



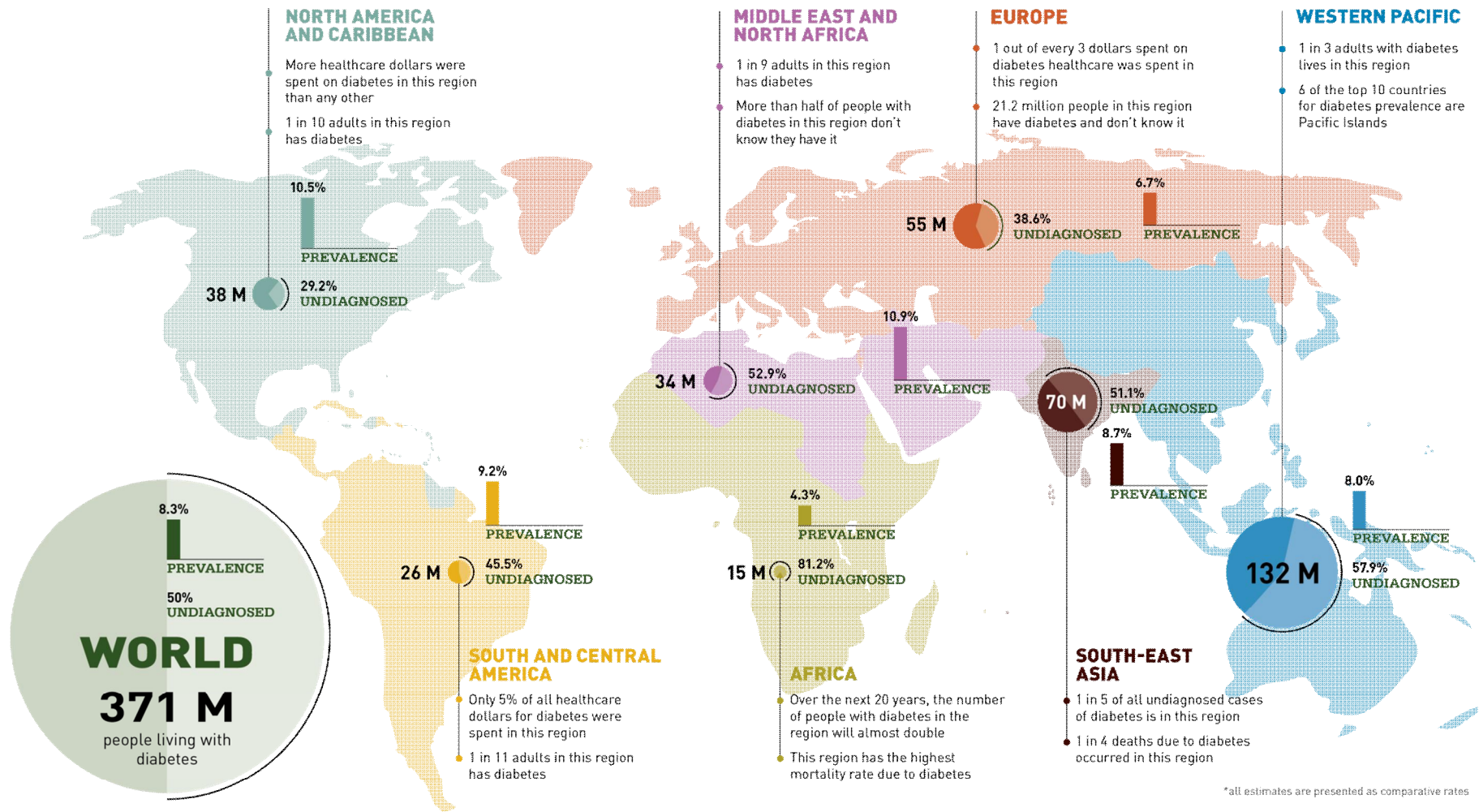
University  
of Glasgow

# Physical activity and the prevention and treatment of diabetes

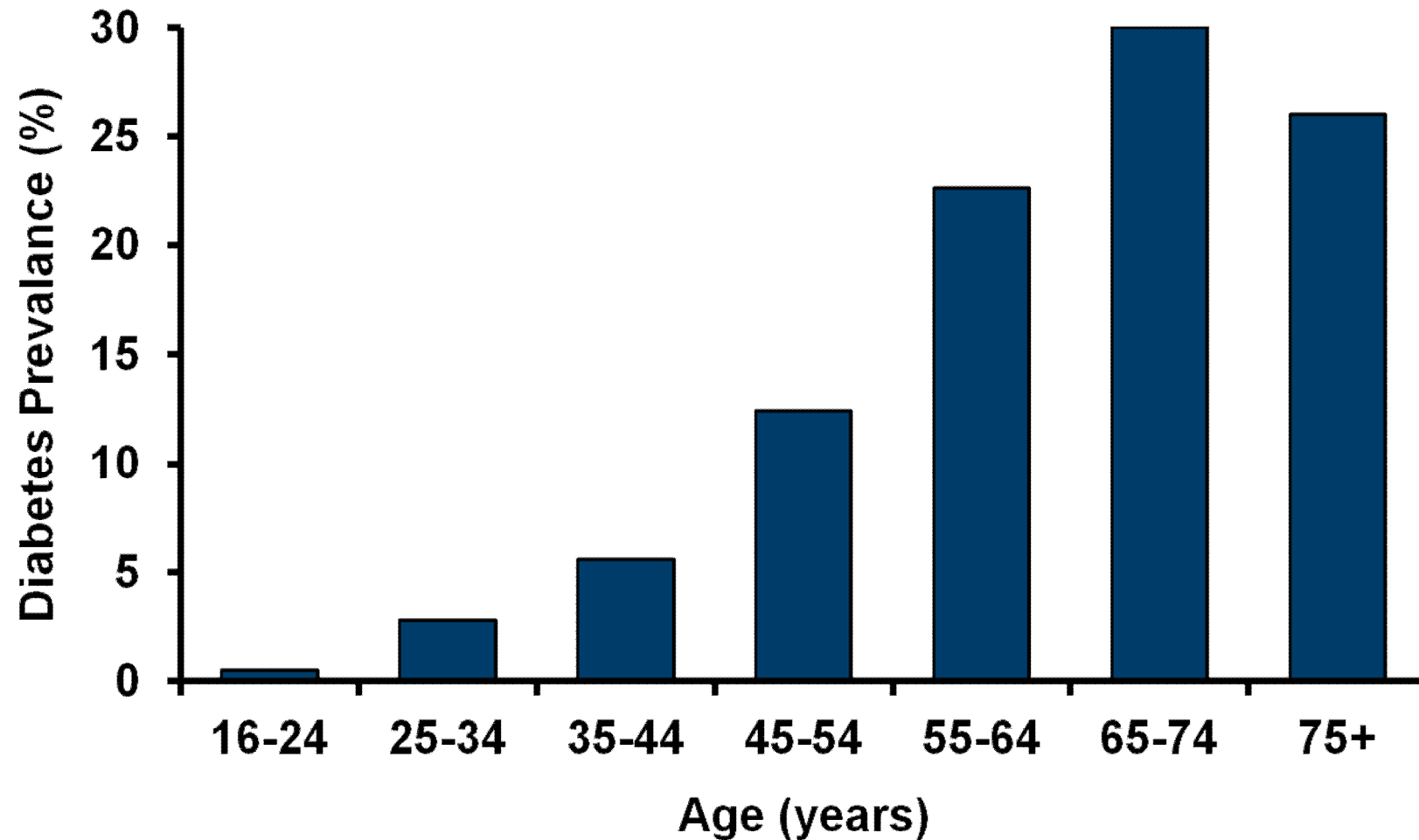
Dr Jason Gill

Institute of Cardiovascular and Medical Sciences  
University of Glasgow





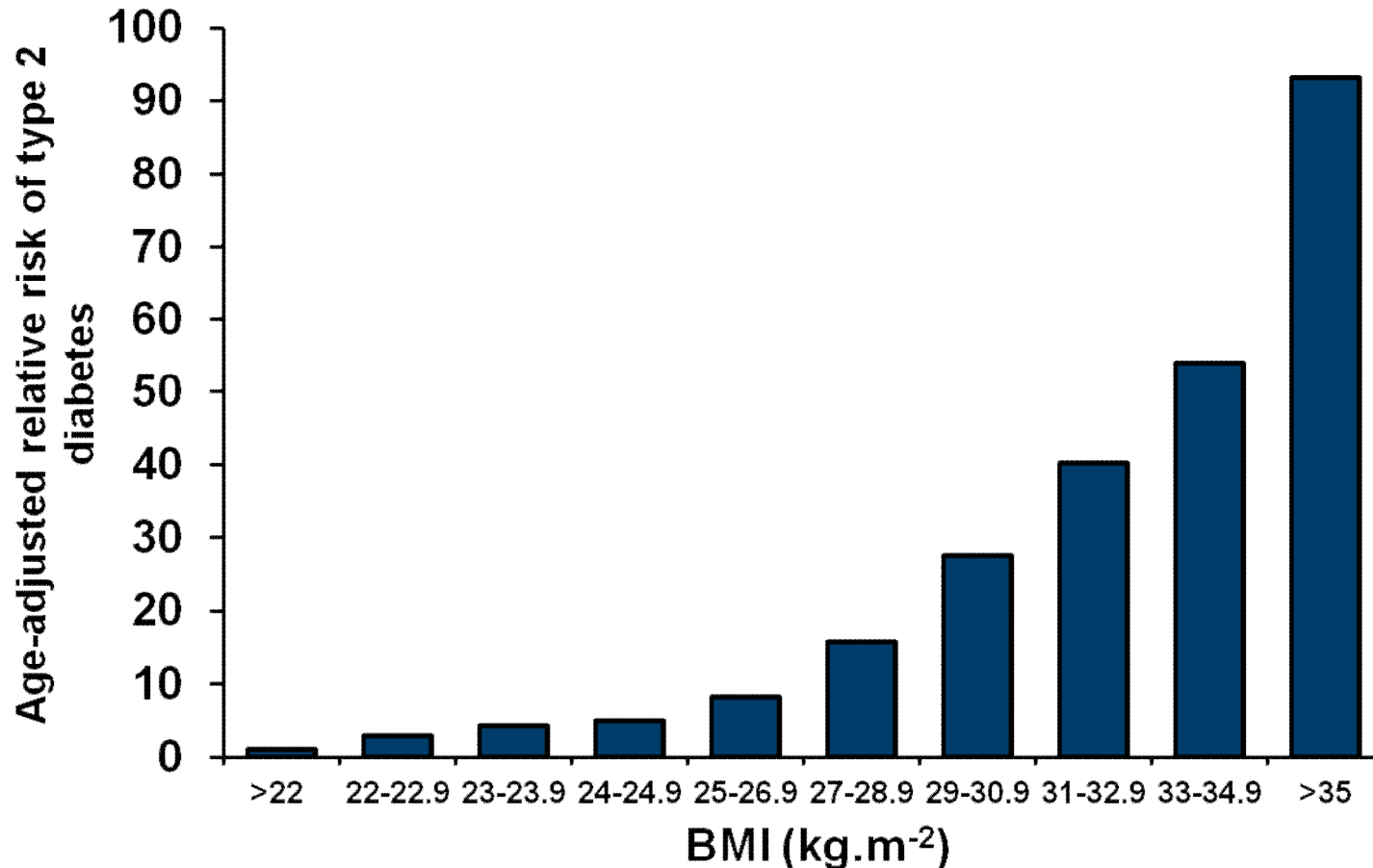
# Diabetes prevalence in Northern Ireland



Northern Ireland Health and Social Wellbeing Survey 2005/2006



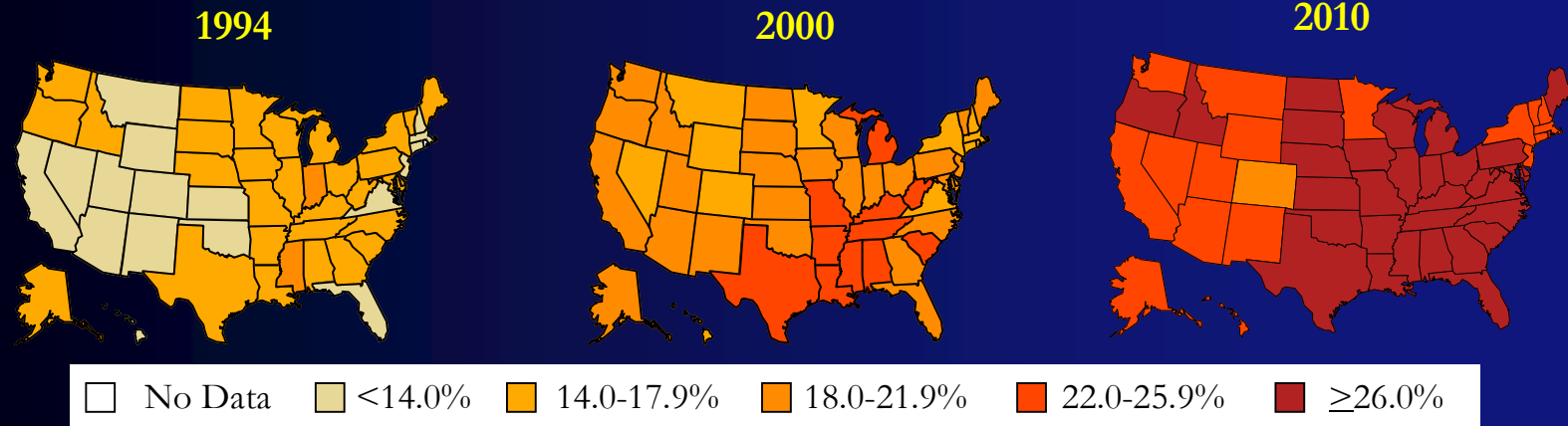
# Obesity and type 2 diabetes: The Nurses Health Study



