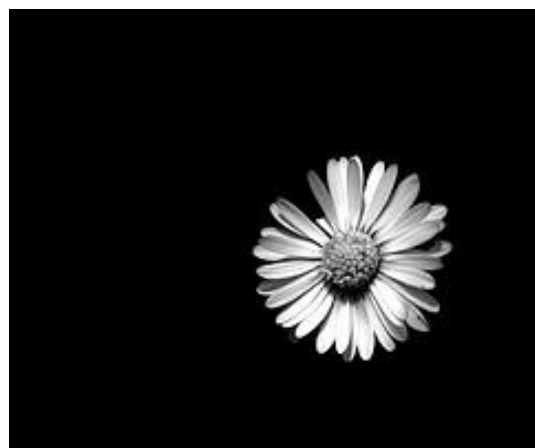


**The association between central obesity  
and psychological distress in a group of  
urban Africans: the SABPA study**

**J. Botha**



2012



**The association between central obesity and  
psychological distress in a group of urban  
Africans: the SABPA study**

**J. BOTHA (M.A.)**  
12618144

Thesis submitted for the degree Doctor of Philosophy at the  
Potchefstroom campus of the North-West University

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May 2012

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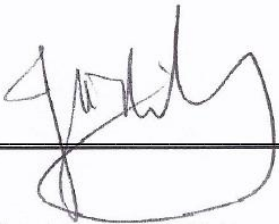
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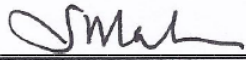
# Declaration

The co-authors of the articles which form part of this dissertation, Prof J. Hans de Ridder (promotor), Prof Leoné Malan (co-promotor), Prof Johan Potgieter (co-promotor) and Prof Faans Steyn (assistant-promotor), hereby give permission to the candidate, Me Judy Botha, to include the three articles as part of a Doctoral dissertation. The contribution, both supervisory and supportive, of these co-authors was kept within limits, thereby enabling the candidate to submit this dissertation for examination purposes. This dissertation, therefore, serves as fulfilment of the requirements for the Ph.D. degree within the School of Biokinetics, Recreation and Sport Science in the Faculty of Health Sciences at the North-West University, Potchefstroom Campus.



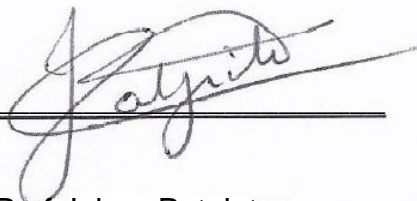
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Promotor and co-author



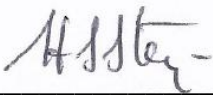
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Prof Leoné Malan  
Co-promotor and co-author



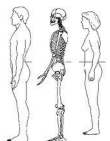
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Prof Johan Potgieter  
Co-promotor and  
Co-author



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Prof Faans Steyn  
Assistant promotor and  
Co-author



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# Summary

## Introduction:

Obesity plays an important role in the development of the metabolic syndrome, with central obesity being included as a required prerequisite in the most recent definition of the metabolic syndrome by the International Diabetes Federation (IDF). The following components were included in the Joint Statement Consensus (JSC) definition: raised blood pressure (systolic BP  $\geq 130$  or diastolic BP  $\geq 85$  mm Hg), raised triglycerides (Trig) ( $\geq 1.7$  mmol/L) and lowered high-density lipoprotein cholesterol (HDL) ( $< 1.03$  mmol/L in males and  $< 1.29$  mmol/L in females), raised fasting glucose ( $\geq 5.6$  mmol/L), and central obesity. Population- and country-specific definitions for waist circumference is recommended, although the IDF cut points are to be used for non-Europeans until more data are available.

## Aim:

Consequently, the aim of the study was to determine a population-specific waist circumference (WC) cut off, comparing the new proposed waist circumference model (NPM) with the current cut offs proposed by the Joint Statement Consensus (JSC). Association between the new proposed cut off and perception of own health in a group of urban African teachers was also investigated.

## Method:

WC, sphygmomanometer blood pressure, fasting bloods (glucose, HDL and triglycerides) and ultrasound carotid intima-media thickness (CIMT) were obtained for 171 black urban teachers from the sympathetic activity and ambulatory blood pressure in Africans (SABPA) study. Perception of own health was determined via the General Health Questionnaire-28 (GHQ-28). Gender-separate receiver operating curve (ROC) analyses were performed for each of the metabolic syndrome components to determine a new population-specific waist circumference cut off. Subsequently Logistic Regression and Neural Networks analyses were performed in order to validate the NPM. Thereafter the association between the NPM and perception of own health was considered.



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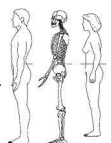
### Results:

The respective blood pressure cut offs corresponded best with WC pathology, and a NPM was proposed with the new WC cut off for the males be set at 90 cm as opposed to the current 94 cm; whilst the female cut off be set at 98 cm as opposed to the existing cut off of 80 cm. Thereafter ROC analyses (not adjusting for covariates), Logistic Regression and Neural Networks statistics (both adjusting for a priori confounders, age, BMI and physical activity) validated that the NPM model was comparable to the JSC model. Both models correlated with CIMT, an indicator of structural vascular disease. When comparing the JSC and NPM WC models, only the NPM model was associated with perception of own poorer health.

