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Complications of Diabetic Ischemia



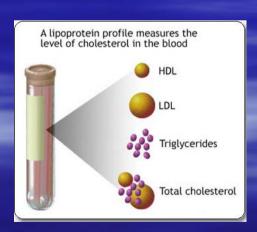
Adie Viljoen
MBChB, MMed, FRCPath, FCP, MBA
Lister Hospital, Stevenage
Honorary Consultant Cambridge University Hospitals NHS Trust

Job number: PHGB/VOK/0515/0031 Date of Preparation: May 2015











Case 1*

- 56 year old man
- Admitted on 24th Dec 2013 with acute onset of chest pain

- Past medical history
 - Type 2 diabetes (diagnosed 3 years ago)
 - Hypertension
 - Current smoker 10-20/day
 - Cholecystectomy

Medication

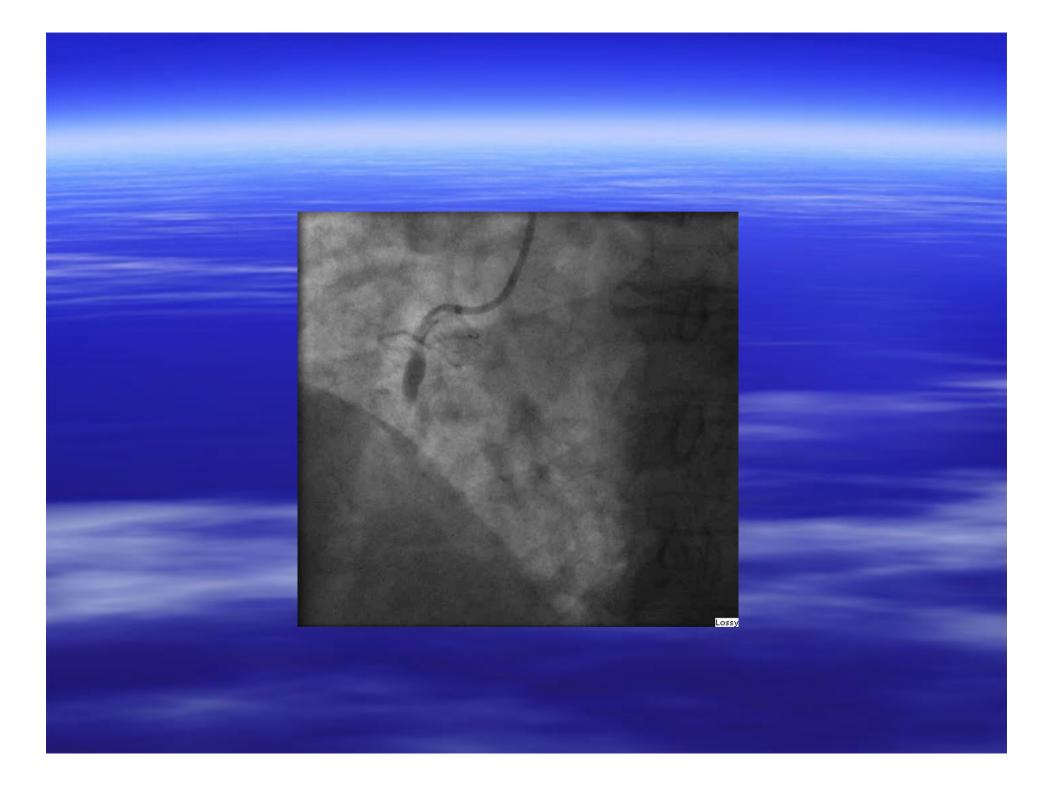
- Simvastatin 40 mg
- Candesartan 8 mg
- Metformin 1 g bd
- Gliclazide 160 mg bd

