1 Associations between anthropometric measurements and cardiometabolic risk

2 factors in White European and South Asian adults in the UK

- ³ Farah F. Kidy, MBChB, M.Sc.¹, Nafeesa Dhalwani, Ph.D.², Deirdre M. Harrington, Ph.D.
- ⁴^{2*}, Laura J Gray, Ph.D.¹, Danielle H. Bodicoat, Ph.D.², David Webb, MBChB, Ph.D.²,
- 5 Melanie J. Davies, MB, ChB, MD² and Kamlesh Khunti, MD, Ph.D.²
- ⁶ ¹Department of Health Sciences, University of Leicester, Leicester, United Kingdom
- ⁷² Diabetes Research Centre, University of Leicester, Leicester, United Kingdom
- 8 *Corresponding author details
- 9 Diabetes Research Centre (Origin wing), Leicester General Hospital, Leicester, LE5 4PW
- 10 Email: <u>dh204@le.ac.uk</u>
- 11
- 12 Financial support and conflict of interest disclosure:
- 13 ADDITION-Leicester was supported by the Department of Health and UK Support for
- Sciences, the NIHR Health Technology Assessment Programme (grant reference no:
- 15 08/116/300), National Health Service research and development support funding
- 16 (including the Primary Care Research and Diabetes Research Networks Leicestershire,
- 17 Northamptonshire and Rutland Collaborative for Leadership in Applied Health Research
- and Care) and the NIHR Leicester Loughborough Lifestyle Biomedical Research Unit.

Professor Melanie Davies has acted as consultant, advisory board member and speaker for Novo Nordisk, Sanofi-Aventis, Lilly, Merck Sharp & Dohme, Boehringer Ingelheim, AstraZeneca and Janssen, an advisory board member for Servier and as a speaker for Mitsubishi Tanabe Pharma Corporation and Takeda Pharmaceuticals International Inc. She has received grants in support of investigator and investigator initiated trials from Novo Nordisk, Sanofi-Aventis, Lilly, Boehringer Ingelheim and Janssen.

25	KK has acted as a consultant and speaker for Astra Zeneca, Novartis, Novo Nordisk,
26	Sanofi-Aventis, Lilly, Merck Sharp & Dohme, Janssen and Boehringer Ingelheim. He has
27	received grants in support of investigator and investigator initiated trials from Astra Zeneca,
28	Novartis, Novo Nordisk, Sanofi-Aventis, Lilly, Boehringer Ingelheim, Merck Sharp & Dohme
29	and Roche. KK has served on advisory boards for Astra Zeneca, Novartis, Novo Nordisk,
30	Sanofi-Aventis, Lilly, Merck Sharp & Dohme, Janssen and Boehringer Ingelheim.
31	The other authors have no conflict of interest related to this paper.
32	
33	
34	
35	
36	
37	
38	
39	
40	
41	
42	
43	
44	
45	
46	
47	
48	
49	

50 Abstract

51 *Objective:* To investigate the association of four anthropometric measurements with 52 cardiometabolic risk factors in a UK bi-ethnic sample in the UK of South Asians (SA) and 53 white Europeans (WE).

Patients: Baseline data from adults of WE and SA origin participating in the ADDITION Leicester study between August 2004 to December 2007.

Methods: Overall, 6,268 WE and SA adults had measures of body mass index (BMI), waist 56 circumference (WC), waist:hip ratio (WHR) and waist:height ratio (WHtR) assessed 57 between August 2004 and December 2007. Hypertension, dyslipidaemia and 58 dysglycaemia were established from venous blood samples using standard definitions. 59 Crude and adjusted (covariates used were age, sex, ethnicity, smoking and alcohol 60 consumption) odds ratios were calculated using multivariate logistic regression. Receiver 61 operating characteristic curves (ROC) and the area under the curve (AUC) were used to 62 calculate optimal cut points overall and for both ethnic groups. 63

Results: Increases in all anthropometric measurements resulted in higher odds of each of the risk factors in both the crude and adjusted models (P<.001). Adjusted odds of dyslipidaemia, hypertension and dysglygaemia ranged from 1.30 – 1.35, 1.36 – 1.52 and 1.62 – 1.75 (all P<.001), respectively, for WE. Adjusted odds of dyslipidaemia, hypertension and dysglygaemia ranged from 1.50 – 1.65 (P<.01), 1.40 – 1.60 (P<.01) and 1.96 – 2.11 (P<.001), respectively, for SA.

AUROCs for all of the anthropometric measurements had low accuracy (P<.70) for the whole cohort and when stratified by ethnicity and sex.

Conclusion: There is insufficient evidence to recommend replacing BMI with another
 anthropometric measurement for the ethnically diverse population in the UK.

