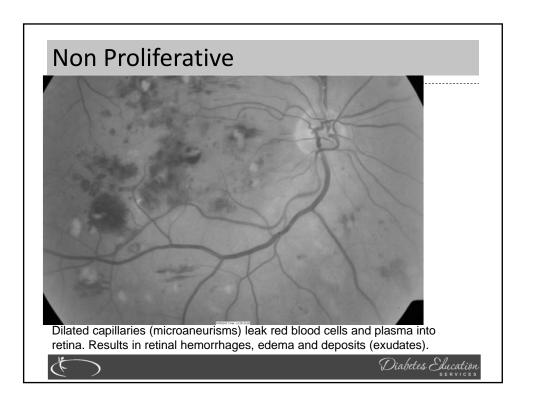


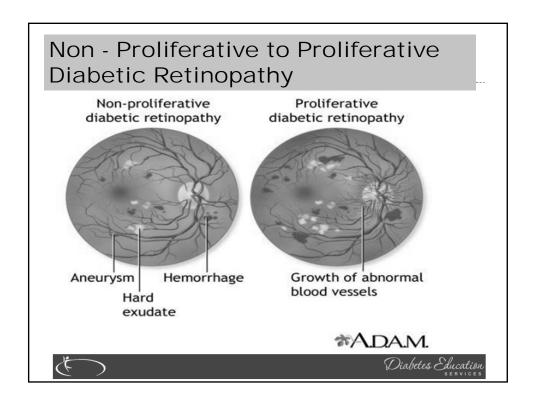
# Severe non-proliferative retinopathy

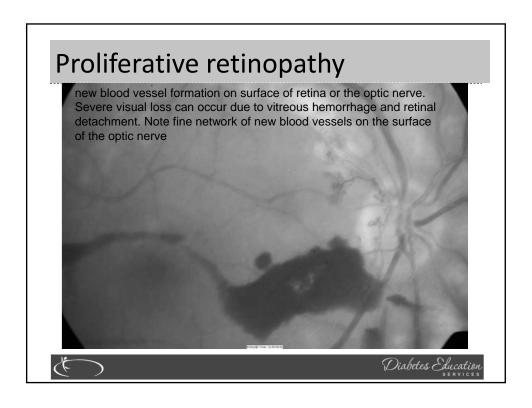
- ▶ Any of the following:
  - ▶ 20+ intraretinal hemorrhages in each 4 quadrants
  - ▶ Venous beading in 2 or > quadrants
  - ▶ Prominent intraretinal microvascular abnormalities in 1 or more quadrant
  - ▶ No signs of proliferative disease
- ▶ Reexamination several times a year

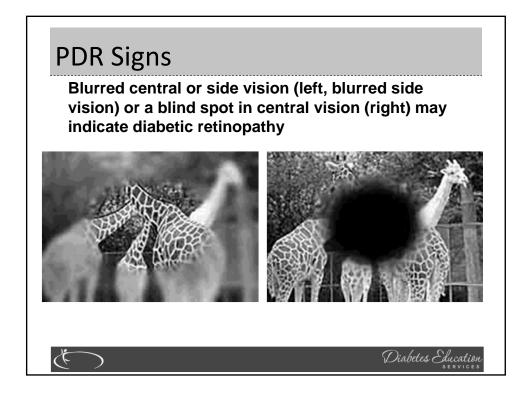












# Retinopathy Changes How We See



View of boys by person with normal vision



View of boys by person with diabetic retinopathy.





### Proliferative Diabetic Retinopathy (PDR)

- ▶ Clinical Findings
  - ▶ Ischemia induced neovascularization
    - ▶ at the optic disk (NVD)
    - ▶ elsewhere in the retina (NVE)
  - ▶ Vitreous hemorrhage
  - ▶ Retinal traction, tears, and detachment
  - ▶ Diabetes Macular Edema must also be evaluated