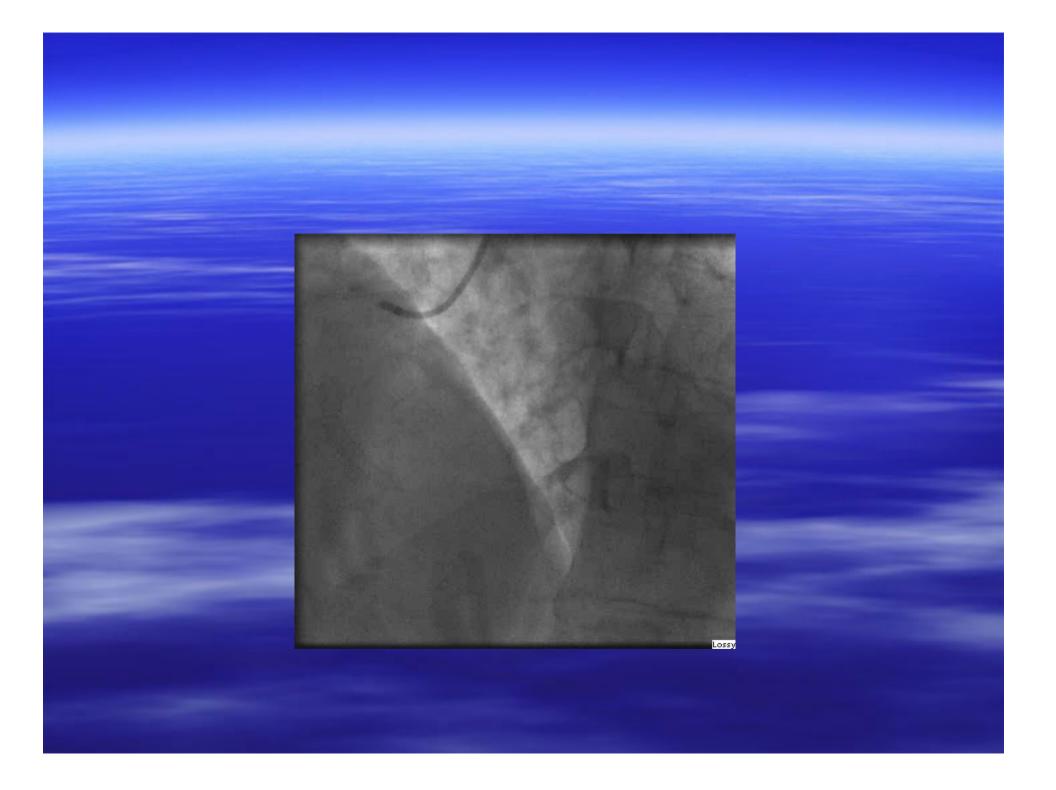


Examination & results

- Weight 111 kg
- BMI 36
- BP 127/83 mm Hg
- O₂ saturation 93%

Status: Printed Report Date: 1





Case 2*

- 58 year old male
- Type 2 diabetes
- Diagnosed 4 years ago



- Family history:
 - Mother †(68 MI), had type 2 diabetes for 8 years
- PMH:
 - Not significant
- Life style:
 - Doesn't get much exercise (lorry driver)
- Medication:
 - Metformin 1g b.d.

- Clinical examination
 - Overweight
 - Weight 101 kg (222 lb, 16 stone)
 - BMI = 33 kg/m²
 - Waist circumference 103 cm (40 inches)
 - BP 144/86 mm Hg
 - Rest examination NAD

- Special Investigations
 - HbA1c 7.7% (61 mmol/mol)
 - Fasting glucose 10.5 mmol/L
 - Total cholesterol 6.4 mmol/L
 - HDL-C 0.8 mmol/L
 - TG 4.6 mmol/L
 - Creatinine 88 umol/L
 - ACR 2.9 mg/mmol (<2.5)