74 Clinicaltrial.gov identifier NCT00318032

75	Abbreviations
76	AUC = area under the curve
77	BMI = body mass index
78	CI = confidence interval
79	OR = odds ratio
80	ROC = receiver operating characteristic curve
81	SA = South Asian
82	WC = waist circumference
83	WE = White European
84	WHR = waist to hip ratio
85	WHtR = waist to height ratio
86	
87	
88	
89	
90	
91	
92	
93	
94	
95	
96	

97 Introduction

Obesity is a recognised, modifiable risk factor for cardiovascular disease,¹ type 2 98 diabetes,^{2, 3} dyslipidaemia,⁴ hypertension⁵ and stroke.⁶ As obesity is often a precursor to 99 these chronic conditions it is important to have an assessment of adiposity that can identify 100 101 those at elevated risk. Adiposity based risk status can be assessed in a variety of ways including body mass index (BMI), waist circumference (WC), waist:hip ratio (WHR) and 102 waist:height ratio (WHtR). Evidence for the best measure at detecting those with increased 103 cardiometabolic risk remains equivocal. Available evidence is further complicated by ethnic 104 differences in the relationships between measures of adiposity and individual 105 cardiometabolic risk factors and a paucity of information on some populations such as 106 those of South Asians origin (countries in the Indian sub-continent). In order to add to the 107 body of literature regarding the use of anthropometric measurements to identify risk we 108 109 investigated four common anthropometric measurements to predict precursors to chronic disease in a bi-ethnic population from the UK. 110

111

112 Methods

113 Study population

Data have been taken from the population-based screening phase (baseline) of the 114 ADDITION-Leicester study,⁷ that formed part of ADDITION-Europe. Overall 6,749 South 115 Asian (SA) and white European (WE) adults who were not known to have diabetes, were 116 recruited through 20 general practices across Leicestershire, UK between August 2004 117 and December 2007. Potential participants were identified through the practice list and 118 invited to as assessment visit that took place at a hospital site or a mobile screening unit 119 located within their community. The age inclusion criteria was 40 – 75 years for WE and, 120 in acknowledgement of type 2 diabetes developing in younger people of minority 121

background, 25 – 75 years for SA. Those with complete data on all anthropometric
measurements and risk factors (n = 6268) are included herein. Those on antihypertensive
(n = 1425) and lipid lowering (n = 712) treatment were excluded from analyses of
hypertension and dyslipidaemia, respectively. Ethical approval was obtained from the
University Hospitals of Leicester (UHL09320) and Leicestershire Primary Care Research
Alliance (64/2004) local research ethics committees. Written informed consent was
obtained from all participants.

129

130 Anthropometric measurements

Anthropometric measurements were performed by trained staff following standard 131 operating procedures. Height was measured to the nearest 0.1 cm using a rigid 132 stadiometer. Weight was measured in light indoor clothing to the nearest 0.1 kg using a 133 Tanita scale (Tanita, Europe). WC was measured to the nearest 0.1 cm at the mid-point 134 between the lower costal margin and the level of the anterior superior iliac crest. Hip 135 circumference was measured to the nearest 0.1cm at the greatest protrusion of the gluteal 136 muscles. BMI was calculated as weight (kg) divided by height² (m). WHR and WHtR were 137 calculated as WC (cm) divided by hip circumference (cm) and height (cm), respectively. 138

139

140 Cardiometabolic risk factors

Arterial blood pressure was measured three times with the participant seated, using a standardised digital sphygmomanometer (Omron M7, Omron Healthcare, Milton Keynes, UK) with the average of the second and third readings used in the analysis. Participants undertook a 75g oral glucose tolerance test that included fasting and 2-hour venous blood samples. All blood samples were processed in the same pathology laboratory of the University Hospitals of Leicester NHS Trust, UK. Glucose was processed using an Abbott

Aeroset clinical chemistry analyser, which employs the hexokinase enzymatic method.
 HbA1c was analysed by a DCCT aligned Biorad Variant HPLC II system.

149

150 **Covariates**

Participants self-reported their ethnicity, current smoking status, alcohol consumption and occupation via questionnaire. Excess alcohol consumption was defined as more than 21 units per week in males and more than 14 units per week in females. Current and exsmokers were designated as 'ever smokers'.

155

156 **Definition of outcomes**

The criteria proposed by the Third Report of the National Cholesterol Education Program 157 Expert Panel on Detection, Evaluation and Treatment of High Blood Cholesterol in Adults 158 were used in defining cardiometabolic risk factors.⁸ Hypertension was defined as systolic 159 blood pressure ≥ 140 mmHg and/or diastolic blood pressure ≥90 mmHg. Raised total 160 cholesterol was defined as levels ≥5.2 mmol/l, raised low density lipoprotein cholesterol as 161 ≥3.36mmol/l, low high density lipoprotein cholesterol as <1.03 mmol/l, and raised 162 triglycerides as ≥1.7 mmol/l. Dyslipidaemia was defined as abnormal levels of one or more 163 lipid measurements. Type 2 diabetes was diagnosed using World Health Organisation 164 1999 criteria⁹ of fasting blood glucose \geq 7.0 mmol/l or an oral glucose tolerance test 2-hour 165 value ≥11.1 mmol/l. Impaired glucose tolerance (IGT) (fasting plasma glucose < 7.0 mmol/l 166 and an oral glucose tolerance test 2-hour value \geq 7.8 mmol/l but <11.1 mmol/l) and impaired 167 fasting glucose (fasting plasma glucose ≥6.1 mmol/l but <7.0 mmol/l) were treated as pre-168 diabetes (n = 865) and were combined with type 2 diabetes (n = 197) and designated as 169 dysglycaemia for the purposes of analyses. 170