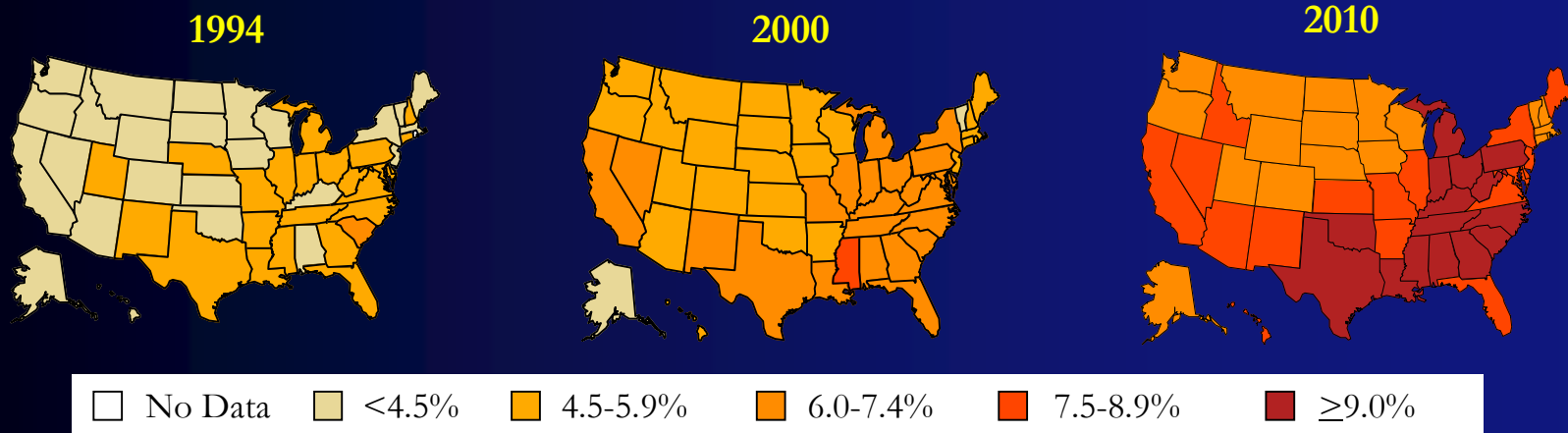
**BMI >35 kg.m<sup>-2</sup> associated with 93 times the risk of type 2 diabetes compared with BMI <22 kg.m<sup>-2</sup>**

# Age-Adjusted Prevalence of Obesity and Diagnosed Diabetes Among U.S. Adults Aged 18 Years or older

## Obesity (BMI $\geq 30$ kg/m<sup>2</sup>)



## Diabetes



CDC's Division of Diabetes Translation. National Diabetes Surveillance System available at <http://www.cdc.gov/diabetes/statistics>



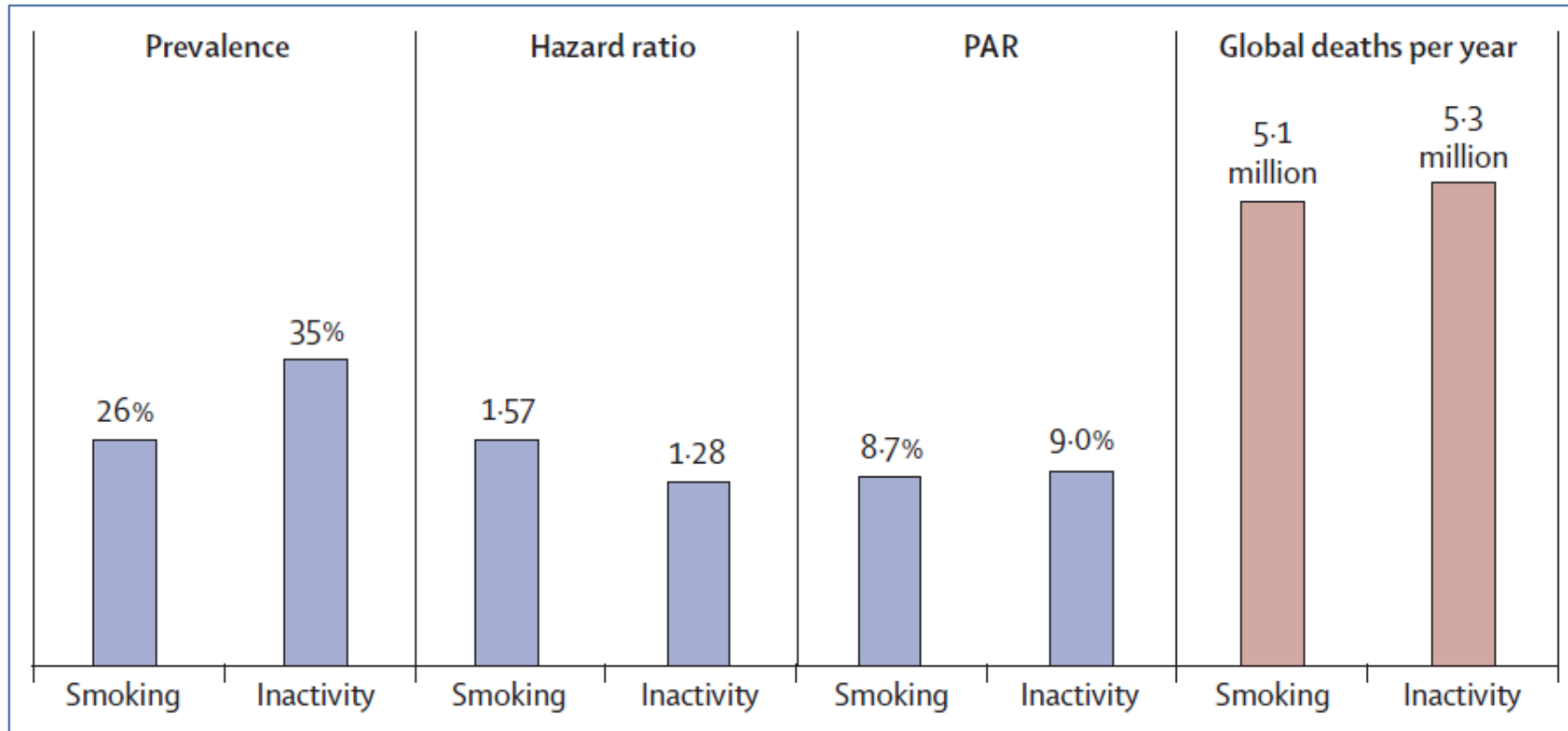


University  
of Glasgow

# **Epidemiology of physical activity, fitness & sedentary behaviour, and risk of diabetes**



# Global burden of physical inactivity



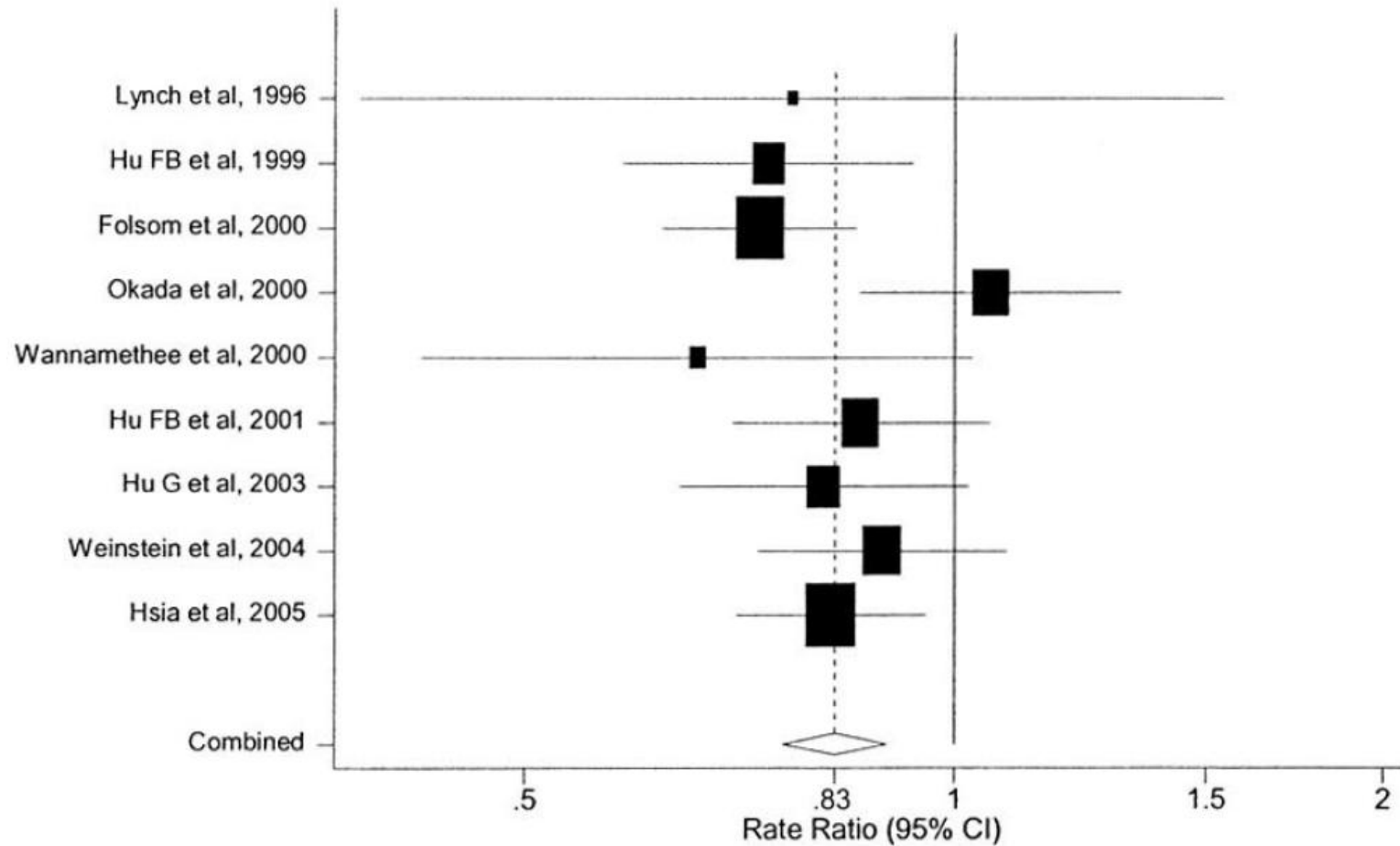
**Figure: Comparison of global burden between smoking and physical inactivity**

Prevalence of smoking, population attributable risk (PAR), and global deaths for smoking were obtained from WHO.<sup>7</sup> Hazard ratio for all-cause mortality of smoking was obtained from meta-analysis studies.<sup>8,9</sup> All inactivity data were obtained from Lee and colleagues.<sup>5</sup>



# Meta-analysis of effects of physical activity on type 2 diabetes risk

RR of type 2 diabetes with adjustment for BMI



Jeon et al, 2007, Diabetes Care, 30:744-752





# UK Physical Activity Guidelines

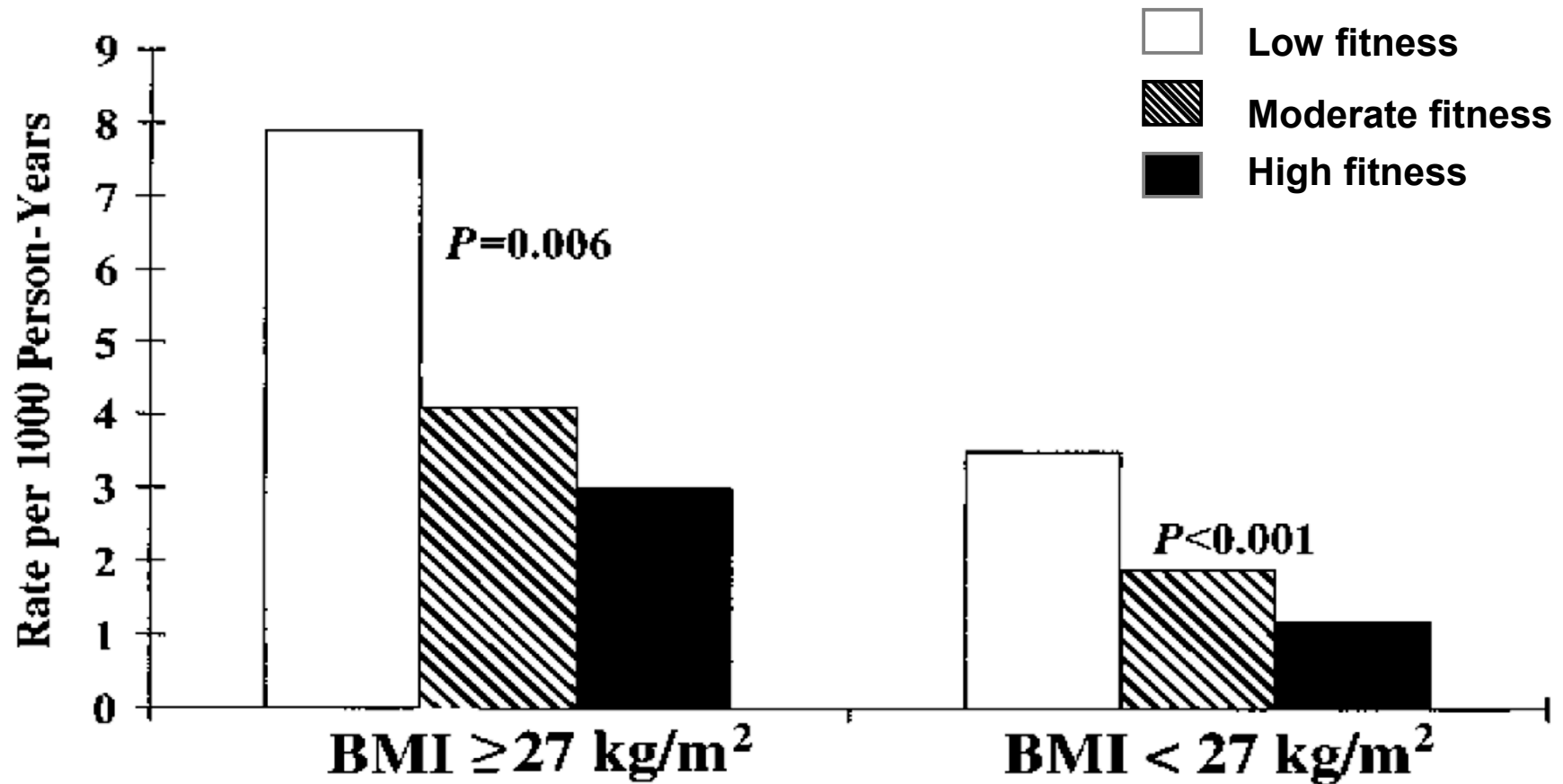
## **ADULTS (19–64 years)**

1. Adults should aim to be active daily. Over a week, activity should add up to at least 150 minutes (2½ hours) of moderate intensity activity in bouts of 10 minutes or more – one way to approach this is to do 30 minutes on at least 5 days a week.
2. Alternatively, comparable benefits can be achieved through 75 minutes of vigorous intensity activity spread across the week or a combination of moderate and vigorous intensity activity.
3. Adults should also undertake physical activity to improve muscle strength on at least two days a week.
4. All adults should minimise the amount of time spent being sedentary (sitting) for extended periods.