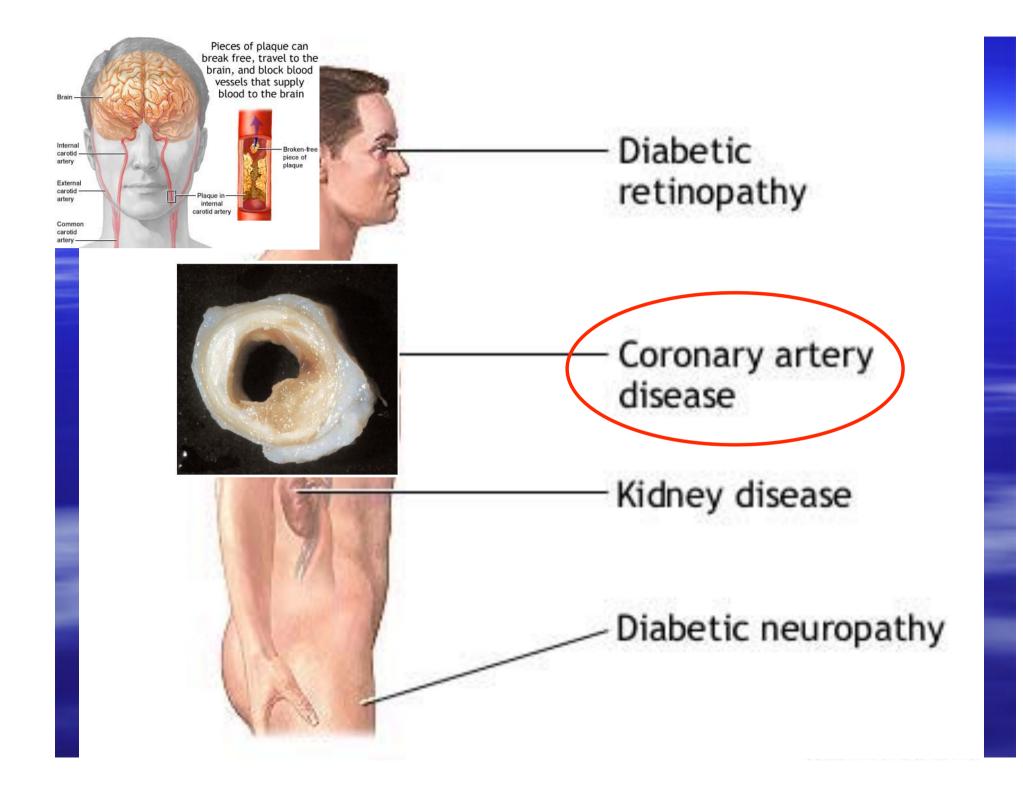
Case vignette - Treatment

- Life style
- Medication
 - Control hyperglycaemia?
 - Lipid lowering medication?
 - Anti-hypertensive?

What is he of risk at?
When did the problems start?
How should this be managed?

What is he of risk at? When did the problems start? How should this be managed?

What do patients with diabetes die of?