171

172 Statistical Analysis

Continuous data are presented as mean (standard deviation) and categorical data as 173 frequency (percentage). Differences between WE and SA were assessed using t-tests for 174 continuous data and chi-squared test for categorical data. Standardised odds ratios (OR) 175 176 with 95% confidence intervals (95% CI) for cardiometabolic risk factors in relation to BMI, WC, WHR and WHtR were calculated using univariate and multivariate logistic regression. 177 The interaction between each anthropometric measure and ethnicity was assessed using 178 Wald's test. Although these were not significant data are still presented stratified by 179 ethnicity. For each model, age, gender, ethnicity, smoking status (smokers vs. ever 180 smokers) and excess alcohol intake were included as a priori confounders in the 181 multivariate analysis. ORs were standardised by using transformed observations 182 ([observation-mean]/SD) in the models. Crude and age-adjusted receiver operating 183 characteristic (ROC) curves were plotted and the area under the curve (AUC) calculated 184 for BMI, WC, WHR and WHtR, first for the cohort as a whole and then stratified by ethnicity 185 and sex. The optimal cut point for each measure of adiposity in detecting cardiometabolic 186 risk factors was chosen as the point on the curve with the highest Youden Index (sensitivity 187 + specificity -1). The age-adjusted AUCs generated for each anthropometric measure were 188 formally compared within each risk factor using the method suggested by DeLong et al.¹⁰ 189 A *P*-value of less than .05 was considered statistically significant. All data were analysed 190 using Stata IC version 14. 191

192

193 **Results**

194 Participant characteristics

Demographic data of the 6,268 participants included in the analyses herein are shown in Table 1. WEs in this sample had significantly higher BMI and WC (*P*<.001) compared to

SAs but there were no differences in WHR or WHtR. There were more than double the 197 percentage of those that ever smoked in the WE group compared with the SA group (WE 198 51% vs. % SA 17%, P<.001). A similar difference in proportions was seen in those with 199 200 excess alcohol consumption (WE 13% vs. SA 6%, P<.001). Dyslipidaemia was the most commonly seen risk factor, being present in 80% of the total population, 82% of WEs and 201 74% of SAs (*P*<.001). There were significantly more hypertensive WEs than SAs (47% vs. 202 35%, P<.001). There were significantly fewer participants with dysglycaemia amongst WEs 203 than SAs (16 % vs. 20%, P<.001). 204

205

Association of anthropometric measurements with cardiometabolic risk factors

The associations between each anthropometric measurement and cardiometabolic risk 207 factors stratified by ethnicity are shown in Table 2. Increases in all anthropometric 208 measurements resulted in higher odds of each of the risk factors in both the crude and 209 adjusted models (P<.001) except for WHR and dyslipidaemia in SA adults (P=.08). Odds 210 of dyslipidaemia, hypertension and dysglygaemia ranged from 1.30 – 1.35, 1.36 – 1.52 and 211 1.62 – 1.75, respectively, for WE and 1.29 – 1.65, 1.40 – 1.60 and 1.96 – 2.11 respectively 212 for SA. Due to overlapping confidence intervals, the odds were not significantly different 213 214 between anthropometric measurements.

215

216 Cut points for anthropometric measurements

The AUROC curves (95% CI) and optimum cut-points for predicting dyslipidaemia, hypertension and dysglycaemia for each of the anthropometric measurements are presented in Table 3. Although significantly different, the AUROCs for all of the anthropometric measurements had low accuracy¹¹ for detecting each cardiometabolic risk factor in both the crude and age-adjusted analyses. For dyslipidaemia, the optimum cut-

points were 24 kg/m² (sensitivity = 80, specificity = 34, AUROC = 0.582) for BMI, 85 cm 222 (sensitivity = 75, specificity = 41, AUROC = 0.599) for WC, 0.86 (sensitivity = 64, specificity 223 = 52, AUROC = 0.598) for WHR and 0.51 (sensitivity = 78, specificity = 37, AUROC = 224 225 0.588) for WHtR. For hypertension, the optimum cut-points were 25 kg/m² (sensitivity = 74, specificity = 42, AUROC = 0.599) for BMI, 92 cm (sensitivity = 59, specificity = 59, AUROC 226 = 0.613) for WC, 0.92 (sensitivity = 43, specificity = 72, AUROC = 0.597) for WHR and 227 0.54 (sensitivity = 0.65, specificity = 0.52, AUROC = 0.607) for WHtR. For dysglycaemia, 228 the optimum cut-points were 27 kg/m² (sensitivity = 67, specificity = 53, AUROC = 0.633) 229 for BMI, 91 cm (sensitivity = 74, specificity = 47, AUROC = 0.640) for WC, 0.91 (sensitivity 230 = 55, specificity = 62, AUROC = 0.606) for WHR and 0.54 (sensitivity = 81, specificity = 231 43, AUROC = 0.666) for WHtR. The age-adjusted values presented in Table 3 were slightly 232 higher but the AUROCs still being considered low accuracy at <0.70. 233

234

Table 4 shows the results of ROC analyses stratified by ethnicity. Similar to the analysis of 235 the cohort as a whole, the AUCs were all low for the crude and age-adjusted analyses. The 236 optimal BMI cut point for predicting dyslipidemia was higher in WEs (24 kg/m²) than SAs 237 (23 kg/m^2) , was the same (25 kg/m^2) for hypertension and slightly higher in WEs (28 kg/m^2) 238 than SAs (27 kg/m²) for dysglycaemia. The optimal WC cut for dyslipidaemia was higher in 239 South Asians (89 cm) than WEs (84 cm) but for dysglycaemia was lower in SAs (91 cm) 240 than WEs (97 cm). For further clinical applicability Table S1 presents the results stratified 241 by both ethnicity and sex. Again, all AUROCs were low between anthropometric 242 measurements and between groups. 243

244

We also investigated the performance (i.e. the sensitivity and specificity) of commonly used BMI and WC cut-points on the cohort as a whole. For BMI of 30 kg/m² the performance was 29 and 76, 31 and 78, 43 and 74 for dyslipidaemia, hypertension and dysglycaemia,

respectively. For WC of 102 cm the performance was 26 and 82, 29 and 82 and 39 and 76
for dyslipidaemia, hypertension and dysglycaemia, respectively.