(Department of Health, 2011)



# Fitness, BMI and risk of type 2 diabetes: The Aerobics Center Longitudinal Study





University of Glasgow

MailOnline



NEWS

Home News U.S. Sport TV&Showbiz Femail Health Science&Tech Money Deb

News Home Arts Headlines Pictures Most read News Board

Building the plane that builds economies.

Watch the full story

## Sitting can be more dangerous than smoking, study shows

By DAILY MAIL REPORTER

Last updated at 11:20 PM on 9th June 2011

Comments (5) Add to My Stories Share

Like Confirm

Going to work could be more harmful than smoking, a new study shows. That is if your job involves sitting around for long periods of time.

Like Confirm

You like College News. Admin Page Insights Error

## Sitting may be worse for your health than you thought

Physical inactivity increases risk of life-threatening conditions

Madeline Rossman

## Sitting Too Long Raises Death Risk

Huffington Post Amanda Chan First Posted: 06/24/11 03:36 PM ET Updated: 06/27/11 05:08 PM ET

React Amazing Inspiring Funny Scary Hot Crazy Important Weird

Read more Healthy Living Health News, Death Risks Sitting, New Research, Sit Death, Sitting Death, Sitting Death Risk, Sitting Down Death, Healthy Living News

SHARE THIS STORY

Like

You like Sitting Too Long Raises Death

332

180

55

3

share

tweet

email

+1

Get up off your chair -- your life is at stake, a new study suggests.

Sitting more than six hours a day raises your risk of death, even if you work out, said the study, published in the American Journal of Epidemiology.

Even if they exercised, women who sat six or more hours a day were 37 percent more likely to die over a 13-year period than people who sat less than

06/15/2011 10:19 AM

ID: 89737

## TV Watching Increases Risk of Dying Rather Young

Several recently conducted studies found that watching TV can actually increase the risk of health problems and of dying at an earlier age.

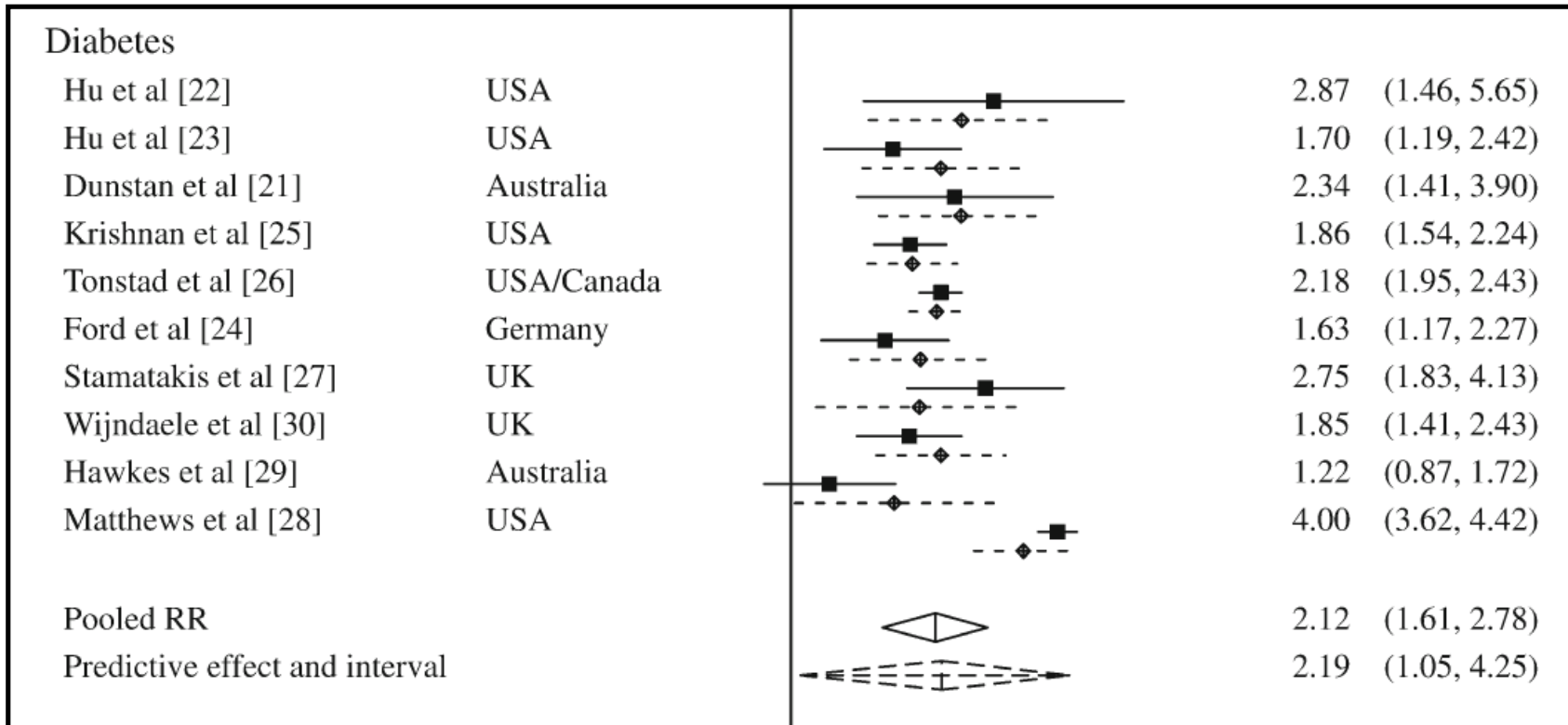
Two hours more in front of the TV on a typical day increase the risk of type 2 diabetes by 20 percent and the risk of developing a heart disease 15 percent up.

For every additional 3 hours watching the telly, the risk of dying from heart disease increases 13 percent.

The average American spends 5 hours daily sitting in front of the tube



# Meta-analysis of effects of sedentary time on diabetes risk



# Sitting time and glycaemia in South Asians (n = 1228)

	Two-hour glucose concentration	
	Difference in glucose (mmol/L)	<i>P</i>
Age (per year)	0.032 (0.016–0.049)	<0.0005
Sex (men compared with women)	0.000 (–0.360 to 0.360)	0.999
Waist (per cm)	0.057 (0.040–0.074)	<0.0005
Sitting time (per h/day)	0.097 (0.036–0.158)	0.002
Walking time (per h/day)	–0.040 (–0.312 to 0.232)	0.774
Moderate activity time (per h/day)	–0.083 (–0.410 to 0.244)	0.620
Vigorous activity time (per h/day)	–0.819 (–1.672 to 0.034)	0.060

Data are means (95% CI).

**Effect of each hour of sitting on glycaemia equivalent to 3 years in age or 2 cm in waist**

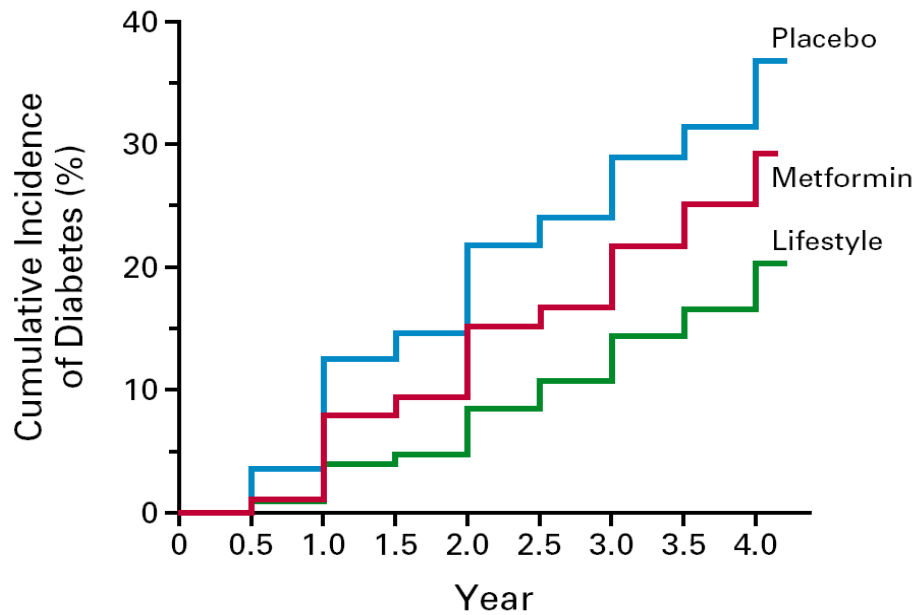


University  
of Glasgow

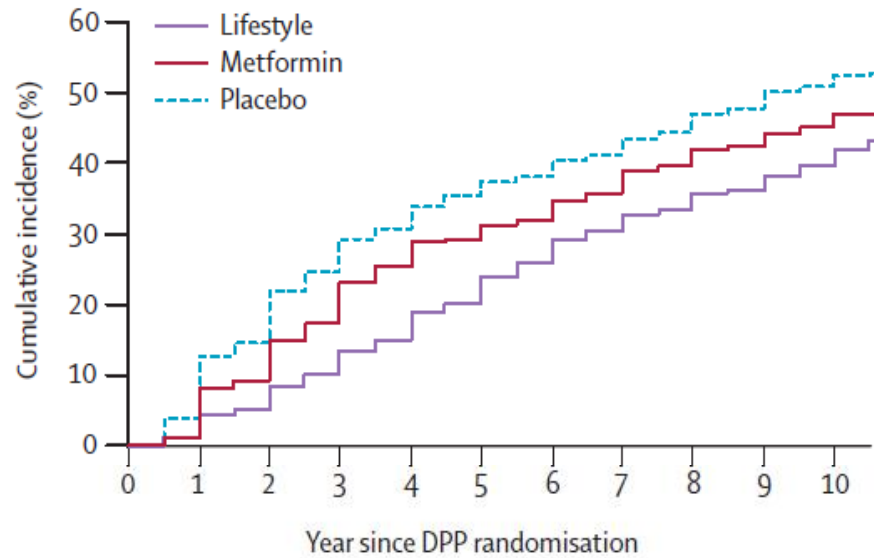
# **Lifestyle intervention trials for diabetes prevention**



# Long-term follow-up in US Diabetes Prevention Program



Knowler et al (2002) NEJM, 346:393-403

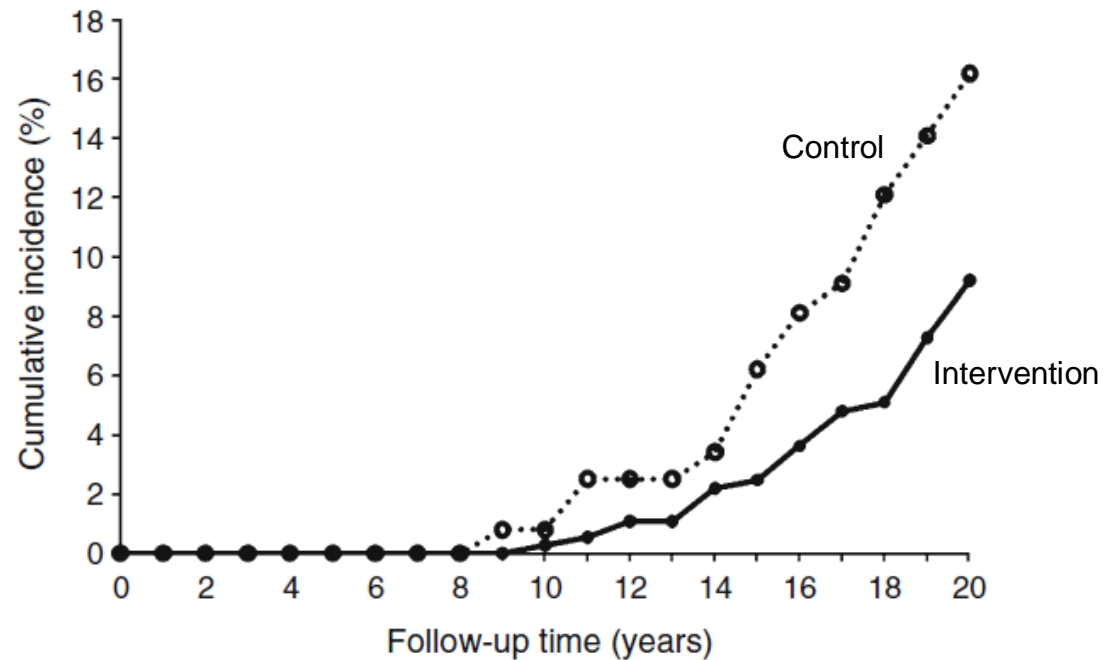


Knowler et al (2009) Lancet, 374:1677-86



# Long-term effects of lifestyle intervention in IGT on diabetes-related microvascular complications: The Da Qing Study

## Cumulative incidence of severe retinopathy



Number at risk:

Control	133	132	131	128	128	121	113	107	98	91	82
Intervention	407	407	402	393	387	376	364	355	339	316	290





University  
of Glasgow

# **Lifestyle intervention trials for diabetes treatment**

# Aerobic training, resistance training or both on glycaemic control in type 2 diabetes

Variable	Mean (SD) Value			Difference in Change from Baseline to 6 Months (95% CI)	P Value
	Baseline	3 mo	6 mo		
<b>Hemoglobin A<sub>1c</sub> [patients], % [n]†</b>					
Combined exercise group	7.46 (1.48) [64]	6.99 (1.56) [60]	6.56 (1.55) [58]	–	–
Aerobic training group	7.41 (1.50) [60]	7.00 (1.59) [58]	6.98 (1.50) [49]	–	–
Resistance training group	7.48 (1.47) [64]	7.35 (1.57) [62]	7.18 (1.52) [56]	–	–
Control group	7.44 (1.38) [63]	7.33 (1.49) [62]	7.51 (1.47) [59]	–	–
<b>Intergroup comparisons</b>					
Aerobic training vs. control	–	–	–	–0.51 (–0.87 to –0.14)	0.007
Resistance training vs. control	–	–	–	–0.38 (–0.72 to –0.22)	0.038
Combined exercise vs. aerobic training	–	–	–	–0.46 (–0.83 to –0.09)	0.014
Combined exercise vs. resistance training	–	–	–	–0.59 (–0.95 to –0.23)	0.001

“ **Control group** (n = 63) . no exercise

“ **Aerobic training group** (n = 60) . building to 45 mins cycling/treadmill exercise @ 75% max HR 3 x per week

“ **Resistance training group** (n = 64) . 2-3 sets of 7 resistance exercises 3 x per week

“ **Combined training group** (n = 64) . the full aerobic AND the full resistance intervention



University  
of Glasgow

# ACSM/ADA recommendations for Physical activity in treatment of type 2 diabetes

- Persons with type 2 diabetes should undertake at least 150 min/week of moderate to vigorous aerobic exercise spread out during at least 3 days during the week, with no more than 2 consecutive days between bouts of aerobic activity.
- In addition to aerobic training, persons with type 2 diabetes should undertake moderate to vigorous resistance training at least 2–3 days/week.
- Supervised and combined aerobic and resistance training may confer additional health benefits, although milder forms of PA (such as yoga) have shown mixed results. Persons with type 2 diabetes are encouraged to increase their total daily unstructured PA. Flexibility training may be included but should not be undertaken in place of other recommended types of PA.