### Conclusion:

A new population and ethnicity-specific WC cut off was recommended (NPM). Subsequently the NPM was validated via Logistic Regression and Neural Networks statistical analyses. The NPM was comparable with the JSC cut offs which are currently in use in predicting structural vascular disease via CIMT. It is proposed that the NPM cut offs be used in this population due to the strong association between blood pressure and the proposed WC cut offs, validated by Logistic Regression and Neural Networks statistical analyses. Furthermore, associations were demonstrated between the NPM and perception of own health in a group of urban Africans.

**Keywords:** [central obesity, metabolic syndrome, Setswana, ROC, Logistic Regression, Neural Networks, perception of health]



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# Opsomming

## Inleiding:

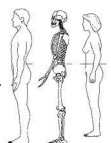
Obesiteit speel 'n belangrike rol in die ontwikkeling van die metaboliese sindroom, met obesiteit ingesluit as voorvereiste in die mees onlangse definisie van die metaboliese sindroom deur die Internasionale Diabetes Federasie (IDF). Die volgende komponente is by die *Joint Statement Consensus* (JSC)-definisie ingesluit: verhoogde bloeddruk (sistoliese bloeddruk  $\geq 130$  of diastoliese bloeddruk  $\geq 85$  mm Hg), verhoogde trigliseriedes ( $\geq 1.7$  mmol/L) en verlaagde hoë densiteit lipoproteïen cholesterol (HDL) ( $< 1.03$  mmol/L in mans en  $< 1.29$  mmol/L in dames), verhoogde vastende glukose ( $\geq 5.6$  mmol/L), en sentrale obesiteit. Populasie- en land-spesifieke definisies vir maagomtrek word voorgestel, alhoewel die IDF afsnyppunte vir nie-Europeërs gebruik word totdat meer data beskikbaar is.

## Doelwit:

Gevolgtrek was die eerste doel van die studie om 'n populasie-spesifieke maagomtrek (WC) afsnyppunt te bepaal (NPM), dit te vergelyk met die afsnyppunt wat tans deur die JSC voorgestel word en ondersoek daarna in te stel of die nuut voorgestelde afsnyppunt verband hou met persepsie van eie gesondheid in 'n groep Afrikaan onderwysers.

## Metode:

WC, sphygmomanometer bloeddruk, vastende bloed (glukose, HDL en trigliseried waardes) en karotis intima-media dikte (CIMT) is vir 171 swart stedelike onderwysers bepaal in die *sympathetic activity and ambulatory blood pressure in Africans* (SABPA)-studie. Persepsie van eie gesondheid is bepaal via die *General Health Questionnaire-28* (GHQ-28). Geslagte is afsonderlik geanaliseer deur *receiver-operating curve* (ROC) analyses vir elk van die metaboliese-sindroom-komponente om 'n nuwe populasie-spesifieke maagomtrek afsnyppunt te bepaal. Daarna is Logistiese Regressie en Neurale Netwerk-analises gedoen om sodoende die NPM te valideer. Laastens is die assosiasie tussen die NPM en persepsie van eie gesondheid bepaal.



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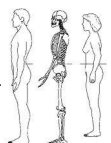
### Resultate:

Daar is gevind dat die onderskeie bloeddruk (BP) afsnypunte die beste korrespondeer met maagomtrek-patologie. Nuwe maagomtrek-afsnypunte is gestel op 90 cm teenoor die huidige 94 cm, terwyl die dames afsnypunt gestel word op 98 cm teenoor die huidige 80 cm. Daarna is ROC-analises (waar nie vir enige veranderlikes gekorrigeer word nie) en Neurale Netwerke-statistiek (waar daar vir a priori veranderlikes ouderdom, BMI en fisieke aktiwiteit gekorrigeer word) gebruik om die NPM te valideer. Die NPM was vergelykbaar met die JSC-model in korrelasie CIMT, 'n aanduider van strukturele vaskulêre siekte. Vervolgens, het die NPM assosiasie tussen sentrale obesiteit en persepsie van eie gesondheid uitgewys.

### Gevolgtrekking:

'n Populasie en etnisiteit spesifieke WC afsnypunt is voorgestel (NPM). Daaropvolgend is die NPM via Logistiese Regressie en Neurale Netwerk statistiese analises gevalideer. Die NPM was vergelykbaar met die JSC-afsnypunte wat huidiglik in gebruik is in die voorspelling van strukturele vaskulêre siekte via CIMT. Daar word voorgestel dat die NPM-afsnypunte in hierdie populasie gebruik moet word as gevolg van die sterk assosiasie tussen bloeddruk en die voorgestelde afsnypunte, gevalideer deur Logistiese Regressie en Neurale Netwerk statistiese analises. Verder is assosiasies ook gevind tussen die NPM en persepsie van gesondheid in 'n groep stedelike Afrikane.