250

251 Discussion

Using data from a large bi-ethnic cohort, we found that a number of common 252 anthropometric measurements had similarly low, although statistically significant different, 253 associations with cardiometabolic risk factors. As obesity continues to be a global problem, 254 measurements that are acceptable to patients and healthcare professionals alike are 255 needed to identify people in the population who are most risk of developing cardiometabolic 256 morbidity and mortality in order to signpost for appropriate testing or intervention.¹² The 257 results herein would suggest that each of these measurements have similarly low utility in 258 identifying those who may benefit from further confirmatory tests or general lifestyle based 259 prevention strategies. 260

261

262 In the sample as a whole our analysis has shown that all four measures of adiposity (BMI, WC, WHR and WHtR) had a low capacity to predict individual cardiometabolic risk factors 263 and, similar to a study investigating the ability of these measures in predicting type 2 264 diabetes,¹⁴ no clear pattern emerged for any measure that was superior. Although the 265 AUCs reported in table 3 were statistically different, they are all lower than those reported 266 on in previous cross-sectional,¹³ meta-analysis¹⁵ and bi-racial analysis from the US.¹⁶ 267 However, the differences in populations (none of the included studies were UK based or 268 had South Asian cohort) and definitions of the risk factors may account for these 269 differences. 270

Studies have reported on ethnic differences in the performance of common anthropometric 272 measurements.^{17, 18}As the AUROCs were low we did not formally test for differences in the 273 performance of the measurements by ethnic group. However, we did find ethnic 274 275 differences in the optimal cut-point for dyslipidaemia (84 cm vs. 89 cm), hypertension (92 cm vs. 90 cm) and dysglycemia (97 cm vs. 91 cm). The optimal cut points that reduce the 276 level of false positives would suggest that lower cut points for South Asians would be 277 supported. National and international guidelines do support the use of ethnic specific cut-278 points.^{19, 20} as reviews have pointed out the large disparity in optimal cut-points between 279 and within ethic groups.²¹ 280

281

Meta-analytical strategies suggest that a measure of central obesity, such as WC or WHtR, 282 is superior to BMI for identifying hypertension, type 2 diabetes and dyslipidemia.^{15, 21, 22} 283 However, papers have cautioned that the discriminatory capability differences between 284 BMI and individual measures of central obesity were clinically non-significant.²¹ As none of 285 the studies included in these reviews included a large cohort of South Asian adults the 286 results herein add to the body of evidence comparing the utility of common anthropometric 287 measurements in those of South Asian background. As obesity is a heterogeneous 288 condition referring to excess adipose tissue deposited both subcutaneously and 289 viscerally^{23, 24} it is both the excess total fat and its distribution which are important to 290 assess. It is unlikely therefore that any single measure of adiposity will be adequate to 291 correctly identify all those at risk in a given population. Even in those with a normal BMI 292 there is value in further exploration using WC,²⁵ dual-energy X-ray absorptiometry²⁶ or % 293 294 body fat from air displacement plethysmography.^{27, 28} Although suggested by guidelines,^{29,} ³⁰ the practicality of even adding a simple WC or % body fat measurement to a BMI 295 measurement in routine clinical care may be difficult given the constraints on healthcare 296 professional time and the limitations to bioelectrical impedance outputs.³¹ Although WC is 297

²⁹⁸ often more correlated with body fat than BMI, WC is just as correlated with total body fat ²⁹⁹ as with abdominal fat.¹⁶

300

To our knowledge, this is one of the largest studies to date to compare the utility of common 301 anthropometric measures in predicting cardiometabolic risk within two different ethnic 302 groups in the UK. However, we used data from the screening phase of ADDITION-303 Leicester, thus only making use of cross-sectional data with no account of longitudinal risks 304 or the inclusion of a hard clinical end point. Data on the inter- and intra- technician reliability 305 of the anthropometric measurements was not collected, however, variability would be 306 minimised as the technicians were trained and followed the same standardised operating 307 protocol which is more that would happen if that measure were collected in routine clinical 308 practice. Previous analyses reported that BMI, WC and WHR had similar correlates with 309 10-year risk of fatal cardiovascular disease³² while both BMI and WC were associated with 310 increased all-cause, cardiovascular disease and cancer mortality risk³³ indicating that 311 these measures have similar value from a longitudinal point of view for diabetes.^{22, 34} 312 Although the site used herein (mid-point between the iliac crest and the lowest floating rib) 313 is recommend by the World Health Organization³⁰ as a WC measurement site, the iliac 314 crest is often used, and recommended for use,³¹ in US contexts. Although the absolute 315 value of the measurement can differ between sites, the mid-point site has been equally 316 well correlated with cardiometabolic risk factors compared with the iliac crest site.³⁵ The 317 SAs enrolled in this study were members of a migrant population and the duration of time 318 spent in the UK was not assessed. Due to potential heterogeneity in lifestyle and dietary 319 320 factors, these results cannot be generalised across all SA populations. Further research is needed to confirm whether anthropometric measurements such as WHR or WHtR adds 321 any additional information to composite risk scores which already include either BMI or WC 322 or both. Particularly in the SA population, more work is needed to assess the utility of these 323

anthropometric measurements in a longitudinal fashion. Our data would suggest that there
 is little to be gained by simply replacing BMI or WC with another measure.