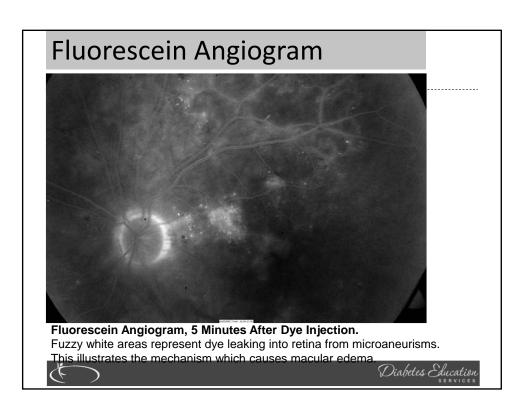


# PDR Management

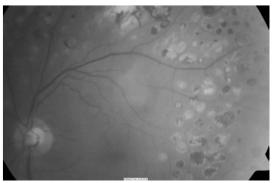
- ▶ Management/Treatment
  - ▶ 2-4 month follow-up
  - ▶ Color fundus photography
  - ▶ Panretinal photocoagulation (3-4 month follow-up)
  - ▶ Vitrectomy if bleeding into vitreous
  - ▶ If macular edema present: fluorescein angiography and injected meds







# Pan Retinal Photocoagulation



Decreases risk of severe vision loss by 50% or more Destroys 12% of retina and loss of visual field. Once stabilized, can achieve excellent control of PDR if B/P and BG well controlled.





# **Retinopathy Prevention**

- ▶ To reduce the risk or slow the progression of retinopathy
  - ▶ Optimize glycemic control
  - ▶ Optimize blood pressure control





### **Quick Question 3**

- ▶ Which of the following is correct regarding eye screening for people with diabetes?
  - A. All people with diabetes must get a complete eye exam every year
  - B. All people diagnosed with type 1 should receive an immediate eye exam.
  - C. All people diagnosed with type 2 should receive an immediate eye exam.
  - D. People with diabetes over age of 60 should receive an eye exam every 6 months.



# **Retinopathy Screening**

- Screen with initial dilated and comprehensive eye exam by ophthalmologist or optometrist
- ▶ Type 2 at diagnosis, then every 1 to 2 years
- ▶ Type 1 within 5 yrs of dx, then every 1-2 years
- ▶ Can use high quality fundus photography as screening tool- Initial exam should be done in person
- Promptly refer pts with macular edema, and severe non-proliferative disease to trained specialist



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### High Quality Fundus Photography to Screen for Retinopathy

Can detect most clinically significant diabetic retinopathy

- ▶ Interpretation of the images
  - ▶ Performed by a trained eye care
- May serve as a screening tool for retinopathy, it is not a substitute for a comprehensive eye exam
- Perform comprehensive eye exam at least initially and at intervals thereafter



### **Retinopathy Screening**

- ▶ Women with preexisting diabetes who are planning pregnancy or are pregnant
  - ▶ Comprehensive eye examination in the first trimester
  - ▶ Close follow-up throughout pregnancy and for 1 year postpartum



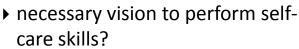
# **Ongoing Retinopathy Screening**

After initial exam, then...

- Annual exam
- Less frequent (every 2-3 yrs) can be considered if 1 or more normal eye exam
- More frequent exams if retinopathy progressing



# Assess adaptation to low vision





- ▶ insulin
- ▶ BGM
- ▶ read instructions
- shopping/home safety/transportation
- ▶ refer to rehab education (800-AFBLIND)
- psychosocial issues



# Prodigy Voice Meter – A+ Access Award Am Fed Blind

Prodigy Meter only completely accessible meter for sale in U.S. independent access for visually impaired -

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- •800-243-2636
- •Independent set up
- Self coding
- Audible Memory
- Audible warning/error
- •Cost: \$84.95
- •50 Strips \$34.95







# **Nephropathy Objectives**

- ▶ Epidemiology of diabetes nephropathy / Kidney Disease
- ▶ Basic functions of the kidney
- ▶ Major stages in progression of nephropathy
- ▶ Diagnostic tests to assess and monitor renal function
- ▶ Treatment and prevention







### **Quick Question 4**

- ▶ John's dad had diabetes and kidney failure. He wants to learn the risk factors for kidney disease. Which of the following describes those at increased risk for kidney disease?
  - A. Excessive alcohol intake and daily Tylenol
  - B. Family history of kidney disease, smoking
  - C. High protein diet and excessive trans-fat intake
  - D. Diet high in processed foods and sodium





# **Kidney Physiology**

- ▶ Size and shape of Idaho potato retroperitoneal
- ▶ Filter entire blood volume every 30 minutes
- excretory organ:
  - removes water, urea, waste
  - maintains blood volume
  - acid base balance and lytes
  - ▶ regulates B/P
  - synthesizes erythropoietin RBC
  - ▶ Maintains calcium /phosphorus levels, activates vitamin D – helps absorb calcium