**Sleutelwoorde:** [sentrale obesiteit, metaboliese sindroom, Setswana, ROC, Logistiese Regressie, Neurale Netwerke, persepsie van gesondheid]



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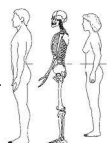
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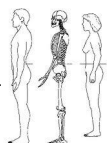
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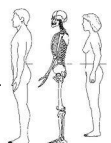
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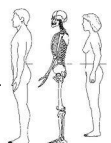
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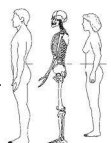
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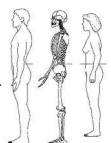


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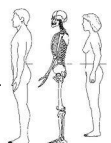
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# List of Abbreviations

|                              |  |
|------------------------------|--|
| <u>Anthropometry:</u>        | WC_____Waist Circumference                 |
|                              | kg_____Kilogram/s                          |
|                              | m_____Meter/s                              |
|                              | BMI_____Body Mass Index                    |
| <br>                         |  |
| <u>Physiological:</u>        | BP_____Blood Pressure                      |
|                              | HDL_____High Density Lipoprotein           |
|                              | Trig_____Triglyceride                      |
|                              | CIMT_____Carotid intima-media thickness    |
| <br>                         |  |
| <u>Psychological:</u>        | GHQ-28_____General Health Questionnaire-28 |
| <br>                         |  |
| <u>Organizations:</u>        | WHO_____World Health Organization          |
|                              | IDF_____International Diabetes Federation  |
|                              | JSC_____Joint Statement Consensus          |
| <br>                         |  |
| <u>Statistical analyses:</u> | NPM_____New Proposed Model                 |
|                              | ROC_____Receiver operating characteristic  |
|                              | AUC_____Area under the curve               |
|                              | CI_____Confidence Interval                 |
|                              | 95% CI_____95 % Confidence Interval        |
|                              | SD_____Standard Deviation                  |
|                              | SE_____Standard Error                      |
|                              | LR_____Logistic Regression                 |
|                              | OR_____Odds Ratio                          |
|                              | NN_____Neural Networks                     |
|                              | MLP_____Multilayer perceptron              |

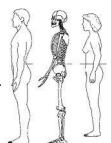


- 1.1 Introduction
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  - 1.3 Hypotheses
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**1.1 Introduction**

A cluster of risk factors for cardiovascular disease and type 2 diabetes mellitus have become known as the metabolic syndrome. These factors include increased blood pressure, dyslipidaemia (raised triglycerides and lowered high-density lipoprotein cholesterol), increased fasting glucose, and central obesity (Alberti *et al.*, 2009:1640). Most patients with risk factors such as cardiovascular disease and co-morbidities such as diabetes, dyslipidaemia, and hypertension have also been found to be associated with clearly altered carotid intima-media thickness (CIMT) (Kotliar *et al.*, 2008:61). In addition, the degree of its alteration increases with the number of coexisting risk factors and the time of exposure, especially in the presence of metabolic syndrome (Kotliar *et al.*, 2008:61).

The current epidemic of type 2 diabetes and metabolic syndrome could be a direct result of our energy-dense diet and affluent sedentary lifestyle, where such a lifestyle could increase the likelihood of individuals eating more than they need (Després & Lemieux, 2006:885). This positive energy balance consequently leads to abdominal obesity and insulin resistance in the presence of an unfavourable genotype and other permissive factors (Després & Lemieux, 2006:885).



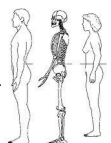
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Stress, anxiety, alcohol and smoking may also contribute to the endocrine abnormalities that promote abdominal obesity (Björntorp, 1995:21; Björntorp, 2001:75). Various studies have shown that a relationship of waist circumference to metabolic abnormalities is age-, gender-, as well as ethnicity-dependent (Lemieux *et al.*, 1996:983; Han *et al.*, 1997:593; Després *et al.*, 2000:1932; Misra *et al.*, 2005:969; Després *et al.*, 2008:1041). The marked differences across racial and ethnic groups in disease risk are likely to be due, in part, to each of genetic, host susceptibility and environmental factors (Forouhi & Sattar, 2006:11, Hamer & Malan, 2010:76, Malan *et al.*, 2010:183, Hamer *et al.*, 2011:237, Malan *et al.*, 2012:549, De Kock *et al.*, 2012).

Urbanization has an effect on the traditional ideal body image among Africans, who have always been inclined towards a larger, fuller body shape. (Szabo & Le Grange, 2001:31, Malan *et al.*, 2008:323). In the traditional body image obesity was associated with dignity, health, wealth and respect (Puoane *et al.*, 2005). Black young South Africans may, however, rapidly become exposed to different belief systems due to Westernization and thereby alter their value systems regarding body size (Caradas *et al.*, 2001:112), which may in turn influence their perception of own health (Malan *et al.*, 2008:323).

Urbanization could also, with accompanying insecurities and disruption in African social relationships, contribute to experiencing poorer health or distress (Van Rooyen *et al.*, 2000:779, Malan *et al.*, 2006:305, Malan *et al.*, 2008:323). A component of the Whitehall II Study examined psychological distress as a risk factor for coronary heart disease (CHD) and found that the experience of psychological distress conferred increased risk of CHD in males, although it did not consistently do so in females (Stansfeld *et al.*, 2002:248). Interestingly, the increased risk of psychological distress for CHD, in the Whitehall II Study, was not explained by health behaviours (Stansfeld *et al.*, 2002:248), although Hamer *et al.* (2011:237) found that the excess burden of sub-clinical vascular disease in black Africans can be largely explained by health behaviours and conventional risk factors.