326

327 Conclusion

Obesity and its associated conditions remain of public health concern and it is important 328 that public health interventions are appropriately targeted. Weight based anthropometric 329 calculations have been used to indicate disease risk historically¹ and currently there is a 330 large number of anthropometric measurements for healthcare professionals, policy makers 331 and researchers to choose from. Although statistically there was a difference in the 332 performance of the indicators of adiposity for each risk factor no clean pattern was seen in 333 the performance as all were similarly low. The variety of anthropometric measurements 334 can be utilised pragmatically as a screening tool to identify adults who may be at risk of 335 chronic disease and who may benefit from further tests/confirmatory tests. Similar to 336 previous reviews²¹ there is insufficient evidence to recommend one anthropometric 337 measurement over another. However, due to its historical use and the amassed 338 epidemiological evidence BMI would seem to be the most suitable measurement to be 339 done alone or in conjunction with an indicator of central adiposity. However, healthcare 340 341 professionals should always be mindful of patient preference, equipment available and the skill of their team. 342

343

344

345

346

347

348

350 **References**

- 1. Hubert HB, Feinleib M, McNamara PM, Castelli WP. Obesity as an independent risk factor for
- cardiovascular disease: a 26- year follow-up of participants in the Framingham Heart Study.
 Circulation. 1983;67(5):968-977.
- 2. Jafar TH, Chaturvedi N, Pappas G. Prevalence of overweight and obesity and their association
- with hypertension and diabetes mellitus in an Indo-Asian population. *Canadian Medical Association Journal.* 2006;175(9):1071-1077.
- 357 3. Wannamethee SG, Papacosta O, Whincup PH, Carson C, Thomas MC, Lawlor DA, et al.
- Assessing prediction of diabetes in older adults using different adiposity measures: a 7 year
- prospective study in 6,923 older men and women. *Diabetologia*. 2010;53(5):890-898.
- 4. Arnlov J, Ingelsson E, Sundstrom J, Lind L. Impact of Body Mass Index and the Metabolic
- Syndrome on the Risk of Cardiovascular Disease and Death in Middle-Aged Men. *Circulation*.
 2010;121(2):230-236.
- 363 5. Dyer AR, Elliott P. The INTERSALT study: relations of body mass index to blood pressure.
- *Journal Human Hypertension*. 1989;3(5):299-308.
- 6. Rexrode KM. A prospective study of body mass index, weight change, and risk of stroke in
- women. JAMA: The Journal of the American Medical Association. 1997;277(19):1539-1545.
- 367 7. Webb D, Khunti K, Srinivasan B, Gray L, Taub N, Campbell S, et al. Rationale and design of the
- ADDITION-Leicester study, a systematic screening programme and randomised controlled trial of multi-factorial cardiovascular risk intervention in people with type 2 diabetes mellitus detected
- 370 by screening. *Trials.* 2010;11(1):16.
- 8. National Cholesterol Education Program Expert Panel on Detection Evaluatiom, Treatment of
- 372 High Blood Cholesterol in Adults. Third Report of the National Cholesterol Education Program
- 373 (NCEP) Expert Panel on Detection, Evaluation, and Treatment of High Blood Cholesterol in Adults
- (Adult Treatment Panel III) final report. *Circulation*. 2002;106(25):3143-3421.
- 9. Alberti KG, Zimmet PZ. Definition, diagnosis and classification of diabetes mellitus and its
- 376 complications. Part 1: diagnosis and classification of diabetes mellitus provisional report of a
- WHO consultation. *Diabetic Medicine*. 1998;15(7):539-553.
- 10. DeLong ER, DeLong DM, Clarke-Pearson DL. Comparing the areas under two or more
- 379 correlated receiver operating characteristic curves: a nonparametric approach. *Biometrics*. 1988;
 380 44(3): 837-45.
- 11. Swets JA. Measuring the accuracy of diagnostic systems. *Science*. 1988;240(4857):1285-1293
- 12. Ng M, Fleming T, Robinson M, Thomson B, Graetz N, Margono C, et al. Global, regional, and
- national prevalence of overweight and obesity in children and adults during 1980–2013: a
- systematic analysis for the Global Burden of Disease Study 2013. *The Lancet.* 2014;384(9945):
 766-781.
- 13. Bosy-Westphal A, Geisler C, Onur S, Korth O, Selberg O, Schrezenmeir J, et al. Value of body
- fat mass vs anthropometric obesity indices in the assessment of metabolic risk factors.
- 388 International Journal of Obesity. 2006;30(3):475-483.
- 14. MacKay MF, Haffner SM, Wagenknecht LE, D'Agostino RB, Hanley AJG. Prediction of type 2
- diabetes using alternate anthropometric measures in a multi-ethnic cohort: The Insulin
 Resistance Atherosclerosis Study. *Diabetes Care*. 2009;32(5):956-958.
- 15. Ashwell M, Gunn P, Gibson S. Waist-to-height ratio is a better screening tool than waist
- circumference and BMI for adult cardiometabolic risk factors: systematic review and meta-
- analysis. *Obesity Reviews*. 2012;13(3):275-286.
- 16. Barreira TV, Staiano AE, Harrington DM, Heymsfield SB, Smith SR, Bouchard C, et al.
- 396 Anthropometric correlates of total body fat, abdominal adiposity, and cardiovascular disease risk
- factors in a biracial sample of men and women. *Mayo Clinic Proceedings.* 2012;87(5):452-460.