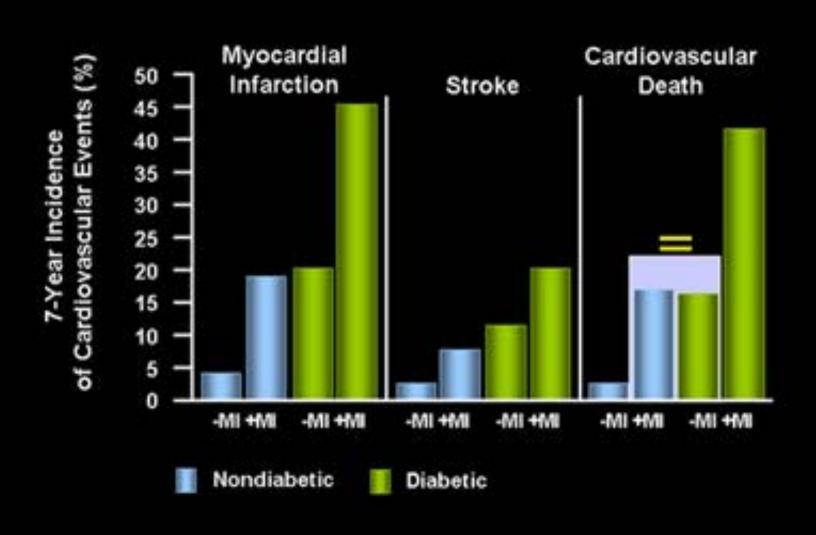


Diabetes = CVD

Up to 80% of diabetics will die of Cardiovascular disease

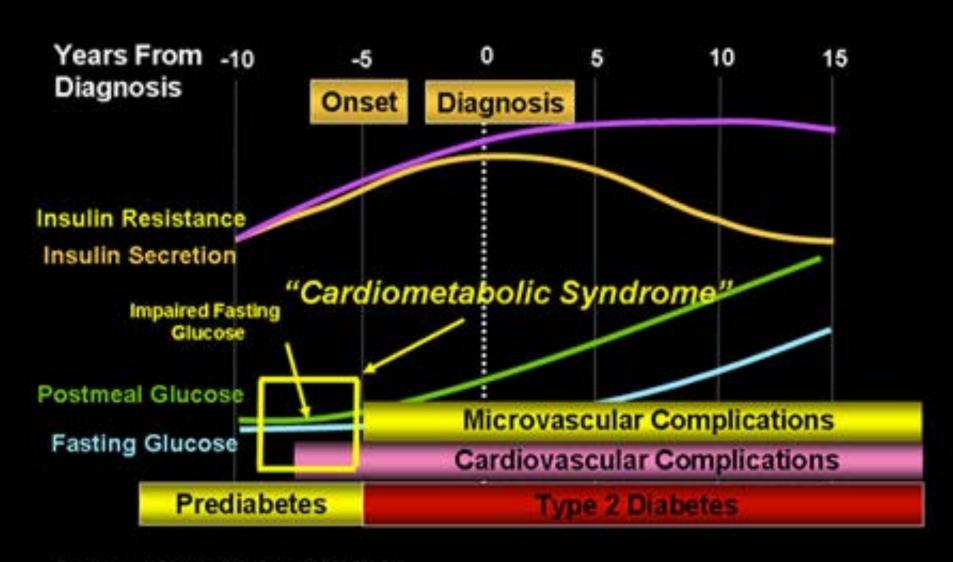


Diabetes Is a "CHD Equivalent"



What is he of risk at? When did the problems start? How should this be managed?

Natural History of Type 2 Diabetes



Ramlo-Halsted BA, et al. Prim Care. 1999;26:771-789. Nathan DM. N Engl J Med. 2002;347:1342-1349.

What is he of risk at? When did the problems start? How should this be managed?