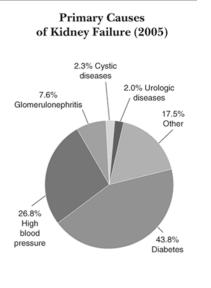
# Risk Factors of Kidney Disease

- ▶ 2 leading risk factors: Hypertension and hyperglycemia
- ▶ Other risk factors:
  - Kidney stones, obesity, smoking and CV disease
  - Family history of kidney disease and age 60 or older
- Kidney disease often has no symptoms, can undetected until very late



## **Diabetic Nephropathy**

- Most new cases of Chronic Kidney Disease (CKD) are attributed to diabetes.
- 220,000 people in US have kidney failure due to diabetes (2013)
- Minorities experience higher than average rates of nephropathy and kidney disease



# Diabetes and Chronic Kidney Disease (CKD) Considerations

- ▶ CVD leading cause of death in CKD
  - ▶ albuminuria = increased risk of CVD
- ▶ 1/4 to 1/3 of insulin cleared by kidney
- renal retinal syndrome
- ▶ 70 80% of people with diabetes DON'T get kidney disease
- ▶ Early and aggressive intervention crucial

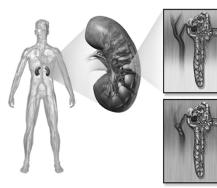






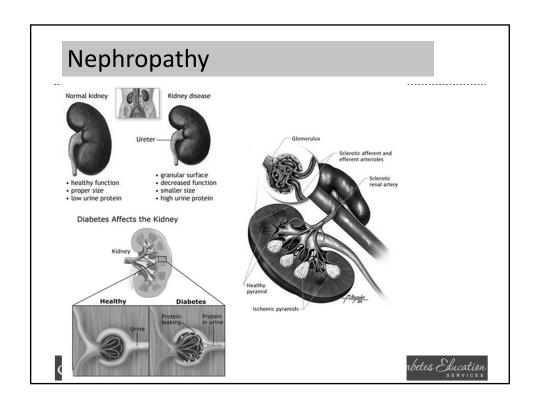
### What is Nephropathy?

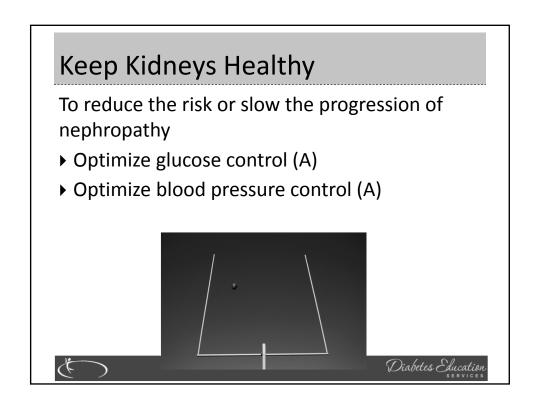
- ▶ Hyperglycemia causes renal hyperfiltration and glomerular capillary hyperperfusion.
- Causes functional and structural damage to glomeruli, increasing permeability, proteinuria, mesangial expansion and sclerosis... destroys nephrons
- ▶ Due to insufficient insulin, glycosylation, increased growth hormone, glucagon, and vasoactive hormones.











# Screening for Kidney Disease

- ▶ Screen for:
  - ▶ Urine Albumin-Creatinine Ratio (UACR) and
  - ▶ Glomerular Filtration Rate (GFR):
- ▶ Type 2 at dx then yearly
- ▶ Type 1 with diabetes for 5 years, then yearly
- Measure serum creatinine and GFR yearly
- ▶ Treat hypertension and intensify as needed







#### **Definitions of Albumin Excretion**

▶ Urine albumin – creatinine ratio (spot collection)

mg/g creatinine Category

<30 ▶ normal

▶ Increased urinary albumin excretion 30-299

- ▶ 2 of 3 tests w/in 3-6 mo abnormal to confirm
- ▶ Exercise within 24 h, infection, fever, CHF, marked hyperglycemia, and marked hypertension may elevate urinary excretion over baseline values.