- 17. Bodicoat DH, Gray LJ, Henson J, Webb D, Guru A, Misra A, et al. Body mass index and waist
- 399 circumference cut-points in multi-ethnic populations from the UK and India: the ADDITION-
- Leicester, Jaipur heart watch and New Delhi cross-sectional studies. *PloS One.* 2014;9(3):e90813.
- 18. Shen W, Punyanitya M, Chen J, Gallagher D, Albu J, Pi-Sunyer X, et al. Waist circumference
- 402 correlates with metabolic syndrome indicators better than percentage fat. *Obesity*.
- 403 2006;14(4):727-736.
- 19. National Institute for Health and Care Excellence. *Type 2 Diabetes: Prevention in People at High Risk.* 2012.
- 20. International Diabetes Federation. The IDF Consensus Worldwide Definition of the MetabolicSyndrome. 2006.
- 408 21. Lee CMY, Huxley RR, Wildman RP, Woodward M. Indices of abdominal obesity are better
- discriminators of cardiovascular risk factors than BMI: a meta-analysis. *Journal of Clinical Epidemiology.* 2008;61(7):646-653.
- 411 22. Qiao Q, Nyamdorj R. Is the association of type II diabetes with waist circumference or waist-
- to-hip ratio stronger than that with body mass index? *European Journal of Clinical Nutrition*.
 2010;64(1):30-34.
- 414 23. Despres JP. Body Fat Distribution and risk of cardiovascular disease: An update. *Circulation*.
 415 2012;126(10):1301-1313.
- 416 24. Vega GL, Adams-Huet B, Peshock R, Willett D, Shah B, Grundy SM. Influence of body fat
- 417 content and distribution on variation in metabolic risk. *Journal of Clinical Endocrinology and* 418 *Metabolism.* 2006;91(11):4459-4466.
- 25. Janssen I, Katzmarzyk PT, Ross R. Waist circumference and not body mass index explains
 obesity-related health risk. *American Journal of Clinical Nutrition*. 2004;79(3):379-384.
- 421 26. Camhi SM, Bray GA, Bouchard C, Greenway FL, Johnson WD, Newton RL, et al. The
- relationship of waist circumference and BMI to visceral, subcutaneous, and total body fat: sex
- and race differences. Obesity. 2011;19(2):402-408.
- 424 27. Gómez-Ambrosi J, Silva C, Galofré JC, Escalada J, Santos S, Gil MJ, et al. Body adiposity and
- 425 type 2 diabetes: Increased risk with a high body fat percentage even having a normal BMI.
- 426 *Obesity*. 2011;19(7):1439-1444.
- 427 28. Gómez-Ambrosi J, Silva C, Galofré JC, Escalada J, Santos S, Millán D, et al. Body mass index
- 428 classification misses subjects with increased cardiometabolic risk factors related to elevated
- adiposity. *International Journal of Obesity*. 2012;36(2):286-294.
- 430 29. National Institutes of Health. *Clinical Guidelines on the Identification, Evalaution and*
- 431 Treatment of Overweight and Obesity in Adults. 1998.
- 432 30. World Health Organization. Waist circumference and waist-hip ratio: report of a WHO expert
- consultation, Geneva, 8-11 December 2008. Geneva: World Health Organization; 2011 2011.
- 434 31. Cornier M-A, Després J-P, Davis N, Grossniklaus DA, Klein S, Lamarche B, et al. Assessing
- adiposity a scientific statement from the American Heart Association. *Circulation*.
- 436 2011;124(18):1996-2019.
- 437 32. Marques-Vidal P, Bochud M, Mooser V, Paccaud F, Waeber G, Vollenweider P. Obesity
- markers and estimated 10-year fatal cardiovascular risk in Switzerland. *Nutrition, Metabolism and Cardiovascular Diseases*. 2009;19(7):462-468.
- 33. Staiano A, Reeder B, Elliott S, Joffres M, Pahwa P, Kirkland S, et al. Body mass index versus
- waist circumference as predictors of mortality in Canadian adults. *International Journal of Obesity*. 2012;36(11):1450-1454.
- 34. Tulloch-Reid MK, Williams DE, Looker HC, Hanson RL, Knowler WC. Do measures of body fat
- distribution provide information on the risk of type 2 diabetes in addition to measures of general
- obesity? Comparison of anthropometric predictors of type 2 diabetes in Pima Indians. *Diabetes*
- 446 *Care.* 2003;26(9):2556-2561.

447 448	35. Mason C, Katzmarzyk PT. Effect of the site of measurement of waist circumference on the prevalence of the metabolic syndrome. <i>The American Journal of Cardiology</i> . 2009;103(12):1716-
449	20.
450	
451	
452	
453	
454	
455	
456	
457	
458	
459	
460	
461	
462	
463	
464	
465	
466	
467	
468	
469	
470	
471	
472	
473	
474	
475	
476	
477	
478	
479	