University  
of Glasgow

# Long-term effects of lifestyle intervention on weight & CVD risk in type 2 diabetes: the Look AHEAD trial

- “ 5145 adults aged 45-76 years with type 2 diabetes randomised to Intensive Lifestyle Intervention (n = 2570) or Diabetes Support and Education (n = 2575).
  
- “ **Intensive Lifestyle Intervention (ILI)**
  - “ dietary goal to induce 7% weight loss and 175 minutes moderate intensity physical activity per week
  - “ Supported by group/individual support sessions weekly for first 6 months; 3 x per months from 6-12 months; once per month (plus 1 telephone/email contact per month) for years 2-4.
  - “ Behaviour change strategies taught (self-monitoring, goal setting, problem solving), weigh-ins at each session,
  
- “ **Diabetes Support and Education (DSE)**
  - “ 3 group sessions per year focused on diet, physical activity or social support.
  - “ No behaviour change strategies taught, no weigh ins

Wing et al (2010) Arch Intern Med, 170:1566-1575



# Long-term effects of lifestyle intervention on weight & CVD risk in type 2 diabetes: the Look AHEAD trial

**Table 1. Mean Changes in Weight, Fitness, and CVD Risk Factors in ILI and DSE Groups and the Difference Between Groups Averaged Across 4 Years**

Measure	Groups, Mean Change (95% CI)		Between-Group Mean Difference (95% CI)	P Value of Difference <sup>a</sup>
	DSE	ILI		
Weight, % initial weight	-0.88 (-1.12 to -0.64)	-6.15 (-6.39 to -5.91)	-5.27 (-5.61 to -4.93)	<.001
Fitness, % METS	1.96 (1.07 to 2.85)	12.74 (11.87 to 13.62)	10.78 (9.53 to 12.03)	<.001
HbA <sub>1c</sub> level <sup>b</sup>	-0.09 (-0.13 to -0.06)	-0.36 (-0.40 to -0.33)	-0.27 (-0.32 to -0.22)	<.001
SBP, mm Hg <sup>b</sup>	-2.97 (-3.44 to -2.49)	-5.33 (-5.80 to -4.86)	-2.36 (-3.03 to -1.70)	<.001
DBP, mm Hg <sup>b</sup>	-2.48 (-2.73 to -2.24)	-2.92 (-3.16 to -2.68)	-0.43 (-0.77 to -0.10)	.01
HDL-C level, mg/dL <sup>b</sup>	1.97 (1.73 to 2.22)	3.67 (3.43 to 3.91)	1.70 (1.35 to 2.04)	<.001
Triglycerides level, mg/dL <sup>b</sup>	-19.75 (-22.11 to -17.39)	-25.56 (-27.91 to -23.21)	-5.81 (-9.14 to -2.48)	<.001
LDL-C level, mg/dL				
Without adjustment for medication use	-12.84 (-13.67 to -12.00)	-11.27 (-12.10 to -10.44)	1.57 (0.39 to 2.74)	.009
Adjusted for medication use	-9.22 (-10.04 to -8.39)	-8.75 (-9.56 to -7.94)	0.47 (-0.67 to 1.60)	.42



# Long-term effects of lifestyle intervention on weight & CVD risk in type 2 diabetes: the Look AHEAD trial

**Table 2. Proportion of DSE and ILI Participants Who Initiated or Maintained Use of Medication for Diabetes, Hypertension, or Lowering Lipid Levels**

	Use of Medication at Follow-up by Group					
	No Use at Baseline		P Value	Continued Use From Baseline		P Value
	DSE	ILI		DSE	ILI	
<b>Diabetes medication</b>						
No. at baseline <sup>a</sup>	348	354		2208	2202	
Follow-up year, %						
1	33.1	10.4	<.001	97.5	89.4	<.001
2	46.3	17.4	<.001	96.3	88.2	<.001
3	58.6	27.3	<.001	95.4	89.2	<.001
4	66.8	41.8	<.001	96.0	90.6	<.001
<b>Insulin</b>						
No. at baseline <sup>a</sup>	2167	2190		408	380	
Follow-up year, %						
1	3.7	1.7	<.001	91.6	80.6	<.001
2	6.7	3.1	<.001	86.4	76.1	<.001
3	8.9	4.3	<.001	86.3	77.7	.004
4	11.5	6.9	<.001	88.0	77.4	<.001

**Table 2. Proportion of DSE and ILI Participants Who Initiated or Maintained Use of Medication for Diabetes, Hypertension, or Lowering Lipid Levels**

	Use of Medication at Follow-up by Group					
	No Use at Baseline		P Value	Continued Use From Baseline		P Value
	DSE	ILI		DSE	ILI	
<b>Hypertension medication</b>						
No. at baseline <sup>a</sup>	684	661		1872	1895	
Follow-up year, %						
1	21.9	16.4	.01	89.9	81.3	<.001
2	31.9	24.7	.005	90.4	81.0	<.001
3	40.3	33.3	.01	91.2	82.9	<.001
4	47.2	43.0	.15	92.7	85.0	<.001
<b>Lipid-lowering medication</b>						
No. at baseline <sup>a</sup>	1313	1310		1243	1246	
Follow-up year, %						
1	25.3	17.6	<.001	92.3	89.6	.03
2	39.8	29.0	<.001	90.7	89.1	.22
3	47.4	38.5	<.001	88.6	90.0	.26
4	53.2	47.2	.004	90.9	90.4	.72



University  
of Glasgow

# Long-term effects of lifestyle intervention on weight & CVD risk in type 2 diabetes: the Look AHEAD trial



U.S. Department of Health and Human Services

**NIH News**

National Institutes of Health

For Immediate Release

Friday, October 19, 2012

Contact:

[Amy Reiter](#)

301-496-3583

**Weight loss does not lower heart disease risk from type 2 diabetes  
Intervention stopped early in NIH-funded study of weight loss in  
overweight and obese adults with type 2 diabetes after finding no  
harm, but no cardiovascular benefits**

An intensive diet and exercise program resulting in weight loss does not reduce cardiovascular events such as heart attack and stroke in people with longstanding type 2 diabetes, according to a study supported by the National Institutes of Health.

## Summary (part 1)

- “ Epidemiological evidence shows strong associations between low physical activity, low fitness and high sedentary time and risk of diabetes.
- “ RCTs indicate that lifestyle intervention is effective over the long-term of preventing diabetes and diabetes complications
- “ Lifestyle intervention is effective at improving glycaemic control, weight loss and reducing medication use, reducing sleep apnoea and improving mobility in type 2 diabetes
- “ Effects of lifestyle intervention is effective at reducing CVD events in highly-treated patients with type 2 diabetes not established





University  
of Glasgow

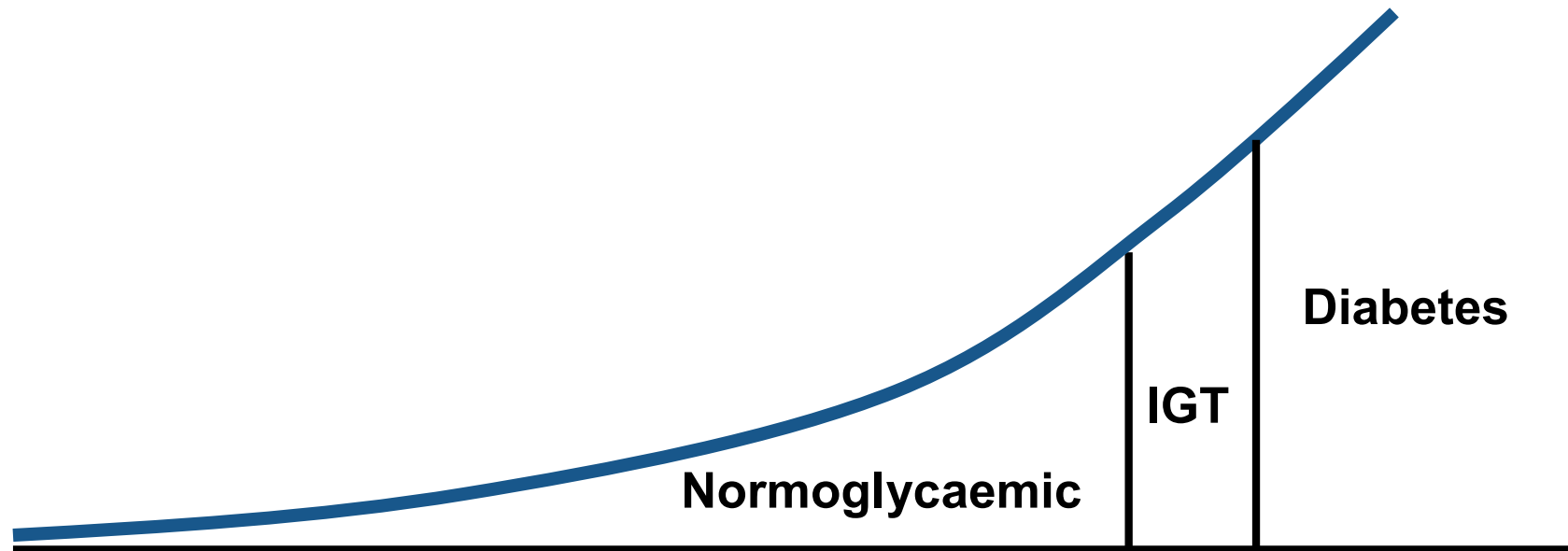
**Should we intervene earlier for  
diabetes prevention?**