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# Stages of Chronic Kidney Disease

Table 9.2	2—Stages of CKD	
Stage	Description	GFR (mL/min/1.73 m <sup>2</sup> )
1	Kidney damage* with normal or increased GFR	≥90
2	Kidney damage* with mildly decreased GFR	60–89
3	Moderately decreased GFR	30–59
4	Severely decreased GFR	15–29
5	Kidney failure	<15 or dialysis

<sup>\*</sup>Kidney damage is defined as abnormalities on pathological, urine, blood, or imaging tests. Adapted from Levey et al. (37).





DA Management of CKD			
Table 9.3—Managem GFR (mL/min/1.73 m <sup>2</sup> )	ent of CKD in diabetes (7)  Recommended management		
All patients	Yearly measurement of creatinine, urinary albumin excretion, potassium		
45–60	Referral to a nephrologist if possibility for nondiabetic kidney disease exists (duration of type 1 diabetes <10 years, persistent albuminuria, abnormal findings on renal ultrasound, resistant hypertension, rapid fall in GFR, or active urinary sediment on ultrasound)  Consider the need for dose adjustment of medications Monitor eGFR every 6 months  Monitor electrolytes, bicarbonate, hemoglobin, calcium, phosphorus, parathyroid hormone at least yearly  Assure vitamin D sufficiency  Consider bone density testing  Referral for dietary counseling		
30–44	Monitor eGFR every 3 months  Monitor electrolytes, bicarbonate, calcium, phosphorus, parathyroid hormone, hemoglobin, albumin, weight every 3–6 months  Consider the need for dose adjustment of medications		
<30	Referral to a nephrologist		

# Kidney disease treatment - ADA

- ▶ ACE or ARB NOT recommended for prevention of kidney disease if BP normal and urinary albumin excretion (UAE) < 30 mg/g (in pts w/ diabetes)
- ▶ ACE or ARB if UAE of >30 mg/g
- ▶ Monitor creat and K+ when on ACE or
- ▶ When GFR < 60, evaluate/manage potential complications of CKD
- Consider referral to specialist when management is difficult and kidney disease is advanced
- Protein restriction no longer recommended







# Treatment of Chronic Kidney Disease (CKD)

There are four primary treatment options for individuals who experience ESRD:

- 1. Hemodialysis
- 2. Peritoneal Dialysis
- 3. Kidney Transplantation
  - ▶ 120, 000 Americans waiting for kidney
  - ▶ Only 17,000 receive one each year
  - ▶ Every day, 12 people die waiting for a kidney
- 4. No treatment





# Psychosocial Issues associated with **Chronic Kidney Failure**



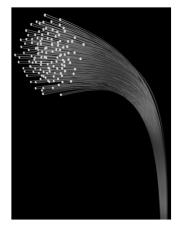
- ▶ depression
- ▶ stress
- anxiety
- support groups, counseling and coping skills





# **Diabetes Nerve Disease Objectives**

- ▶ Causes of neuropathy
- ▶ Different types of neuropathy
- ▶ Detection, prevention and treatment
- ▶ Key info to teach about neuropathy



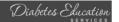




### Microvascular Disease and Polyol Theory

- ▶ hyperglycemia ↑ glucose level in cells
- ▶ sorbitol pathway glucose reduced to sorbitol by aldose reductase
- ▶ polyol pathway sorbitol oxidized to fructose by sorbitol dehydrogenase
- ▶ glucose, sorbitol, fructose toxic to cells
- ▶ Unerve velocity, oxygenation, increases oxidative stress





### Quick question 5

- Mary has had diabetes for 10 years and wants to reduce her risk of neuropathy. What are most important steps she can take to limit risk?
  - A. Lose weight and decrease coffee intake
  - B. Control blood glucose
  - C. Take vitamin B12 daily
  - D. Apply capsaicin cream to extremities twice daily.





# What is Neuropathy?

- ▶ Diabetic Neuropathy (DN) = demonstrable nerve disorder and destruction, either clinical or subclinical- that occurs w/ diabetes, w/out other causes (10% of neuropathy due to other causes)
- ▶ 2 abnormalities present (symptoms, signs, abnormal quantitative test results)







# **Neuropathy Risk Factors**

- Age
- ▶ Hypertension
- ▶ Hyperglycemia
- ▶ Elevated LDL
- Smoking
- ▶ Overweight
- Excess alcohol
- Nutrition (eat lots of omega-3 fatty acids)
- Lack of exercise