Treatment of Diabetes

- **-** A
- <u>-</u> B
- C
- D
- <u>, E</u>
- <u>.</u> F

Treatment of Diabetes - life style

- A
- B
- C
- Diet, Don't smoke
- Exercise, Education
- <u>-</u> F





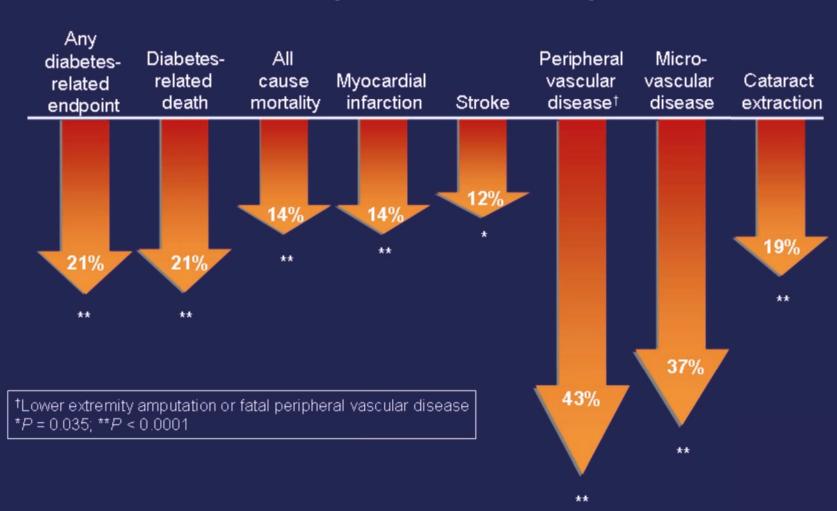
Treating diabetes

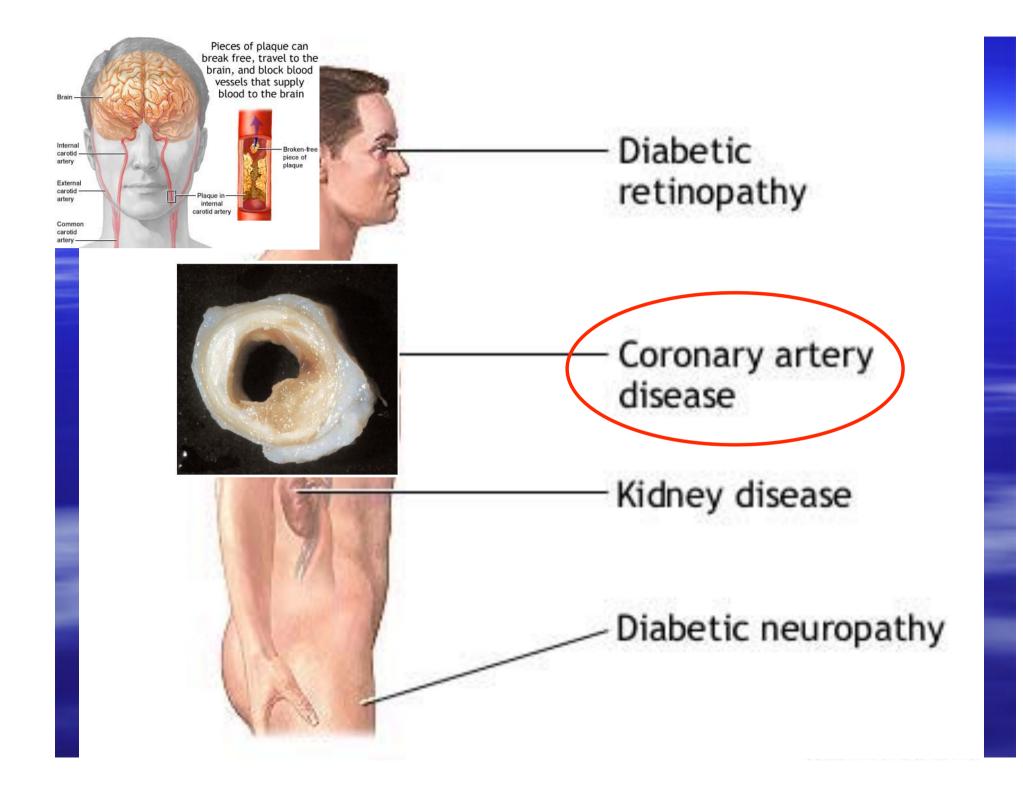
- A HbA1c
- B
- C
- D
- F

Percentage decrease in relative risk corresponding to a 1% decrease in HbA1C

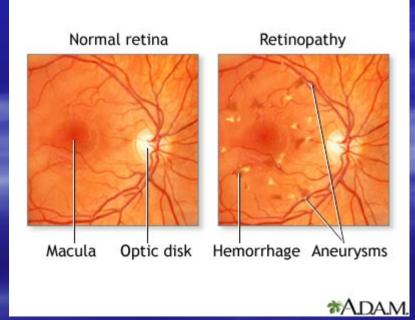
UKPDS: decreased risk of diabetes-related complications associated with a 1% decrease in A1C

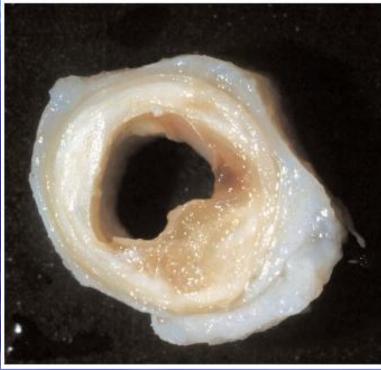
Observational analysis from UKPDS study data











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A correction has been published: N Engl J Med 2007;357(1):100.



Effect of Rosiglitazone on the Risk of Myocardial Infarction and Death from Cardiovascular Causes

Steven E. Nissen, M.D., and Kathy Wolski, M.P.H.

CONCLUSIONS

Rosiglitazone was associated with a significant increase in the risk of myocardial infarction and with an increase in the risk of death from cardiovascular causes that had borderline significance. Our study was limited by a lack of access to original source data, which would have enabled time-to-event analysis. Despite these limitations, patients and providers should consider the potential for serious adverse cardiovascular effects of treatment with rosiglitazone for type 2 diabetes.