**And if so, who should we target  
for early intervention?**



University  
of Glasgow

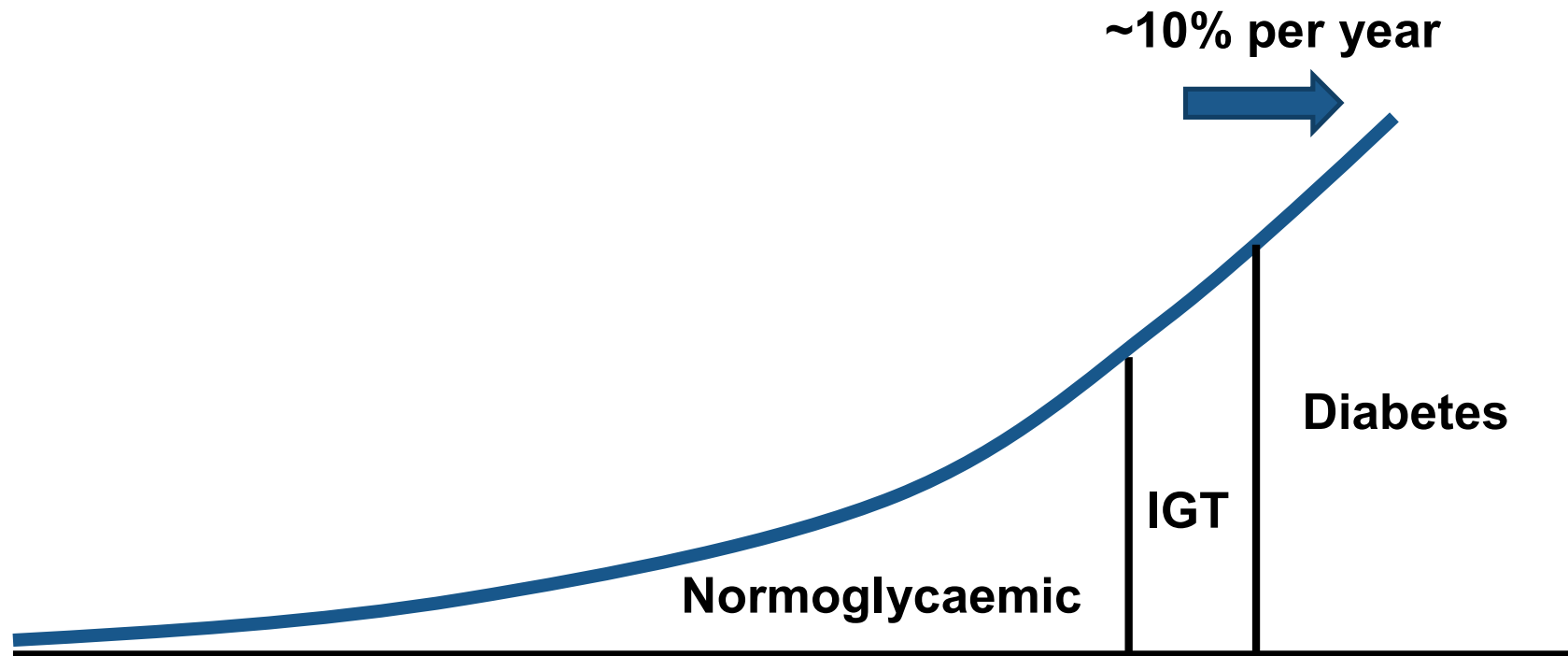
# The diabetes risk continuum





University  
of Glasgow

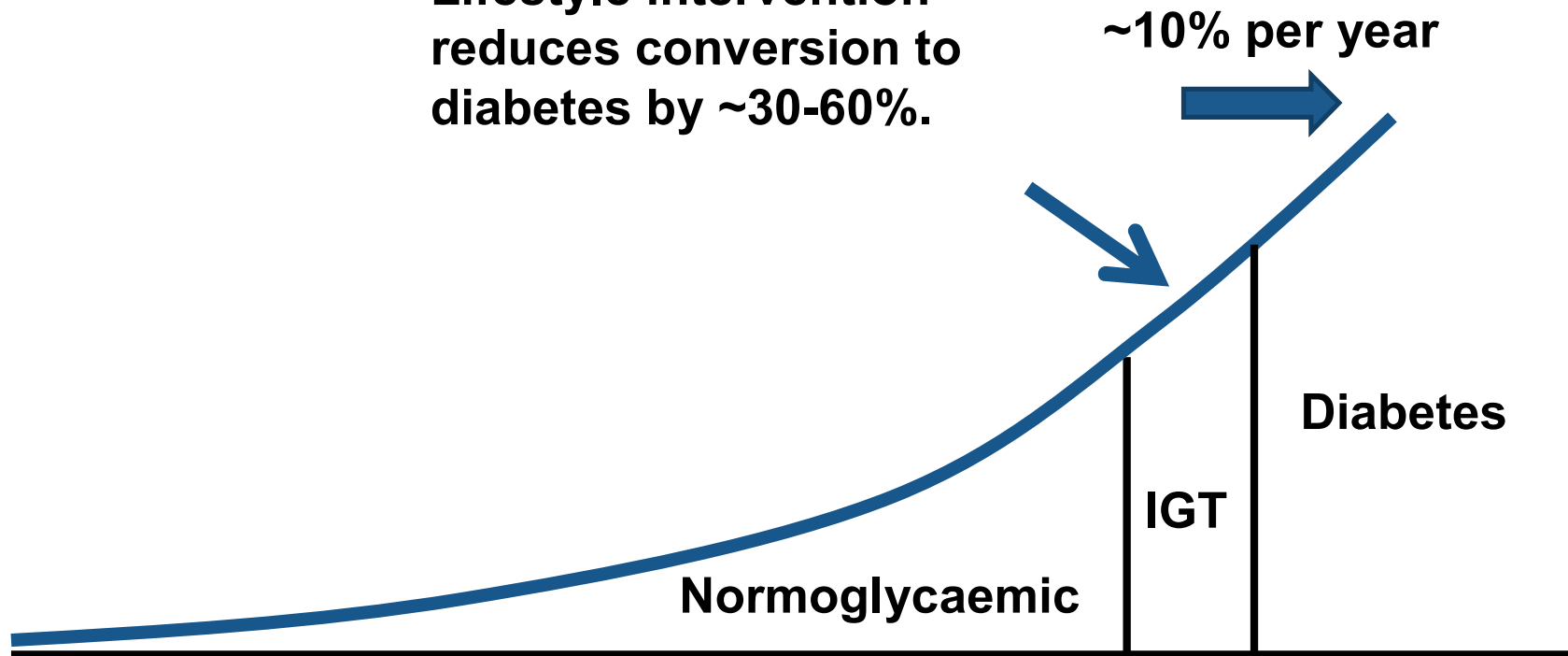
# The diabetes risk continuum





# The diabetes risk continuum

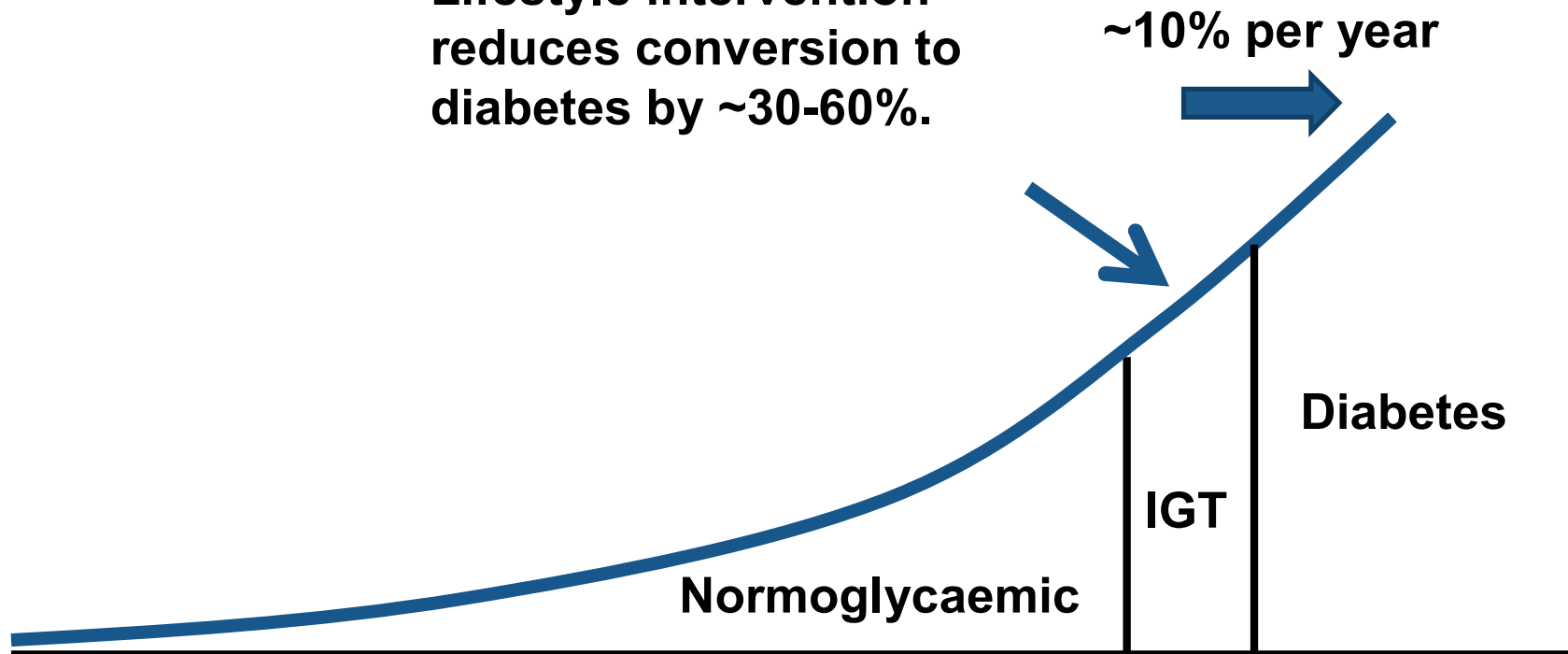
Lifestyle intervention  
reduces conversion to  
diabetes by ~30-60%.





# The diabetes risk continuum

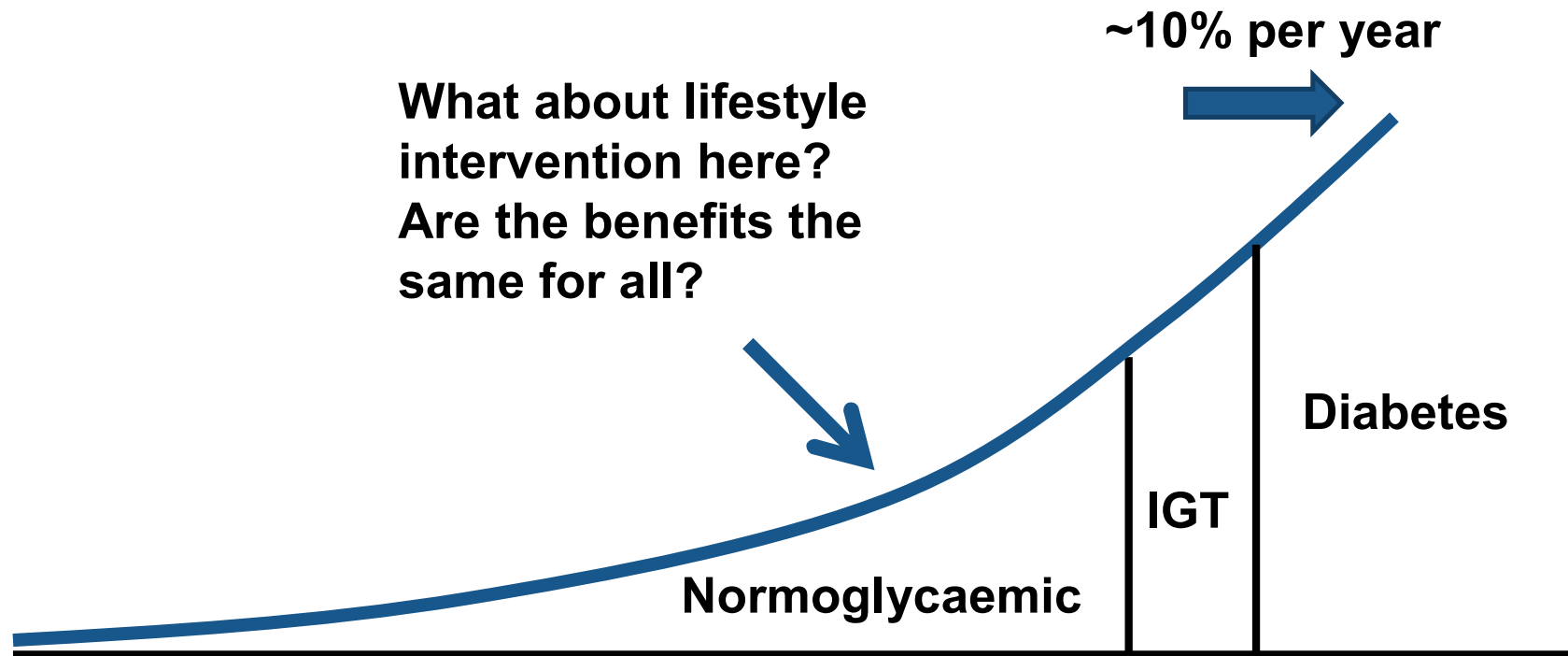
Lifestyle intervention  
reduces conversion to  
diabetes by ~30-60%.