480 **Table 1.** Baseline characteristics

	Total sample (N = 6,268)	White Europeans (N = 4,604)	South Asians (N = 1,664)	P-value (WE vs SA)
Males number (%)	2979 (47.5)	2162 (47.0)	817 (49.1)	.134
Age (years)	56.1 (10.7)	58.5 (9.5)	49.2 (11.1)	<.001
Body mass index (kg/m²)	28.0 (5.0)	28.3 (4.9)	27.3 (5.0)	<.001
Waist circumference (cm)	93 (13)	94 (13)	92 (12)	<.001
Waist to hip ratio	0.89 (0.08)	0.89 (0.08)	0.89 (0.08)	.775
Waist to height ratio	0.56 (0.07)	0.56 (0.08)	0.57 (0.07)	.136
Ever smoker (%)	2597 (42)	2313 (51)	284 (17)	<.001
Excess alcohol consumption (%)	569 (12)	525 (13)	44 (6)	<.001
Dyslipidaemia [†] (%)	4450 (80)	3327 (82)	1123 (74)	<.001
Hypertension [†] (%)	2065 (43)	1589 (47)	476 (35)	<.001
Dysglycaemia (%)	1065 (17)	735 (16)	330 (20)	<.001

481 Note: continuous variables are presented as means with SD in parenthesis and categorical variables are

482 presented as %. WE – White Europeans, SA - South Asians. [†]Analysis of dyslipidaemia and hypertension

483 exclude those on lipid-lowering and antihypertensive treatment, respectively.

		Adjusted OP (95% CI)	Dyalua for
		Adjusted OK (95% CI)	adjusted OR
Dyslipidaemia			
White Europeans			
BMI	1.38 (1.26 – 1.51)	1.30 (1.18 – 1.44)	<.001
WC	1.41 (1.30 – 1.54)	1.32 (1.20 – 1.46)	<.001
WHR	1.36 (1.25 – 1.47)	1.35 (1.20 – 1.53)	<.001
WHtR	1.48 (1.36 – 1.62)	1.32 (1.20 – 1.45)	<.001
South Asians			
BMI	1.18 (1.05 – 1.33)	1.65 (1.29 – 2.11)	<.001
WC	1.41 (0.24 – 1.61)	1.50 (1.17 – 1.93)	.002
WHR	1.57 (1.38 – 1.78)	1.29 (0.97 – 1.72)	.08
WHtR	1.20 (1.07 – 1.36)	1.52 (1.19 – 1.96)	.001
Hypertension	· · · · ·		
White Europeans			
BMI	1.41 (1.31 – 1.52)	1.52 (1.40 – 1.66)	<.001
WC	1.47 (1.36 – 1.57)	1.45 (1.33 – 1.58)	<.001
WHR	1.43 (1.33 – 1.53)	1.36 (1.23 – 1.51)	<.001
WHtR	1.51 (1.41 – 1.63)	1.47 (1.35 – 1.61)	<.001
South Asians			
BMI	1.44 (1.28 – 1.62)	1.60 (1.29 – 1.98)	<.001
WC	1.63 (1.43 – 1.85)	1.60 (1.28 – 2.01)	<.001
WHR	1.49 (1.32 – 1.69)	1.40 (1.06 – 1.85)	.02
WHtR	1.54 (1.37 – 1.74)	1.49 (1.20 – 1.86)	<.001
Dysglycaemia			
White Europeans			
BMI	1.58 (1.46 – 1.70)	1.62 (1.48 – 1.77)	<.001
WC	1.66 (1.54 – 1.80)	1.75 (1.59 – 1.93)	<.001
WHR	1.46 (1.34 – 1.58)	1.73 (1.54 – 1.94)	<.001
WHtR	1.78 (1.64 – 1.93)	1.75 (1.60 – 1.92)	<.001
South Asians			
BMI	1.56 (1.39 – 1.75)	1.96 (1.56 – 2.47)	<.001
WC	1.79 (1.56 – 2.04)	2.11 (1.64 – 2.73)	<.001
WHR	1.51 (1.33 – 1.72)	2.06 (1.49 – 2.85)	<.001
WHtR	1.78 (1.57 – 2.02)	2.03 (1.59 – 2.59)	<.001

Table 2 – Crude and adjusted standardised odds Ratio (95% CI) for cardiometabolic risk factors in relation to anthropometric measures in the whole cohort, stratified by ethnicity.

OR = Odds Ratio, CI = confidence interval, BMI = body mass index, WC= waist circumference, WHR = waist to hip ratio, WHtR = waist to height ratio. Each measure has been transformed.

Adjusted model is adjusted for age, sex, ethnicity, smoking status and excess alcohol consumption. *all models significant at *P*<.01

	Crude AUC (95% CI)	Crude optimal cut point	Sens (%)	Spec (%)	Adjusted AUC (95% Cl)*	Adjusted optimal cut point*	Sens (%)*	Spec (%)*	Р
Dyslipidaer	nia								
BM	l 0.582 (0.562 – 0.601)	24	80	34	0.621 (0.602 – 0.640)	22	72	47	.006
WC	0.599 (0.580 – 0.618)	85	75	41	0.630 (0.611 – 0.650)	93	78	44	
WF	IR 0.598 (0.579 – 0.617)	0.86	64	52	0.632 (0.613 – 0.652)	0.78	73	50	
WF	ltR 0.588 (0.569 – 0.608)	0.51	78	37	0.620 (0.601 – 0.640)	0.50	81	38	
Hypertensio	on								
BM	l 0.599 (0.583 – 0.615)	25	74	42	0.680 (0.665 – 0.695)	30	72	48	<.001
WC	0.613 (0.597 – 0.629)	92	59	59	0.684 (0.669 – 0.699)	106	73	49	
WF	IR 0.597 (0.580 – 0.613)	0.92	43	72	0.677 (0.662 – 0.692)	0.89	67	55	
WF	ltR 0.607 (0.591 – 0.623)	0.54	65	52	0.679 (0.664 – 0.694)	0.54	77	42	
Dysglycaen	nia								
BM	l 0.633 (0.615 – 0.651)	27	67	53	0.663 (0.645 – 0.680)	23	73	47	<.001
WC	0.640 (0.622 – 0.658)	91	74	47	0.664 (0.647 – 0.682)	81	76	46	
WF	IR 0.606 (0.587 – 0.624)	0.91	55	62	0.642 (0.624 – 0.660)	0.88	73	51	
WF	ltR 0.666 (0.649 – 0.684)	0.54	81	43	0.682 (0.665 – 0.699)	0.52	79	40	