The strength of this study lies in the current void in literature where population- and country-specific guidelines for WC have not yet been appropriated for sub-Saharan Africans; also the association between obesity and mental health have not been



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investigated in the black South African population. Subsequently, the research questions we aim to answer by means of this study are: (1) Can we establish new population-specific WC cut off points in a cohort of urban African teachers? (2) Which WC cut off point model (JSC or new proposed model) (NPM) is best associated with structural vascular disease, as indicated by carotid intima-media thickness (CIMT)? and (3) Which of the two WC models will be best associated with perception of own health in this African cohort?

## **1.2. Aims**

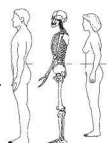
The specific aims of this study were derived from the above-mentioned research questions and are as follows:

- To establish new population-specific WC cut off points in a cohort of urban Africans.
- To determine which WC cut off point model (JSC or NPM) will be best associated with structural vascular disease, as indicated by carotid intima-media thickness (CIMT) in a cohort of urban Africans.
- To examine the association between each of the two WC models (JSC and NPM) and perception of own health in a cohort of urban Africans.

## **1.3. Hypotheses**

The following hypotheses are proposed for this investigation:

- New population-specific WC cut off points in a cohort of urban Africans would differ from those proposed by the JSC.
- New population-specific WC cut off points would predict structural vascular disease (CIMT) in a cohort of urban Africans.
- New population-specific WC cut off points would predict a poorer perception of own health in a cohort of urban Africans.



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## 1.4 Structure of the dissertation

This dissertation is presented in four main parts, namely an introduction (Chapter 1), a literature review (Chapter 2), and three research articles (Chapters 3, 4 and 5). Thereafter a summary with conclusions and recommendations will follow (Chapter 6).

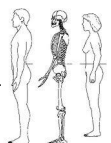
Chapter 1 introduces the problem and states the aim and hypotheses of this study.

Chapter 2, the literature review, focuses on the relationship between obesity and psychological distress.

Chapter 3 will take the form of an article: **Determining the waist circumference cut off which best predicts the Metabolic Syndrome components in urban Africans: the SABPA study.** This article was accepted for publication by Experimental and Clinical Endocrinology & Diabetes, a journal with an impact factor of 1.89 (Prinsloo, J., Malan, L., De Ridder, J.H., Potgieter, J.C. & Steyn, H.S. 2011. Determining the waist circumference cut off which best predicts the metabolic syndrome components in urban Africans: the SABPA study. Experimental and Clinical Endocrinology and Diabetes, 119:599-603).

Chapter 4 consists of the second research article titled: **Comparing performances of two central obesity models to predict structural vascular disease by using ROC analyses, Logistic Regression and Neural networks: the SABPA study.**; this article was prepared for Atherosclerosis and is currently in the review process.

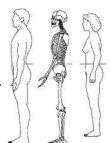
Chapter 5 will also take the form of an article: **Association of waist circumference with perception of own health in a group of urban African males and females: the Sympathetic Activity and Ambulatory Blood Pressure in Africans (SABPA) study.** This article was submitted to the Journal of Endocrinology, Metabolism and Diabetes of South Africa and is in rebuttal.



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**In chapters 3, 4 and 5, the research articles have been prepared according to the guidelines of each respective journal.**

Chapter 6, the final chapter, will wrap up with the conclusion and recommendations of both research articles. Chapter 6 is followed by a list of appendices.



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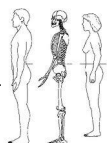
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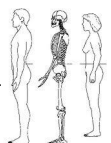
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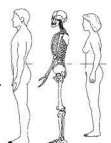
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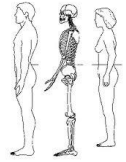
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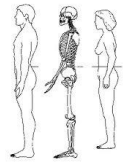
**Chapter****2****The relationship between central obesity and psychological distress**

- 2.1 Introduction
- 2.2 The metabolic syndrome - past and present
- 2.3 Population- and country-specific cut off points for central obesity
- 2.4 Hypertension and carotid intima-media thickness (CIMT)
- 2.5 Psychological distress and perception of health
- 2.6 Conclusion
- 2.7 References

**2.1 Introduction**

Overweight and obesity are associated with increased cardiovascular morbidity and mortality (Rao *et al.*, 2001:1102; Vega, 2001:1108). Most patients with risk factors such as cardiovascular disease and co-morbidities such as diabetes, dyslipidaemia, and hypertension have also been found to be associated with clearly altered carotid intima-media thickness (CIMT) (Kotliar *et al.*, 2008:61). Lorenz *et al.* (2007:459) also confirmed in a meta-analysis that CIMT is a strong predictor of future vascular events. Consequently, CIMT measurement is regarded as an excellent tool to detect preclinical vascular disease (Kotliar *et al.*, 2008:61).