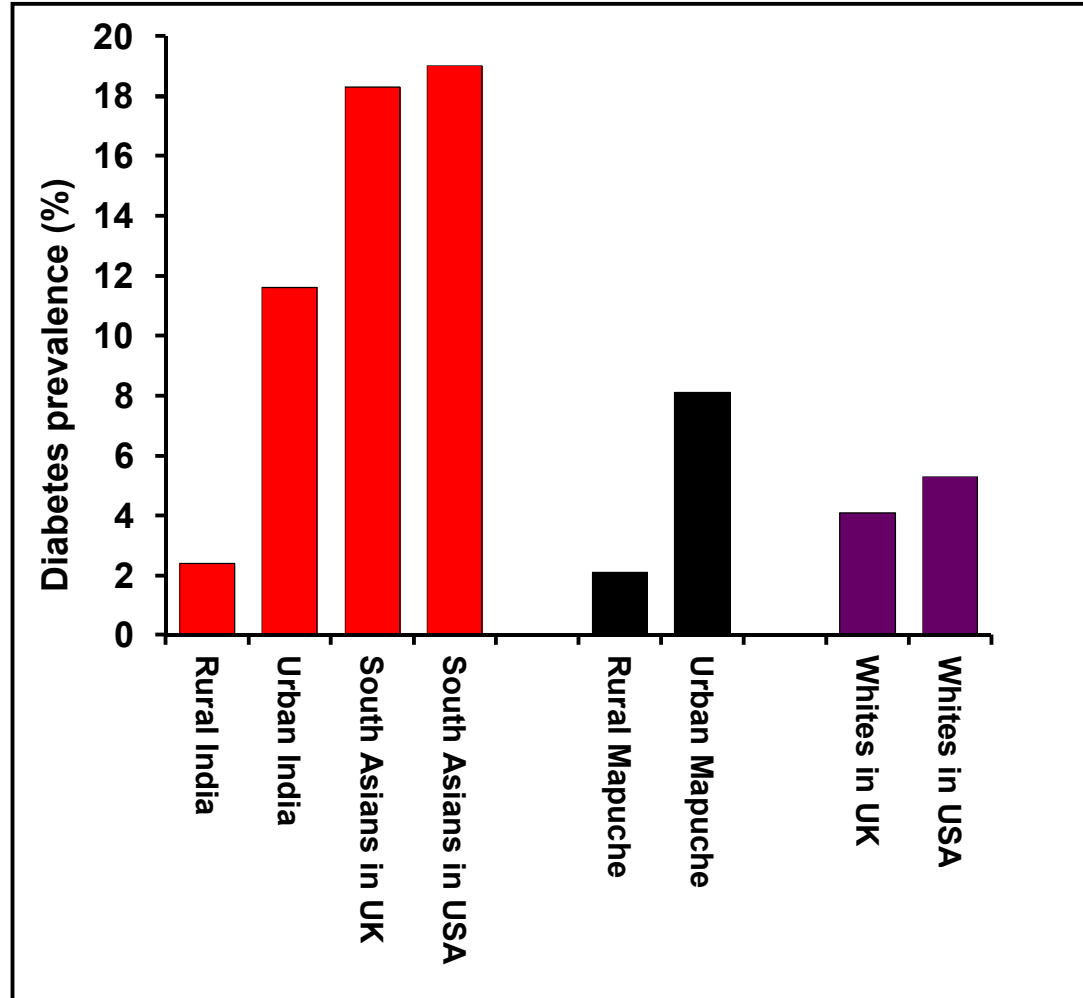


**BUT – even with lifestyle intervention many  
with IGT still develop diabetes.  
Should we aim to intervene earlier?**



# The diabetes risk continuum

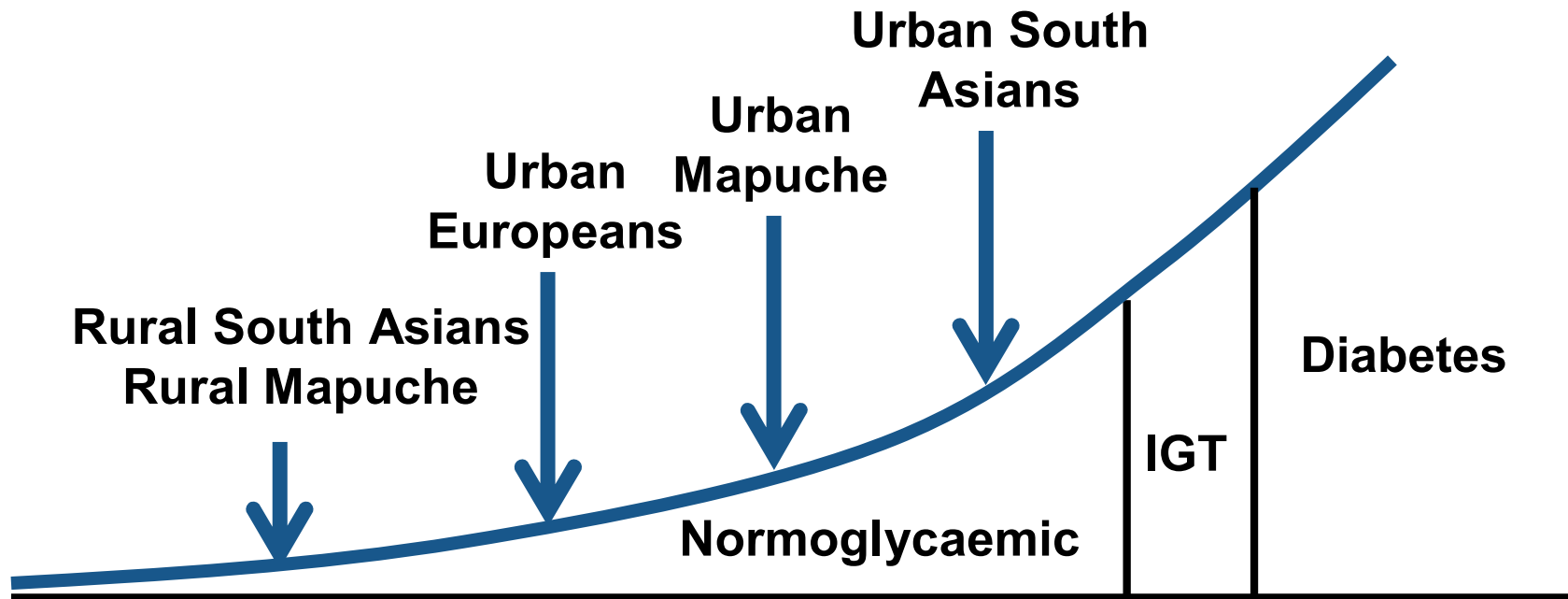




Hall et al (2008) *Future Lipidol* 3: 411-424; Carrasco et al (2004) *Chilean Med J.* 132: 1189-1197.



# The diabetes risk continuum







# Ethnicity in Chile

Distribution of Chilean Population, Census 2002.



Total population: ~16 million

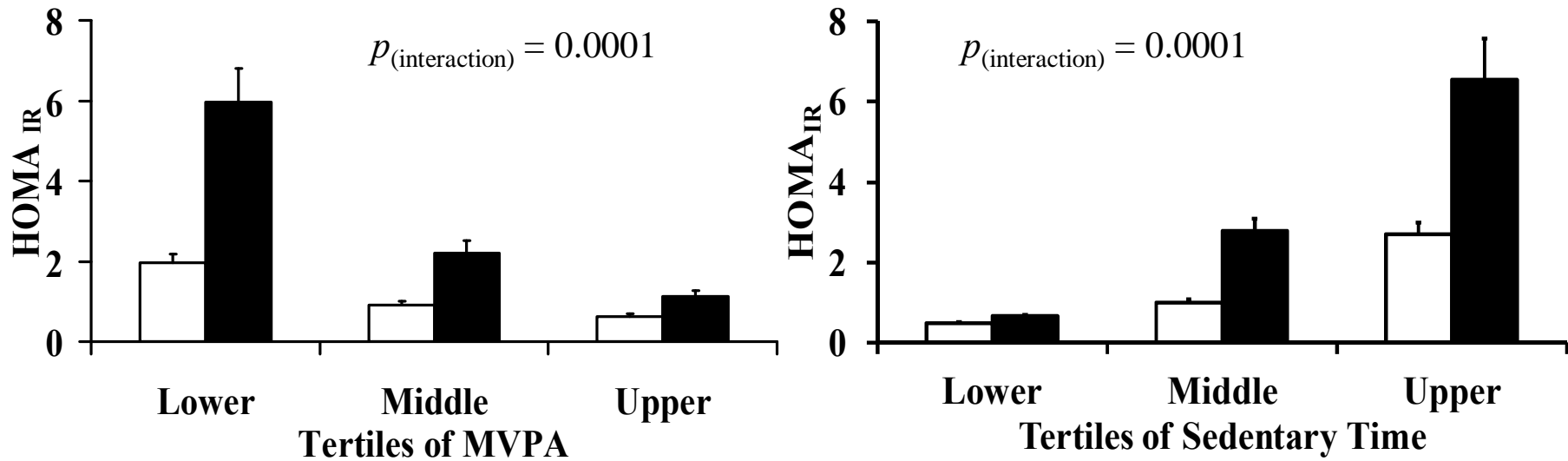
Total Aboriginal Population: 730,000

Mapuche Populations: >600,000





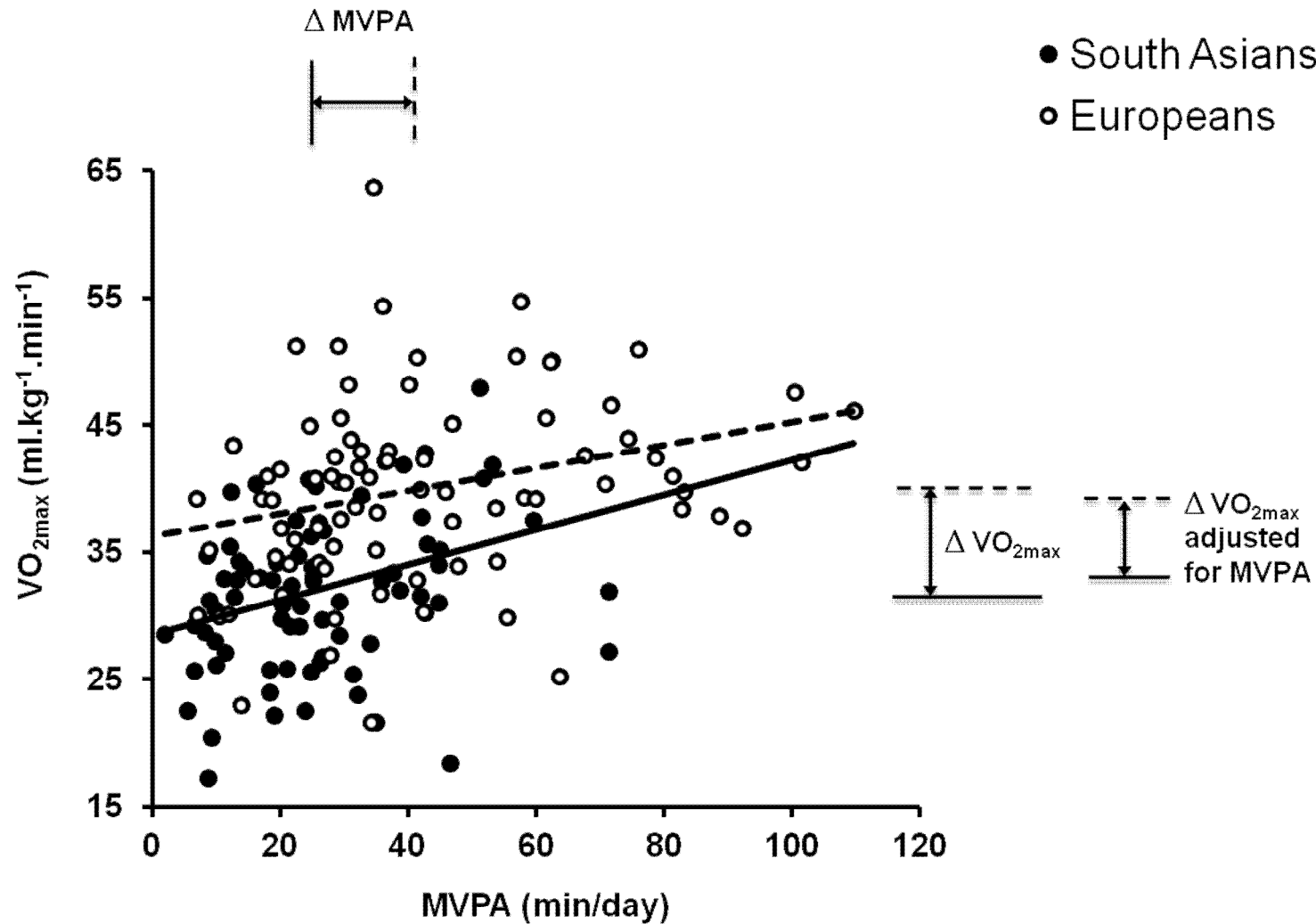
# Effects of physical activity and sedentary behaviour on insulin resistance in European and Mapuche Chilean men





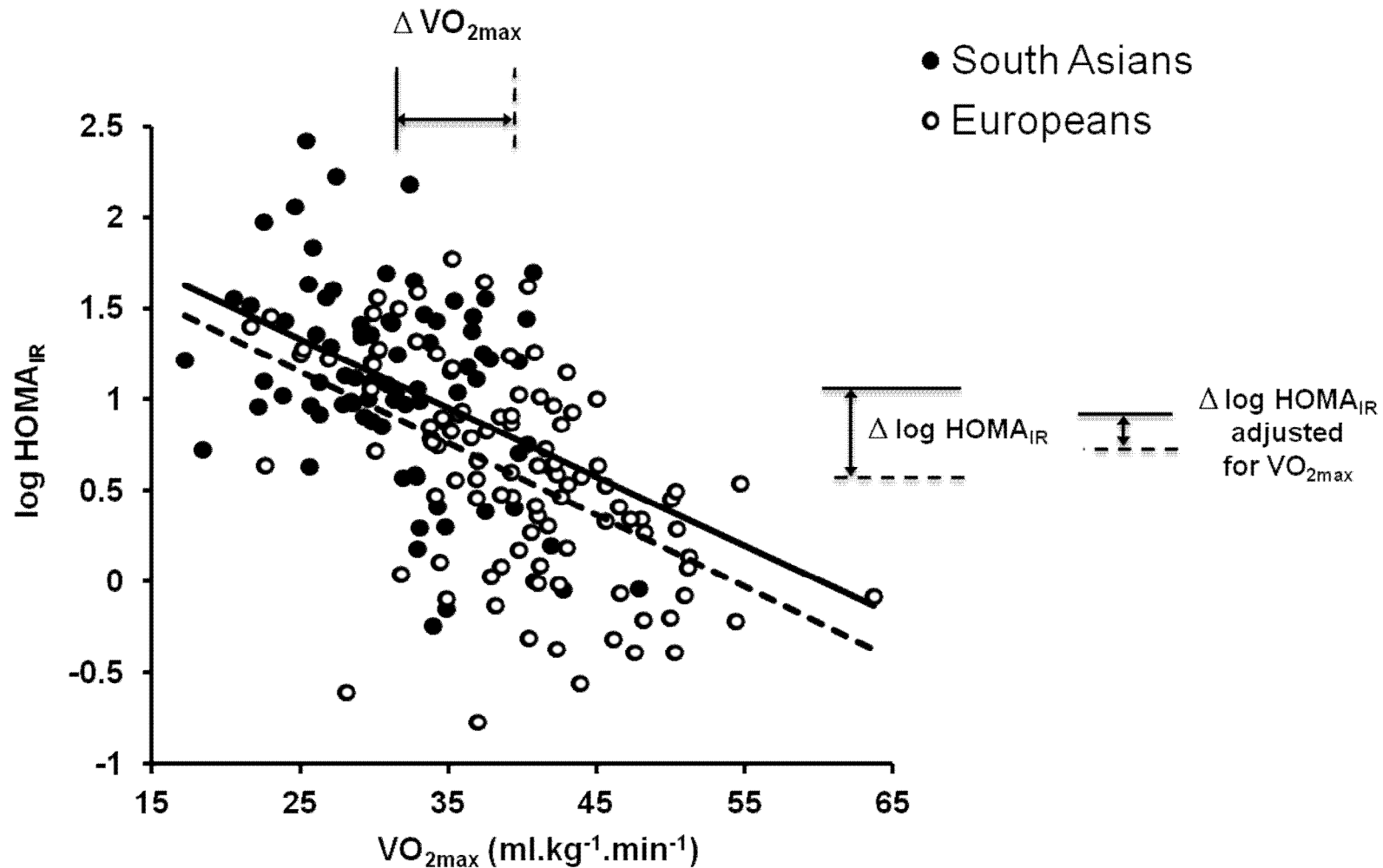


# Relationship between fitness and HOMA in European and South Asian men





# Relationship between fitness and physical activity in European and South Asian men





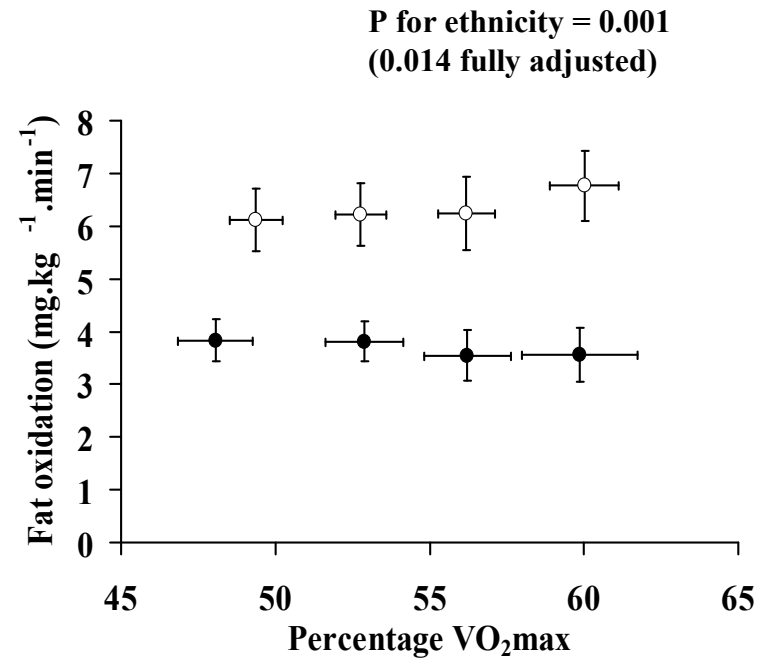
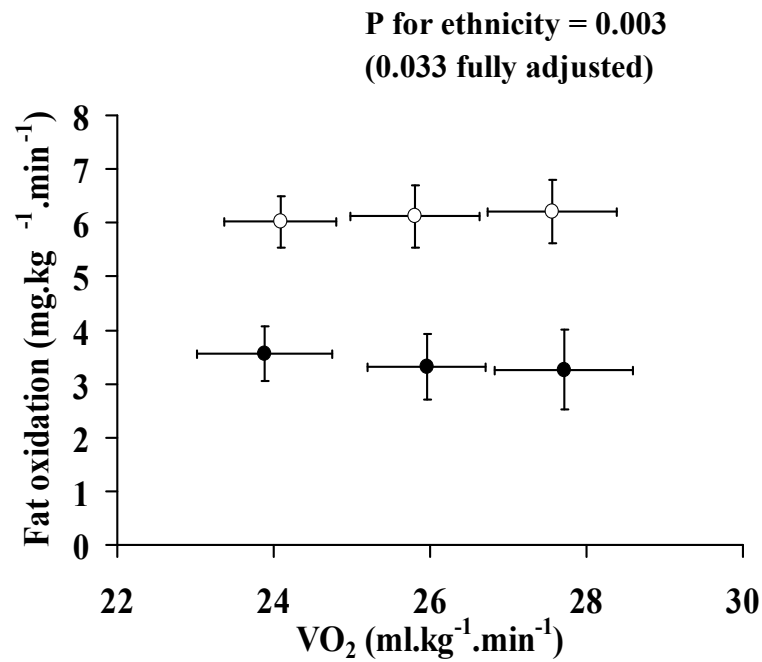
University  
of Glasgow