Table 3. Crude and adjusted AUC and optimal cut points for anthropometric measurements in relation to cardiometabolic risk factors for the whole cohort

AUC = area under the receiver-operating characteristics curve, CI = confidence interval, sens = sensitivity, spec= specificity, BMI = body mass index, WC = waist circumference, WHR = waist to hip ratio, WHtR = waist to height ratio, * = Adjusted model is adjusted for age, *P* value derived by comparing AUC across all four anthropometric measures.

Table 4. Crude and adjusted AUC and optimal cut points for measures of adiposity in relation to cardiometabolic risk factors, by ethnicity

	Crude AUC (95%CI)	Crude optimal cut point	Sens (%)	Spec (%)	Adjusted AUC (95% CI)*	Adjusted optimal cut point*	Sens (%)*	Spec (%)*	Р
Dyslipidaemia									
White Europeans	S								
BMI	0.590 (0.567 – 0.614)	24 (79.7)	82	32	0.631 (0.607 – 0.656)	24	70	50	<.001
WC	0.596 (0.572 – 0.620)	84 (74.2)	77	39	0.633 (0.608 – 0.657)	92	73	50	
WHR	0.587 (0.563 – 0.610)	0.84 (70.3)	73	42	0.626 (0.601 – 0.651)	0.95	69	53	
WHtR	0.609 (0.585 – 0.633)	0.51 (74.6)	78	41	0.635 (0.611 – 0.659)	0.56	67	52	
South Asians									
BMI	0.553 (0.519 – 0.587)	23 (82.4)	85	26	0.555 (0.520 – 0.590)	18	73	48	<.001
WC	0.594 (0.560 – 0.627)	89 (53.9)	58	59	0.594 (0.560 - 0.628)	91	73	47	
WHR	0.627 (0.594 – 0.660)	0.86 (61.4)	67	55	0.628 (0.595 – 0.661)	0.85	75	44	
WHtR	0.555 (0.521 – 0.590)	0.49 (86.0)	89	22	0.555 (0.520 – 0.590)	0.49	72	47	
Hypertension									
White Europeans	5								
BMI	0.594 (0.575 – 0.613)	25 (66.8)	75	41	0.668 (0.650 - 0.686)	27	75	45	.01
WC	0.603 (0.584 - 0.622)	92 (51.5)	60	56	0.671 (0.653 - 0.687)	73	74	48	
WHR	0.597 (0.578 – 0.616)	0.92 (35.2)	43	72	0.666 (0.648 - 0.683)	0.93	67	55	
WHtR	0.608 (0.589 - 0.627)	0.54 (52.2)	61	55	0.668(0.650 - 0.686)	0.51	78	41	
South Asians	, , , , , , , , , , , , , , , , , , ,				, ,				
BMI	0.601 (0.570 - 0.632)	25 (61.8)	72	44	0.680 (0.650 - 0.708)	20	71	48	<.001
WC	0.631 (0.601 – 0.662)	90 (48.8)	63	59	0.687(0.658 - 0.716)	91	72	30	
WHR	0.602(0.571 - 0.634)	0.93 (29.0)	40	77	0.673(0.644 - 0.703)	0.79	76	46	
WHtR	0.618 (0.587 – 0.648)	0.54 (58.5)	71	49	0.677 (0.647 – 0.706)	0.58	78	41	
Dysglycaemia				-			-		
White Europeans	6								
BMI	0.641 (0.619 – 0.662)	28 (43.9)	62	59	0.688 (0.667 - 0.708)	22	73	47	<.001
WC	0.643(0.621 - 0.664)	97 (40.9)	58	62	0.686(0.665 - 0.706)	109	77	44	
WHR	0.602(0.580 - 0.624)	0.91 (40.9)	53	61	0.660 (0.640 - 0.681)	0.78	69	53	
WHtR	0.666(0.645 - 0.687)	0.58 (40.2)	61	64	0.696(0.676 - 0.716)	0.49	70	50	
South Asians		0.00 (10.2)	01	0.		0.10		00	
BMI	0.632 (0.600 - 0.663)	27 (62.3)	61	61	0 679 (0 648 – 0 709)	26	70	51	< 001
WC	0.655(0.625 - 0.686)	91 (49.8)	70	55	0.683(0.652 - 0.713)	Q2	76	46	2.001
WHR	0.616(0.582 - 0.649)	0.91 (39.5)	57	65	0.648 (0.615 - 0.681)	0.73	73	50	
WHtR	0.667 (0.638 - 0.697)	0.54 (61.8)	85	44	0.687 (0.657 - 0.717)	0.75	73	50	

AUC = area under the receiver-operating characteristics curve, CI = confidence interval, sens = sensitivity, spec= specificity, BMI = body mass index, WC = waist circumference, WHR = waist to hip ratio, WHtR = waist to height ratio, * = Adjusted model is adjusted for age, *P* value derived by comparing AUC across all four anthropometric measures.