Central obesity also forms part of the constellation termed metabolic syndrome. The 2009 Joint Statement Consensus (JSC) risk factors include raised blood pressure (systolic BP  $\geq 130$  or diastolic BP  $\geq 85$  mm Hg), raised triglycerides (Trig) ( $\geq 1.7$  mmol/L) and lowered high-density lipoprotein cholesterol (HDL) ( $< 1.03$  mmol/L in males and  $< 1.29$  mmol/L in females), raised fasting glucose ( $\geq 5.6$  mmol/L), and central obesity (Alberti *et al.*, 2009:1640). Three abnormal findings out of five would qualify a person for



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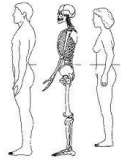
the metabolic syndrome. A single set of cut points would be used for all components except waist circumference, for which further ethnicity- and country-specific work is required (Alberti *et al.*, 2009:1640).

Culturally worthy of note in black African females is their perception regarding body weight and body image. Puoane *et al.* (2005a:6) found that socio-cultural, behavioural and environmental factors seem to influence the development of obesity in this population. It is proposed that, compared with white women, non-Westernized and some groups of Westernized black women adopt a larger ideal body size and they are more accepting towards being overweight (Faber & Kruger, 2005:238; Kruger *et al.*, 2005:493; Puoane, 2005b:92; Schutte & Olckers, 2007:651). Obesity should, however, not be viewed as a benign condition amongst any ethnic population of South Africa (Kruger *et al.*, 2005:497).

Typically, psychological co-morbidity is high in patients with obesity and is associated with a variety of medical and dietary problems, as well as demographic, social and cognitive risk factors (Van der Merwe, 2007:S14). Whether this is the case with the traditional African female is indefinite. Increased knowledge of behavioural risk factors has, however, enabled patients with obesity to be classified on a psychological basis and this needs to be considered part of a patient's clinical assessment and treatment strategy (Van der Merwe, 2007:S14).

## **2.2 The Metabolic Syndrome - past and present**

Prior to the Joint Statement Consensus (Alberti *et al.*, 2009:1640) addressing the definition of the metabolic syndrome, different international organizations and bodies formulated various definitions for the metabolic syndrome. These included the definitions of the World Health Organization (WHO) (Alberti & Zimmet, 1998), the European Group for the Study of Insulin Resistance (EGIR) (Balkau & Charles, 1999:442), the National Education, Cholesterol Program's Adult Treatment Panel III (NCEP-ATP III, 2001) and finally, the International Diabetes Federation (IDF) (Alberti *et al.*, 2009:1640).



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Table 1 illustrates the various organizations' criteria.

In 2009, a Joint Statement Consensus (JSC) was released by the International Diabetes Federation Task Force on Epidemiology and Prevention; National Heart, Lung, and Blood Institute; American Heart Association; World Heart Foundation; International Atherosclerosis Society; and International Association for the Study of Obesity. This JSC decided that a single set of cut points would be used for all components except waist circumference (WC), for which further research is required (Alberti *et al.*, 2009:1640).

Population-specific cut offs have already been accepted for Asian populations by the International Diabetes Federation (Alberti *et al.*, 2009:1642). Sub-Saharan Africans, however, still use the cut offs proposed for the European and Caucasian populations, with the male cut off set at  $\geq 94$  cm and the female WC cut off set at  $\geq 80$  cm.

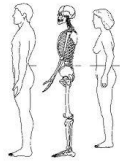
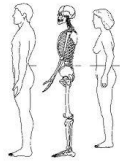
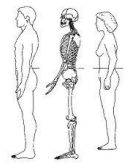


Table 1: Various definitions of the metabolic syndrome - 1998 to 2009.

|  | <b>Classify as</b>   | <b>Blood Pressure</b>  | <b>Dyslipidaemia</b>   | <b>Central Obesity</b>  | <b>Other factors:</b>   |
|--|--|--|--|---|---|
|  | <b>diabetes when:</b>  |  |  |   |   |
| <b>WHO (Alberti &amp; Zimmet, 1998)</b>  | Glucose intolerance, impaired glucose tolerance or diabetes mellitus and/or insulin resistance together, and any 2 of the following: | raised arterial pressure ( $\geq 140/90$ mmHg);  | raised plasma triglycerides ( $\geq 1.7$ mmol <sup>-1</sup> ) and/or low HDL-cholesterol ( $< 0.9$ mg.g <sup>-1</sup> males; $< 1.0$ mg.g <sup>-1</sup> females) | central obesity (waist to hip ratio males $> 0.90$ , females $> 0.85$ ) and/or BMI $> 30$ kg/m <sup>2</sup> | microalbuminuria (urinary albumin excretion rate $\geq 20$ $\mu$ g min <sup>-1</sup> or albumin:creatinine ratio $\geq 30$ mg g <sup>-1</sup> ) |
| <b>European Group for the Study of Insulin Resistance (EGIR) (Balkau &amp; Charles, 1999:442)</b>                | The presence of insulin resistance or fasting hyperinsulinaemia (the highest 25%)  | hypertension (systolic/diastolic blood pressures $\geq 140/90$ mmHg or treated for hypertension) | dyslipidaemia (triglycerides $> 2.0$ mmol/L or HDL cholesterol $< 1.0$ mmol/L or treated for dyslipidaemia)  | central obesity (waist circumference $\geq 94$ cm in males and $\geq 80$ cm in females).                    | hyperglycaemia (fasting plasma glucose $\geq 6.1$ mmol/L, but nondiabetic)  |
| All of these criteria must be measured before it is possible to evaluate the presence of the metabolic syndrome. |  |  |  |   |   |