### **Quick Question 6**

- What 2 office tests can be used to detect diabetes neuropathy?
  - A. Pin prick and electrophysiology testing
  - B. Monofilament and tuning fork
  - C. Hot/Cold discrimination testing
  - D. Babinski reflex assessment





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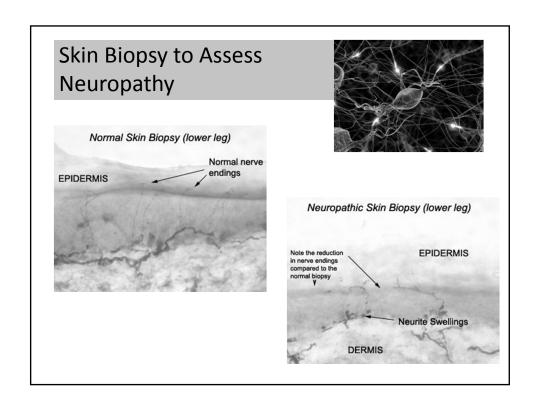
### Nerve disease Screening

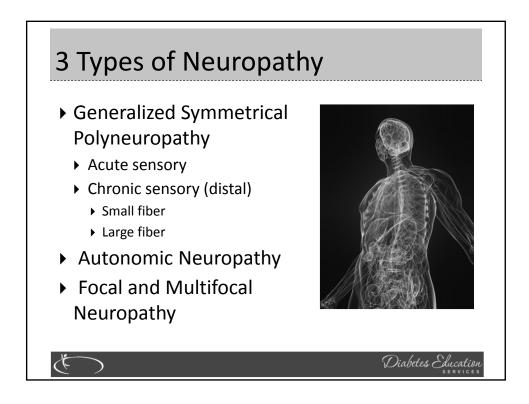
- Screen all patients for nerve disease using simple tests, such as a monofilament
  - ▶ Type 2 at diagnosis, then annually
  - ▶ Type 1 diabetes at 5 years, then annually
  - ▶ Tight glycemic control is the only strategy shown to prevent or delay the development and progression of neuropathy (ADA)
  - ▶ Assess and treat patients to reduce pain and symptoms to improve quality of life.











### Generalized Symmetrical Polyneuropathy -Acute Sensory Neuropathy

- ▶ Severe pain, wasting, weight loss, depression and erectile dysfunction
- ▶ Foot pain- burning, unremitting, deep, sharp, stabbing, "shock like"...worse at night, hypersensitive to light touch
- Associated w/ hyperglycemia or w/ rapid improvement of glucose
- ▶ Goal improve BG resolve in year



#### **Generalized Symmetrical Polyneuropathy** Chronic Sensorimotor Neuropathy Small Nerve Fiber

- ▶ Sensory deficits in distal portions, spreading medially "stocking-glove"
- ▶ Small Nerve Fiber Neuropathy
  - ▶ C-fiber pain = burning and superficial
  - Allodynia (all stimuli interpreted as painful)
  - ▶ Later, loss of pressure and temp sensation
  - Decrease blood flow, sweating
  - Detect w/ Monofilament
  - ▶ High risk for ulceration, Charcot, gangrene





#### **Generalized Symmetrical Polyneuropathy** Chronic Sensorimotor Neuropathy — *Large Nerve Fiber*

- ▶ Involve sensory and/or motor nerves
- ▶ Fibers are myelinated, rapid conductors
- ▶ Can detect destruction w/ nerve testing
- ▶ Symptoms may be minimal:
  - ▶ Impaired vibration perception/position sense
  - ▶ Ataxia "moon-walking", in-coordination
  - ▶ Pain described as deep-seated gnawing
  - ▶ Shortening of Achilles tendon and claw foot
  - Increased blood flow "hot foot"





# **Treating Neuropathy**

- ▶ Improve glycemic control
- ▶ Control pain
- ▶ Relief from depression from chronic pain
  - Massage, stretching, pain control clinic, TENS, avoiding alcohol, relaxation exercises....