N.B. Medicine no longer available in the UK



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Steven Nissen

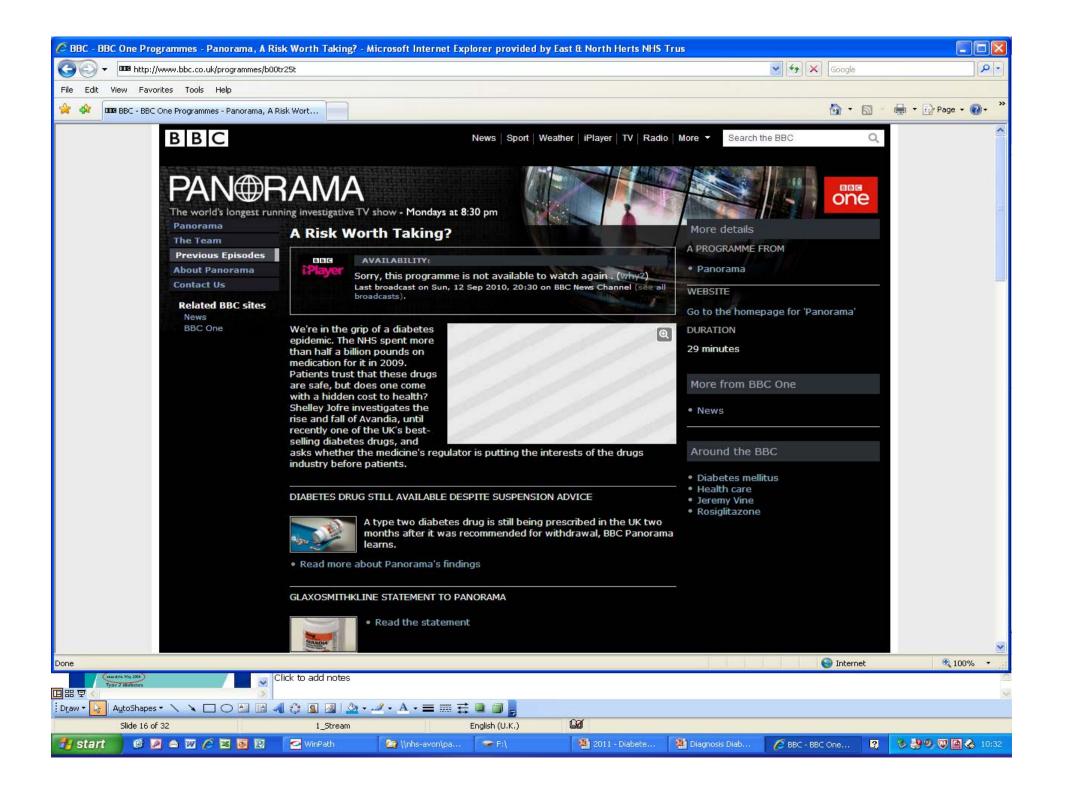
By Dr. Mehmet Oz

n some ways, Steve Vissen's undergraduate rears at the University of Michigan—which ncluded crusading igainst the Vietnam War—were the toughest

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oeriod of his life. But his extracurricular ctivism was just the training required for his





Numerous Studies Assessing CV Outcomes in T2DM Drugs Are Either Recently Completed or Ongoing

Trial Name	Drug	Target Enrollment	Timing*
DPP-4 Inhibitors			
SAVOR	Saxagliptin	N=16,492	Began 2010; Complete
EXAMINE	Alogliptin	N=5384	Began 2009; Complete
TECOS	Sitagliptin	N=14,000	Began 2008; Ending 2014
CAROLINA	Linagliptin	N=6000	Began 2010; Ending 2018
CARMELINA	Linagliptin	N=8300	Began 2013; Ending 2018
GLP-1 Agonists			
ELIXA	Lixisenatide	N=6000	Began 2010; Ending 2014
EXSCEL	Exenatide	N=9500	Began 2010; Ending 2017
LEADER	Liraglutide	N=9340	Began 2010; Ending 2016
REWIND	Dulaglutide 🔻	N=9622	Began 2011; Ending 2019
SUSTAIN 6	Semaglutide**	N=3260	Began 2013; Ending 2016
SGLT-2 Inhibitors			
CANVAS	Canagliflozin	N=4410	Began 2009; Ending 2018
C-SCADE 8	Empagliflozin V	N=7000	Began 2010; Ending 2018
DECLARE	Dapagliflozin	N=17,150	Began 2013; Ending 2019

^{*}Trial ending dates are anticipated based on publicly available information.