|  | <b>Classify as diabetes when:</b>   | <b>Blood Pressure</b>  | <b>Dyslipidaemia</b>  | <b>Central Obesity</b>  | <b>Other factors:</b>                       |
|--|---|--|---|---|---|
| <b>The National Education, Cholesterol Program's Adult Treatment Panel III (NCEP-ATP III) (2001)</b> | Did not list any principal criteria   | hypertension (systolic/diastolic blood pressures $\geq 135/85$ mmHg or treated for hypertension) | raised plasma triglycerides ( $\geq 1.7$ mmol <sup>-1</sup> ) and/or low HDL-cholesterol ( $< 1.03$ mg.g <sup>-1</sup> males; $< 1.29$ mg.gl <sup>-1</sup> females) | central obesity (waist circumference $\geq 94$ cm in males and $\geq 80$ cm in females).  |   |
| <b>Joint Statement Consensus (JSC) (Alberti <i>et al.</i>, 2009:1640)</b>                            | Three abnormal findings out of 5 (raised blood pressure, dyslipidemia and raised fasting glucose) would qualify a person for the metabolic syndrome | raised blood pressure (systolic BP $\geq 130$ or diastolic BP $\geq 85$ mm Hg)                   | raised triglycerides (Trig) ( $\geq 1.7$ mmol/L) and lowered high-density lipoprotein cholesterol (HDL) ( $< 1.03$ mmol/L in males and $< 1.29$ mmol/L in females)  | central obesity (proposed until population and ethnicity-specific guidelines are set: waist circumference $\geq 94$ cm in males and $\geq 80$ cm in females). | raised fasting glucose ( $\geq 5.6$ mmol/L) |



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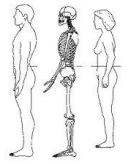
### 2.3 Population- and country-specific cut off points for central obesity

Differences in body composition are observed in different ethnic groups (Wagner & Heyward, 2000:1392); Blacks have increased skeletal muscle mass and bone mineral content, whereas Asians have less skeletal muscle mass, low bone mineral content, and excess body fat for a given BMI (Misra *et al.*, 2005:970). Kumar *et al.* (2006:686) found that ethnic differences in obesity measures persisted despite adjusting for age and known socio-demographic, biological and lifestyle factors.

Factors that might influence waist circumference is laxity of anterior abdominal muscles, poor posture, phases of respiration, as well as water, food and gases contained in hollow viscera (Misra *et al.*, 2005:970). Heterogeneity of composition of abdominal tissues, in particular adipose tissue and skeletal muscle, and their location-specific and changing relations with metabolic factors and cardiovascular risk factors in different ethnic groups do not, however, allow a simple definition of abdominal obesity that could be applied uniformly (Wagner & Heyward, 2000:1392; Misra *et al.*, 2005:969; Carroll *et al.*, 2008:607).

Sex differences in fat distribution are thought to be due to changes in the local levels of sex steroids/hormones i.e. androstenedione to testosterone and estrone to estradiol (Ahima, 2006:243S). Nindl *et al.* (2002:1611) demonstrated that the main gender differences in body composition are that males have more muscle mass in their upper limbs and females carry more of their fat mass in their legs.

Regarding ethnicities, Després *et al.* (2000:1932-1933) found that black females had a greater body fat content and higher levels of visceral adipose tissue than white females, suggesting a lower susceptibility to visceral obesity in black females. White men, on the other hand, had higher levels of visceral adipose tissue than black men, (Després *et al.*, 2000:1932). In a review study, Wagner and Heyward (2000:1399-1400) unequivocally found that the fat free mass of blacks and whites differ significantly; with increased bone mineral content and bone mineral density in blacks, shown by cadaver and in vivo analyses.



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Consequently, population-specific and country-specific definitions for waist circumference (WC) are recommended, although the International Diabetes Federation (IDF) cut points are to be used for non-Europeans until more data are available (Alberti *et al.*, 2009:1640). Several studies have proposed new population-specific cut points (Table 2). Proposed cut offs in the various studies ranged from 71.5 cm to 96 cm for female participants and from 76 cm to 106 cm for male participants.

The European Society of Hypertension position statement on the metabolic syndrome in hypertension states that obesity and insulin resistance have been implicated in the pathogenesis of the metabolic syndrome (Redon *et al.*, 2009:1891). Evidence now indicates that central obesity plays a central role in the development of the metabolic syndrome and appears to precede the appearance of other metabolic syndrome components linked to insulin resistance (Fezeu *et al.*, 2007:70; Cameron *et al.*, 2008:2707; Després *et al.*, 2008:1039).

Conversely, Chambers *et al.* (2008:716) carried out a genome-wide association study and found that a common genetic variation near *MC4R* is associated with risk of adiposity and insulin resistance. In general, the metabolic syndrome components have a high degree of interaction where one component contributes to the establishment of abnormality in other components and vice versa (Redon *et al.*, 2009:1893). Redon *et al.* (2009:1893) adds that obesity and insulin resistance may play an important role in the increment of blood pressure and the development of hypertension, although the precise mechanisms involved remain partially unresolved.