#### Pharmacologic Therapy for Neuropathy

Try Alpha lipoic acid: 600 – 1,800mg /day Prescription Therapy

#### 1<sup>st</sup> line

- Tricyclic antidepressants (ie amitriptyline, nortriptyline
- Calcium channel modulators (ie gababentin, pregabalin)
- Serotonin Norepinephrine Reuptake Inhibitors (SNRI)

#### 2<sup>nd</sup> line

- **Topical Capsaicin Cream**
- Opioids (tramadol, oxycodone)

#### Reasons for treatment failure:

 Dose too low, inadequate trial, pt expecting elimination of symptoms, not changing class when no response Ziegler, D Painful diabetic neuropathy. Diabetes Care, 2009





### **Quick Question 7**

- ▶ Which of the following patients are at most risk for developing diabetes autonomic neuropathy?
- A. Diabetes for 1 year with A1c of 7.6%
- B. Person with diabetes for 16 years with A1c never above 6.9%
- c. Person with type 1 diabetes for 8 years with retinopathy
- D. Person with type 2 for 19 years with A1c less than 7.5%





# "DAN" Diabetic Autonomic Neuropathy

- ▶ 50% of pt's with peripheral neuropathy also have DAN
- DAN increases M & M rates
  - ▶ neurogenic bladder, sexual dysfunction
  - ▶ GI related disorders / gastroparesis
  - orthostatic hypotension
  - ▶ fixed heart rate, silent MI, sudden death
  - hypoglycemia unawareness
  - sudomotor, pupillary





# Sexual Functions as We Age

▶ 20-30 years trice daily ▶ 30-40 years tri weekly ▶ 40-50 years try weekly ▶ 50-60 years try weakly



▶ 60-70 years try oysters

▶ 70-80 years try anything

▶ 80-90 years try to remember

A touch of humor from AADE-New Perspectives on Erectile Dysfunction, 1999



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# **Erectile Dysfunction**



- ▶ Affects about 50% of men with diabetes
- ▶ Loss of erections sufficient for intercourse
- ▶ Due to combo of vascular and nerve damage
- ▶ Tests: penile tumescence to eval if organic or psychogenic
- ▶ Treatment:
  - ► Sildenafil (Viagra), Vardenafil (Levitra), Tadalfil (Cialis)
    - ▶ Use caution if taking nitrate drugs. Check w/ MD first
  - Other meds, vacuum devices, prosthetics
  - ▶ HRT- testosterone gel, patches, injections, pills







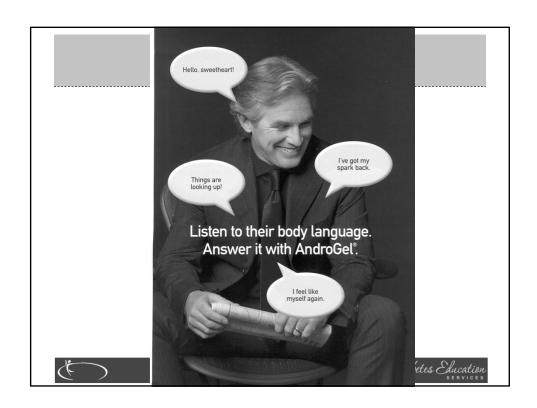
### Take Charge. Talk T.

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- ▶ Men w/ DM, 2x risk of low testosterone levels
  - > Symptoms include low sex drive, ED, depression, lack of energy and vitality
  - ▶ Low T easily diagnosed and managed, only 10% of men currently treated
  - ▶ Initial Screening:
    - ➤ Total testosterone: if < 300 ng/dl = hypogonadal
    - am testing preferred, repeat to confirm
  - ▶ Treatment: determine cause, testosterone replacement therapy







# **Focal Neuropathies**

- ▶ Often occurs in middle aged pt's or those w/ polyneuropathy
- ▶ 4 major focal neuro
  - ▶ mono compression or entrapment
    - carpal tunnel most common
  - plexopathy- femoral neuropathy
    - pain from hip to ant and lat aspects of thigh
  - radioculopathy intercostal neuropathy
  - cranial abrupt onset, HA, eye pain







### **Neuropathy Key Considerations**

- ▶ Very common long-term complication often not recognized and treated
- ▶ Management / treatment complex
- ▶ Thorough history /assessment critical
- ▶ Treatment based on underlying process, presentation, and cost effectiveness
- ▶ Treatable condition with new therapies on horizon.





### The ABC's of Diabetes Control

- A A1c less than 7%
- **B** Blood pressure less than 140/90
- **C** Cholesterol HDL > 40, Triglycerides < 150
- D Drugs- Keep list for emergencies/ MD
- **E** Exercise and Eyes
- F Food and Feet
- **G** Glucose checks and goals
- H- Healthy Coping Hoorah for your hard work!