^{**}Not licensed in UK

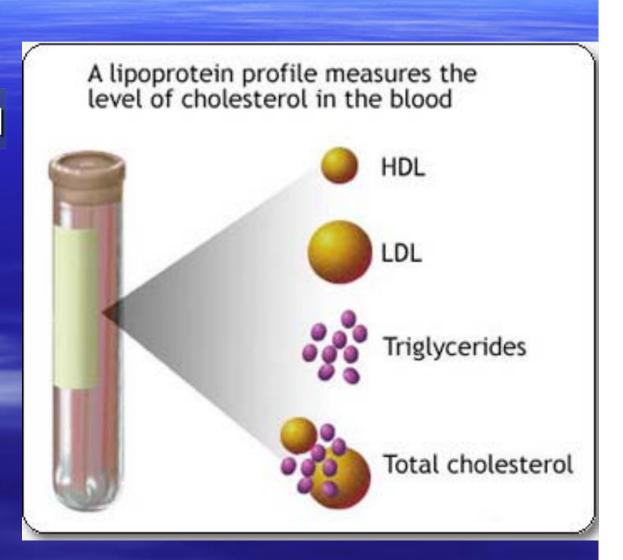
Treating diabetes

- A
- C
- D
- **-** F
- F

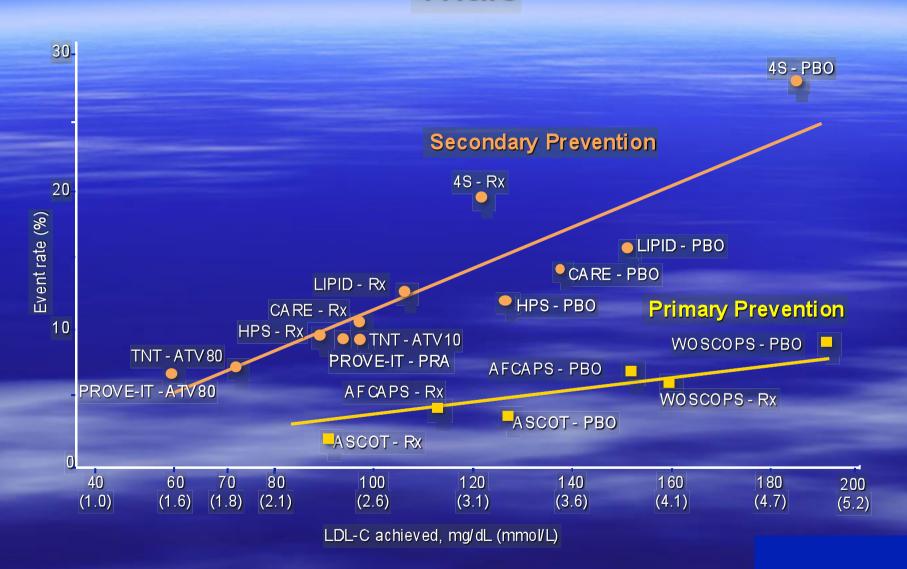
Cholesterol

Lipid profile

- The Lipid Profile
 - Total Cholesterol
 - HDL-cholesterol
 - LDL-cholesterol
 - Triglycerides



On-Treatment LDL and CHD Events in Statin Trials



Adapted from Rosenson RS. Expert Opin Emerg Drugs. 2004;9:269-279. LaRosa JC et al. N Engl J Med. 2005;352:1425-1435.





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The Dangers of Low Blood Cholesterol, Bad Cholesterol: A Myth and a Fraud? www.shirleys-wellness-cafe.com/cholesterol.htm - Cached

The Undiscovered **Danger** of **Statin** Drugs 🕸

8 Aug 2009 ... When you find out you have high cholesterol the doctor prescribes a drug to help out. Is that drug really a safe choice?