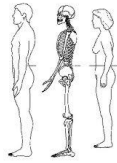
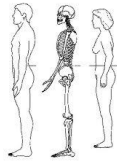
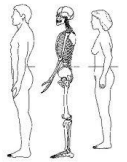


Table 2: Proposed race and gender-specific waist circumference cut off points.

| Source                                | Population / Ethnicity                         | Nr of Participants         | Female cut-off (cm)          | Male cut-off (cm)             |
|---------------------------------------|--|----------------------------|------------------------------|-------------------------------|
| Okusun <i>et al.</i> , 2000a:180      | Nigeria; Cameroon; Jamaica; St Lucia; Barbados | 3983 males; 4763 females   | 72; 82; 85; 86;<br>88        | 76; 81; 80;<br>83; 87         |
| Okusun <i>et al.</i> , 2000b:1279     | White, black and Hispanic Americans            | 7613 males; 7207 females   | 82 - 91; 81 - 90;<br>83 - 92 | 89 - 106; 84 -<br>95; 87 - 97 |
| Berber <i>et al.</i> , 2001:1794      | Mexico   | 2426 males; 5939 females   | 85                           | 90                            |
| Foucan <i>et al.</i> , 2002:992       | Guadeloupe                                     | 5149 females               | 83.5                         |                               |
| Lin <i>et al.</i> , 2002:1232         | Taiwan   | 26359 males; 29204 females | 71.5                         | 80.5                          |
| Mirmiran <i>et al.</i> , 2004:1110    | Tehran   | 4449 males; 6073 females   | 79 - 96                      | 80 - 93                       |
| Wang & Hoy, 2004:1581                 | Australia (Aborigines)                         | 473 males; 442 females     | 91                           | 86                            |
| Shiwaku <i>et al.</i> , 2005:52       | Asia (361 Japanese; 252 Mongolians)            | 388 males; 364 females     | 73; 84                       | 82; 92                        |
| Hara <i>et al.</i> , 2006:1123 & 1124 | Japan  | 408 males; 284 females     | 78                           | 85                            |
| Al-Lawati & Jousilhati, 2007:102      | Omani Arab                                     | 696 males; 725 females     | 84.5                         | 80                            |
| Bouguerra <i>et al.</i> , 2007:859    | Tunisia  | 1244 males; 2191 females   | 85                           | 85                            |
| Hayashi <i>et al.</i> , 2007:120      | Japanese American                              | 344 males; 295 females     | 80 - 90                      | 87 - 90                       |
| Mansour <i>et al.</i> , 2007:1        | Iraq   | 700 males; 300 females     | 99                           | 97                            |
| Neufeld <i>et al.</i> , 2007:159      | Mexico   | 802 females                | 89.3 - 91.2                  |                               |
| Oka <i>et al.</i> , 2007:474          | Japan  | 1061 males; 809 females    | 82.3                         | 89.8                          |



|                                      |                     |                            |  |  |
|--------------------------------------|---------------------|----------------------------|--|--|
| Al-Lawati <i>et al.</i> , 2008:304   | Omani Arab          | 680 males, 704 females     | 84.5   | 78.5   |
| Esteghamati <i>et al.</i> , 2008:104 | Iran                | 1046 males; 1706 females   | 85.5   | 91.5   |
| Matoba <i>et al.</i> , 2008:590      | Japan               | 1658 male; 1116 females    | 80   | 87   |
| Sumner <i>et al.</i> , 2008:841      | African American    | 68 males; 63 females       | ≥98  | ≥102   |
| Bao <i>et al.</i> , 2008:378         | Chinese             | 525 males; 615 females     | 85   | 90   |
| Cameron <i>et al.</i> , 2008:1       | Europid; South Asia | 3525 males, 4201 females   | 84; 76                                       | 97; 79                                       |
| Hadaegh <i>et al.</i> , 2009:1437    | Iran                | 1614 males; 2006 females   | 94.5   | 94.5   |
| Kim <i>et al.</i> , 2009:35          | Korea               | 18551 males; 12525 females | 77 (Hyperten.)<br>77 (Diabetes)<br>76 (MetS) | 84 (Hyperten.)<br>85 (Diabetes)<br>83 (MetS) |
| Ye <i>et al.</i> , 2009:1058         | China               | 114 males; 176 females     | 82   | 88   |
| Ogawa <i>et al.</i> , 2010:117       | Japan               | 3811 males; 2161 females   | 80   | 84   |
| Park <i>et al.</i> , 2010:511        | Korea               | 3574 males; 5243 females   | 80   | 85   |
| Prinsloo <i>et al.</i> , 2011:599    | South Africa        | 80 males, 93 females       | 98   | 90   |



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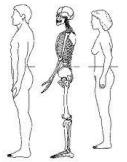
## 2.4 Hypertension and carotid intima-media thickness (CIMT)

Essential hypertension is frequently associated with metabolic abnormalities. These abnormalities include obesity of the central, visceral subtype, insulin resistance with or without impaired glucose tolerance and dyslipidemia, consisting of elevated very low density lipoproteins and low levels of high density lipoproteins - now widely known as the Metabolic Syndrome (Björntorp, 2002:82).