# $VO_{2max}$ , physical activity and insulin resistance in South Asians

- “ Low  $VO_{2max}$  explains over two-thirds of South Asians' greater insulin resistance
- “ This is not explained by lower levels of physical activity
- “ South Asians need to undertake higher amounts of physical activity for similar  $VO_{2max}$  (and  $HOMA_{IR}$ )

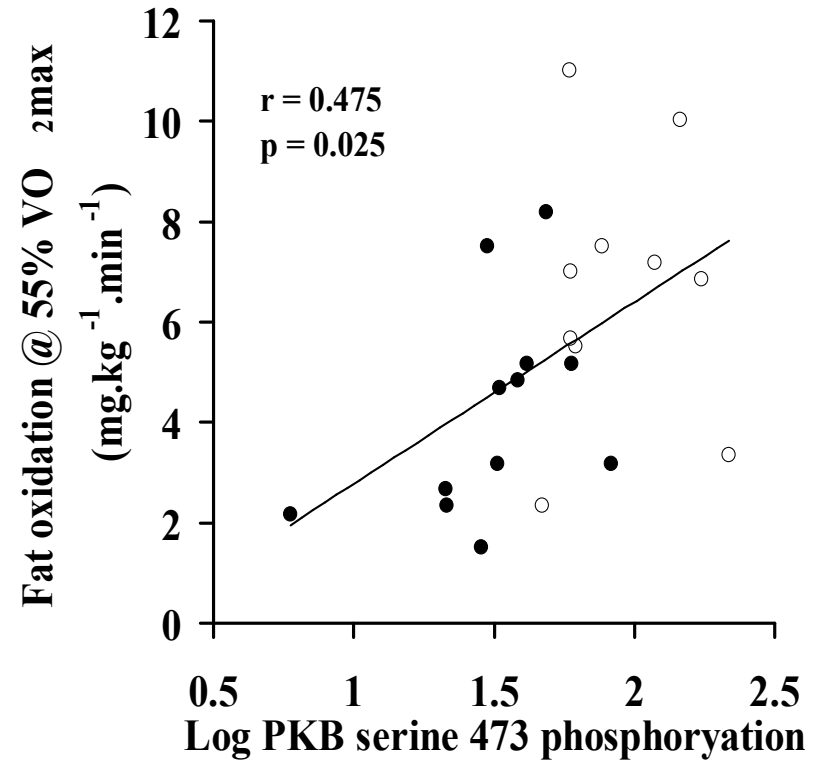
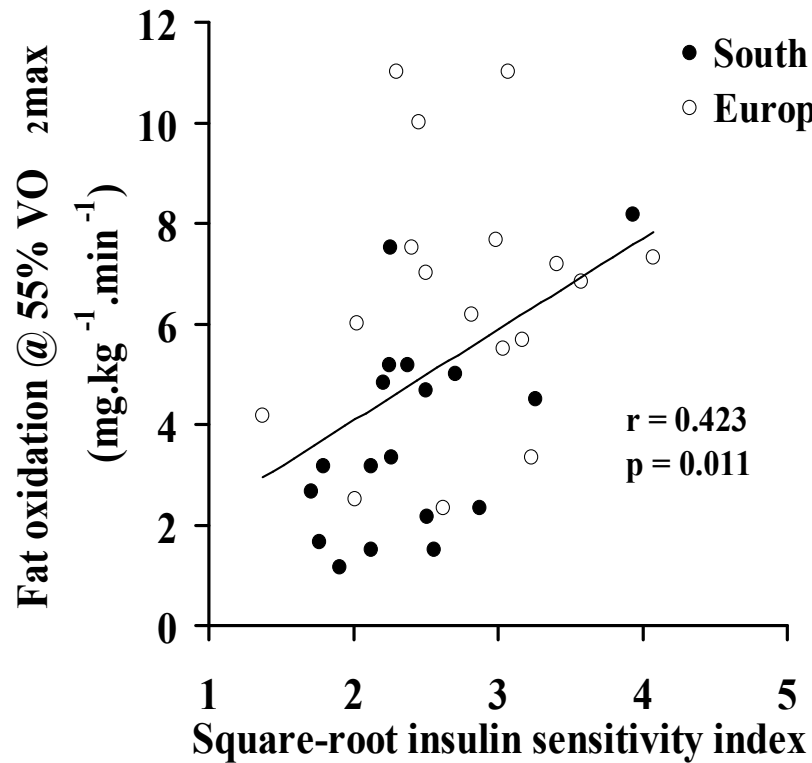


# Fat oxidation during submaximal exercise in South Asian and European men





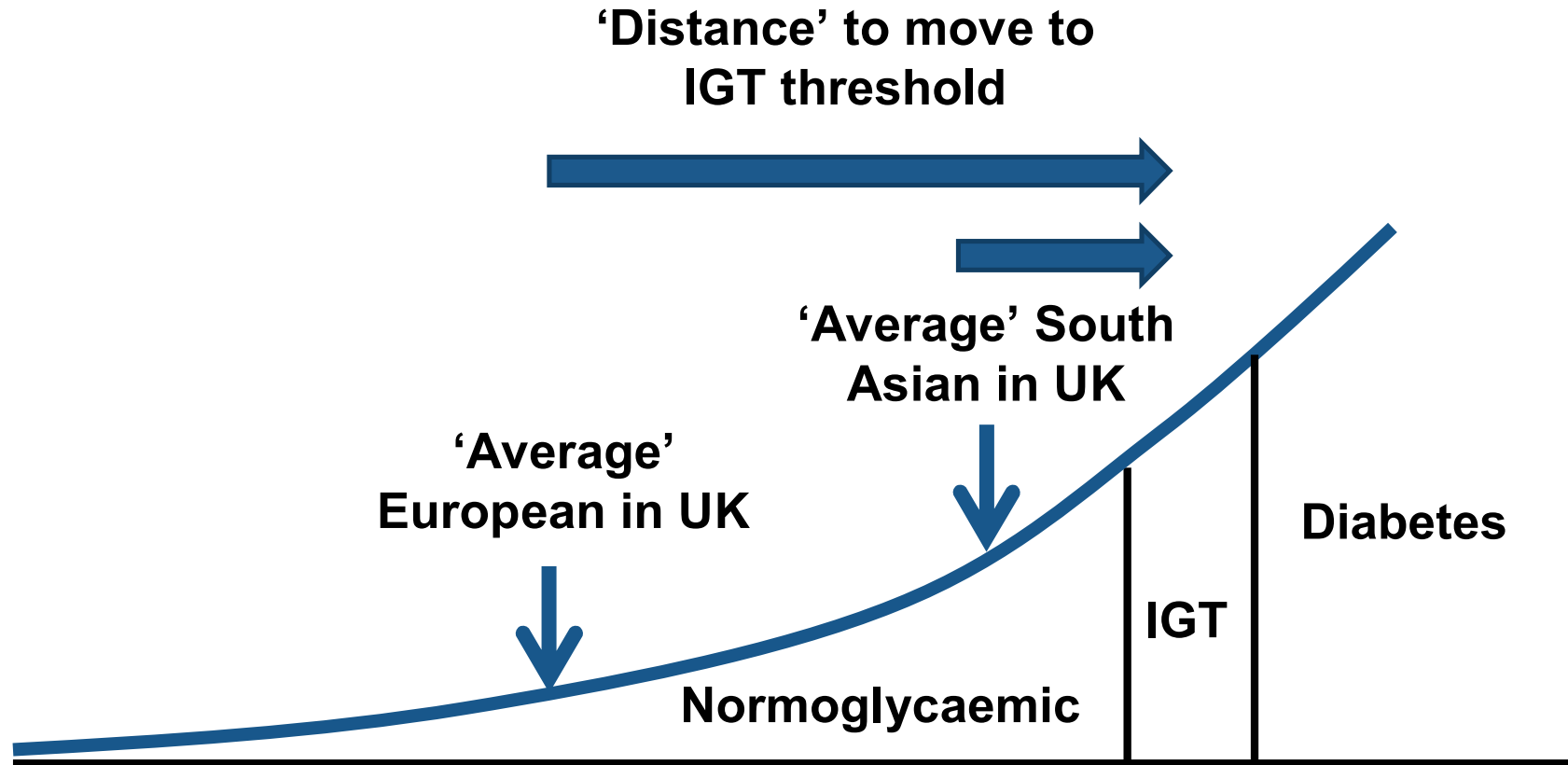
# Substrate utilisation during exercise and insulin sensitivity in South Asian and European men