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15 Apr 2009 ... Researchers revealed that statins not only lowered the overall risk of stroke, but also slowed progression of blockages in the carotid ...

www.dailymail.co.uk/.../Cholesterol-lowering-statins-cut-danger-stroke-20-cent.html - Similar

The Dangers of **Statin** Drugs 😭

Danger: Gastrointestinal disturbances. The most common side effects of statins are common to many drugs. Nausea, upset stomach, constipation, diarrhea, ... www.squidoo.com/dangersofstatindrugs - Cached - Similar

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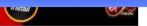
Statin Drug Side Effects

Your doctor might not be aware of the dangers of cholesterol drugs! www.healthy-heart-guide.com

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The other side of statins:
They've saved countless lives but now doctors fear for some,
the side effects could be
devastating

By JO WATERS
Last updated at 9:39 AM on 30th March 2010

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Smallholder Paul Rhoades was counting his sheep - as he did every day - when he stopped, unable to focus or remember what he was doing. 'I couldn't remember which sheep I'd counted and which I hadn't,' he recalls. 'Counting was beyond me.'

Paul, who was then aged 57, was concerned. I had a flicker of panic that it





FEMAIL TODAY

Wayne Rooney prostitute Jennifer Thompson shamelessly breaks cover as disgraced fooballer jets off with England squad Both looked glum



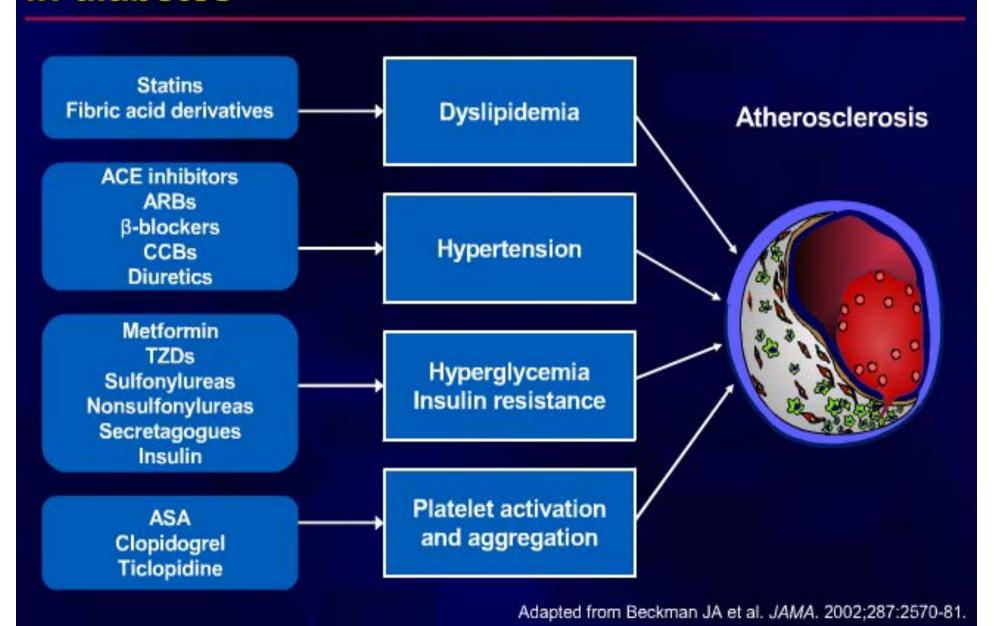
A wannabe WAG who





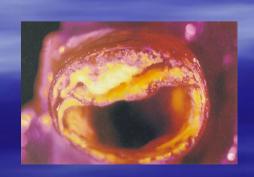


Beyond lifestyle: Aggressive medical therapy in diabetes













Case 3*

- 66 year old female
- Type 2 diabetes
 - No microvascular complications
- Diagnosed 4 years ago
 - HbA1c of 7.8% (62 mmol/mol)
 - Total cholesterol = 4.8
 - BP 140/80
- Questions?
 - Glycaemia, cholesterol



Case 4*

- 88 year old man
- Recently diagnosed diabetes
- HbA1c 8.5% (69 mmol/mol)
- Lives care home
- Co-morbidities
 - Vascular dementia
 - Congestive heart failure NYHA 2-3
 - Renal impairment eGFR 18
 - Previous MI and CVA