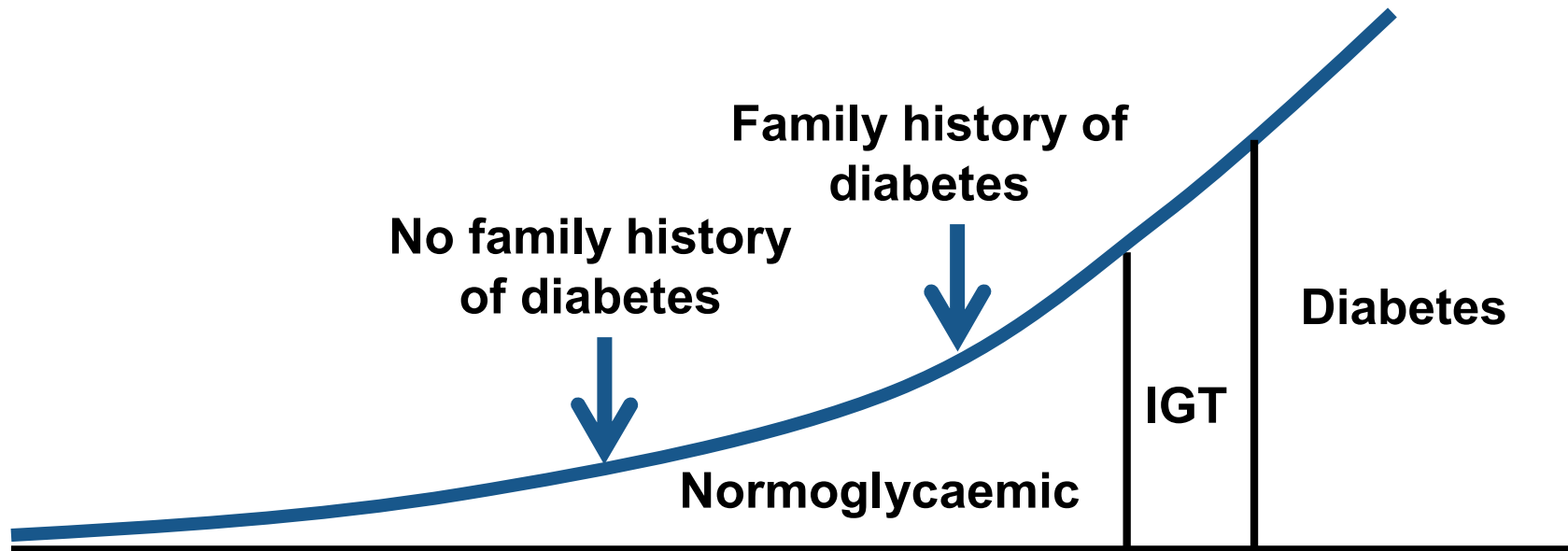


# The diabetes risk continuum





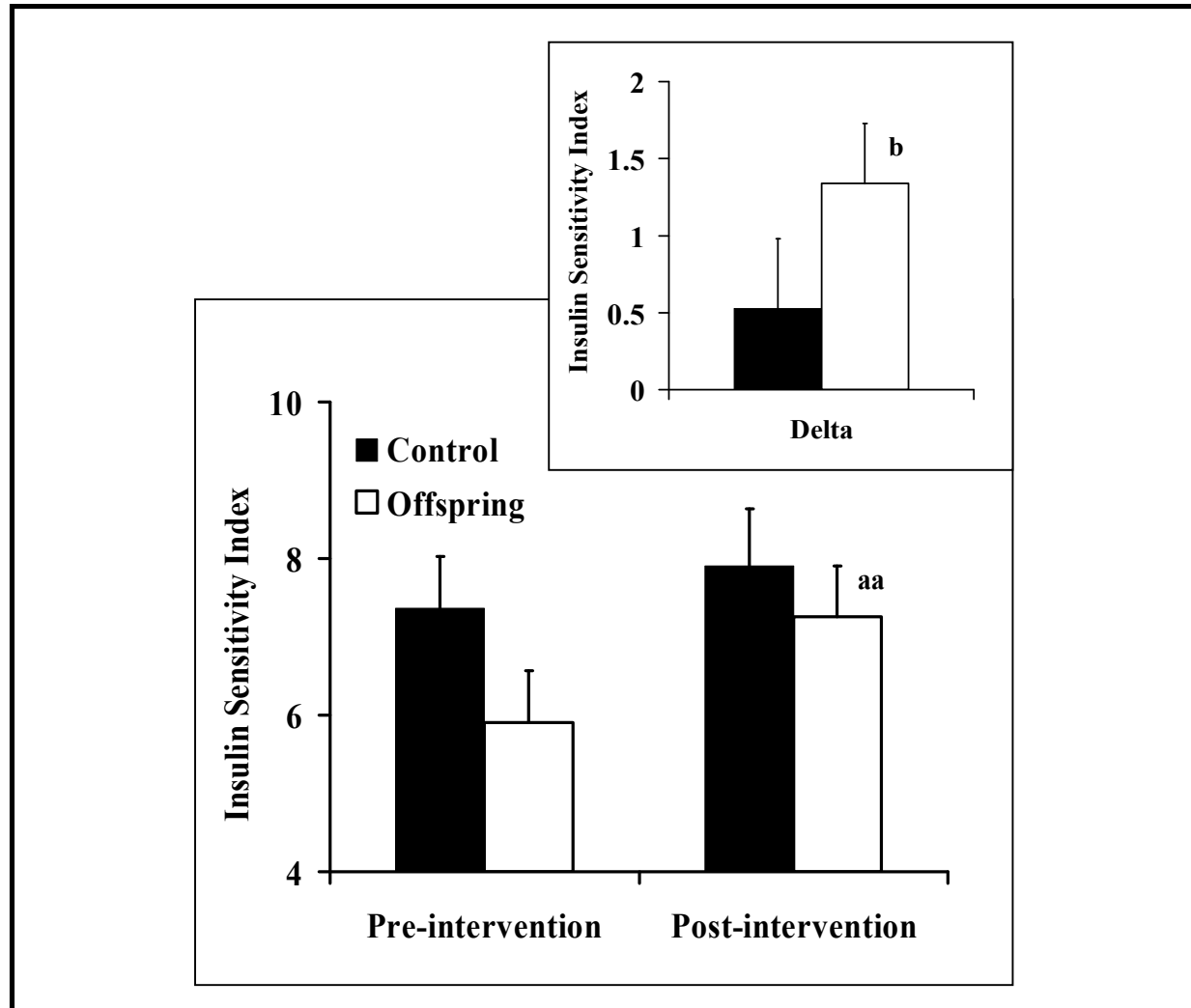
# The diabetes risk continuum





University  
of Glasgow

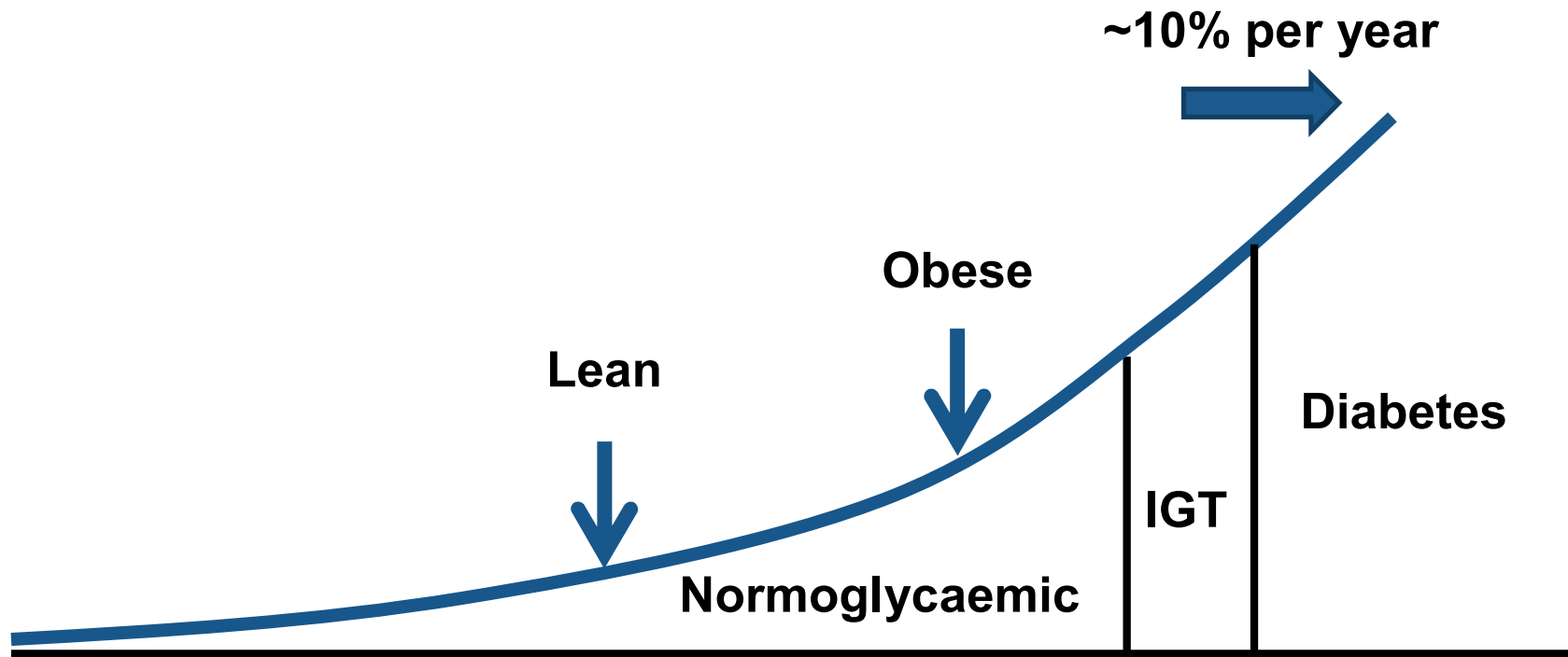
# Change in insulin sensitivity following a 7-week exercise intervention in women with and without a family history of diabetes



Barwell et al (2008) Diabetologia 51:1912-9

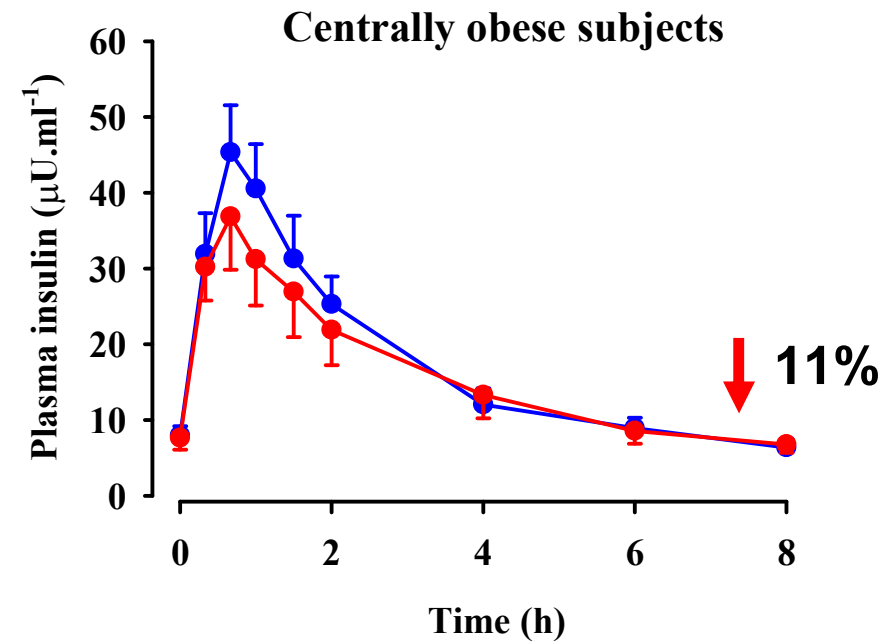
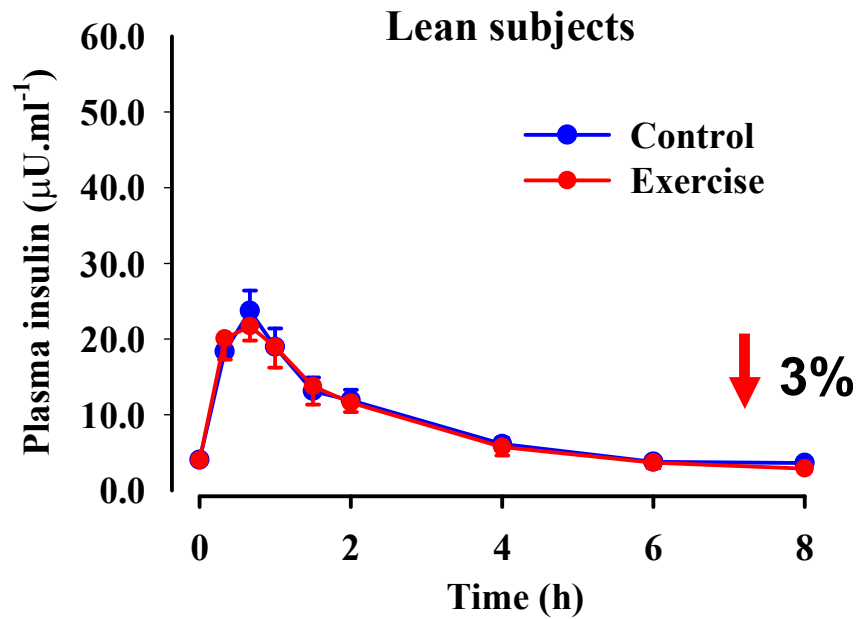


# The diabetes risk continuum





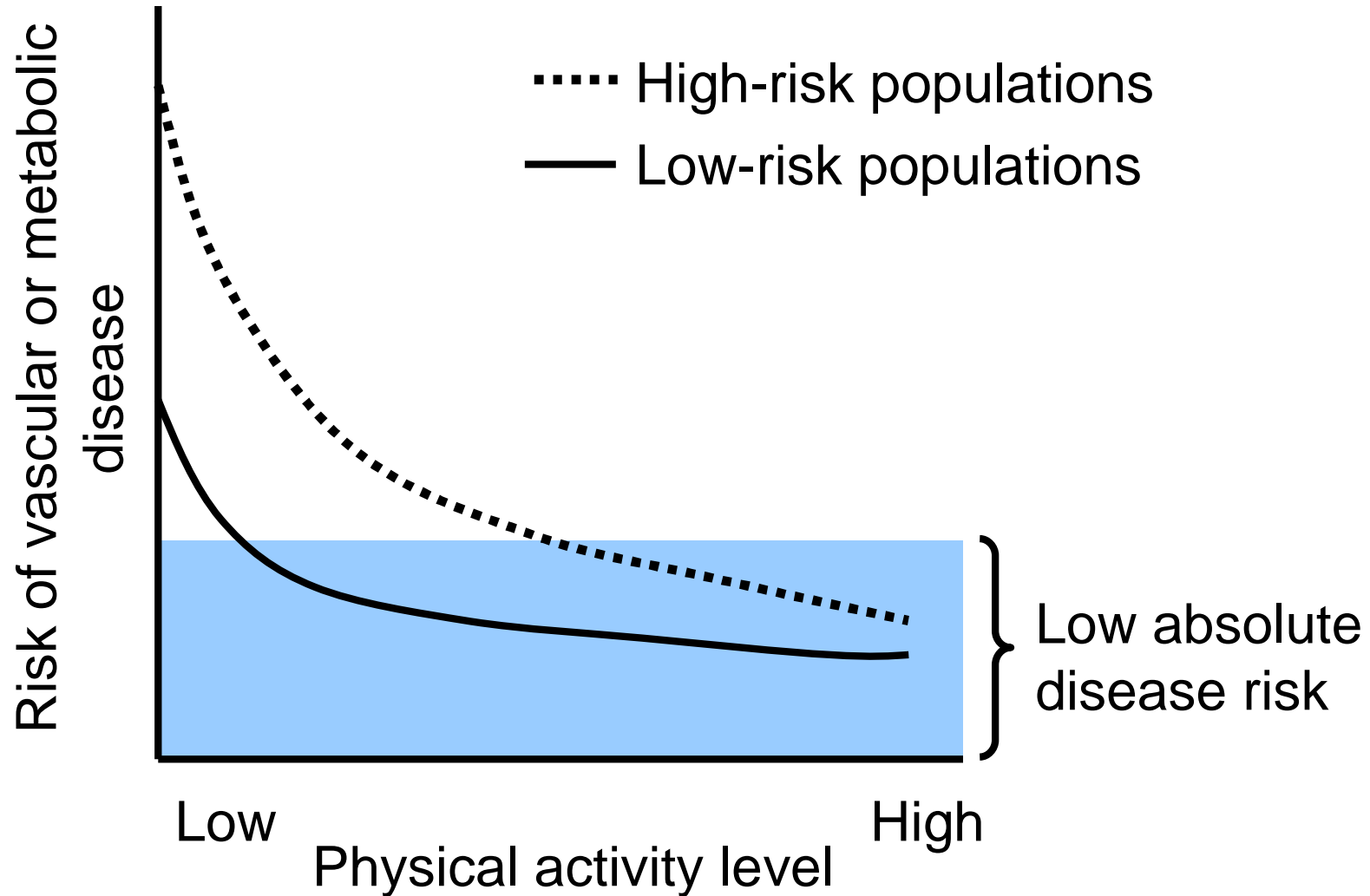
# Moderate exercise and insulin responses in lean and obese men



- “ Lifestyle intervention is effective at preventing diabetes in individuals with IGT
- “ But even with lifestyle intervention many with IGT will still develop diabetes.
- “ Within the normoglycaemic general population, there is a wide range of diabetes risk
- “ In groups at increased diabetes risk (e.g. diabetes family history, obese, South Asian, Mapuche), the benefits of increased physical activity appear particularly large



# How much physical activity do people need to do?





University  
of Glasgow

Should we be targeting these high-risk groups for diabetes prevention while they are still normoglycaemic?







# Acknowledgements

## Funders

- “ **British Heart Foundation**
- “ **Diabetes UK**
- “ **Translational Medicine Research Initiative**
- “ **Chest Heart and Stroke Scotland**
- “ **Government of Chile**



## Research Team

- “ **Dr Nick Barwell**
- “ **Dr Carlos Celis**
- “ **Dr Nazim Ghouri**
- “ **Dr Lesley Hall**
- “ **Dr Colin Moran**
  
- “ **Dr Mark Bailey**
- “ **Prof Carlos Calvo**
- “ **Dr Nita Forouhi**
- “ **Dr Niall MacFarlane**
- “ **Dr Alex McConnachie**
- “ **Prof Francisco Perez**
- “ **Mr David Purves**
- “ **Dr Ian Salt**
- “ **Prof Naveed Sattar**
- “ **Mr John Wilson**