Previous studies found that the African population is a high-risk group regarding the prevalence of hypertension compared to the Caucasian population (Van Rooyen *et al.*, 2000:779; Opie & Seedat, 2005:3652; Dennison *et al.*, 2007:484; Thorogood *et al.*, 2007:326; Sliwa *et al.*, 2008:915; Malan *et al.*, 2012:542). Opie and Seedat (2005) and Seedat (2009:39) additionally proposed that sodium sensitivity be held accountable for the very high prevalence of hypertension observed in African men.

Joffe *et al.* (1992:460), on the other hand, proposed that the African population either inherit or acquire decreased pancreatic  $\beta$ -cell mass and that they are more prone to the development of insulinopenic non-insulin dependent diabetes mellitus. Addo *et al.* (2007:1016) concurred and added that urban populations had a consistently higher prevalence of hypertension compared to that of rural areas, implicating differences in lifestyle. Thus, higher levels of obesity and increased salt and fat intake from consuming more processed foods and engaging in jobs with minimal physical activity were the most likely explanations according to Addo *et al.* (2007:1016). On the upside, the African population is prone to have favourable lipid profiles (low serum total cholesterol and high ratio of HDL cholesterol) as well as genetically determined low homocysteine levels that could protect them from ischaemic heart disease (Vorster, 2002:243).

Primary hypertension may also be a cause of long-term activation of the sympathetic nervous system (Björntorp, 2001:73; Van Lill *et al.*, 2011:355; Hamer & Malan, 2010:76 Malan *et al.*, 2012:546). Stress initially activates the HPA axis and the sympathetic nervous system where central obesity and insulin resistance (metabolic syndrome) may be a consequence of HPA axis activation (Björntorp, 2002:83), whilst higher central



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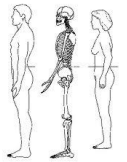
nervous system activity was associated with hypertension (Van Lill, 2011:355; Malan *et al.*, 2012:546).

Yanai *et al.* (2008) proposed visceral obesity, insulin resistance, oxidative stress, endothelial dysfunction, activated renin-angiotensin system, increased inflammatory mediators, and obstructive sleep apnoea to be possible factors to develop hypertension in the metabolic syndrome. These factors may induce sympathetic overactivity, vasoconstriction, increased intravascular fluid, and decreased vasodilatation, leading to development of hypertension in the metabolic syndrome (Yanai *et al.*, 2008).

Risk factors such as cardiovascular disease and co-morbidities such as diabetes, dyslipidaemia, and hypertension have been found to be associated with clearly altered carotid intima-media thickness (CIMT) (Kotliar *et al.*, 2008:61). CIMT assessment allows evaluation of structural changes of the vessel in a particular site of interest, the intimal space, where the atherosclerotic process is initiated. In addition, the degree of its alteration increases with the number of coexisting risk factors and the time of exposure, especially in the presence of metabolic syndrome, which includes central obesity (Kotliar *et al.*, 2008:61). Consequently, CIMT is increasingly used for risk stratification in individuals and as an end point in intervention studies (Lorenz *et al.*, 2007:459).

CIMT is measured via the Rudy Meijer protocol, also known as the Meier Carotid Arc, where the arc is designed to assist and guide the sonographer through a single or multi-angle IMT scan protocol of the carotid artery. When used appropriately, this approach will give a set of reproducible CIMT images, over time, from specific segments of the carotid artery (Liang *et al.*, 2000:127., 2000; Meijer, 2008)

The CIMT measurement method, after a long period of refinement, has been tested in numerous population-based studies and several intervention trials. Lorenz *et al.* (2007:459) also confirmed in a meta-analysis that CIMT is a strong predictor of future vascular events. Hence, CIMT measurement is an excellent tool to detect preclinical vascular disease (Kotliar *et al.*, 2008:61).



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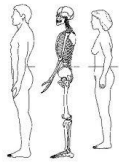
## 2.5 Psychological distress and perception of health

Mortality rates from cardiovascular disease indicate that stroke may be a feature of urbanization of black South Africans (Vorster, 2002:243, Thorogood *et al.*, 2007:325, Connor *et al.*, 2007:269). African urbanization is associated with inevitable stress, dietary changes, and acculturation (Opie & Seedat, 2005:3565, Malan *et al.*, 2008:323, Rose & Bond, 2008:268; Malan *et al.*, 2010:183, Malan *et al.*, 2012:542).

Few studies found that the more urbanized African communities were, the higher the rate of obesity and the less prudent their diets became (Vorster *et al.*, 2000:511, Puoane, 2002:1038). Urbanization could also, with accompanying insecurities and disruption in African social relationships, contribute to experiencing poorer health or distress (Malan *et al.*, 2006:305). In the second national victims of crime survey in South Africa, Burton *et al.* (2004:46) found that feelings of safety have declined markedly since 1998 – adding to chronic stress and decreased mental health (Clark *et al.*, 2007:22).

The importance of acute mental stress as a trigger for cardiac catastrophes (acute myocardial infarction, sudden death) and of depressive illness as a cause of coronary heart disease, is now firmly established (Brunner *et al.*, 2002:2659; Esler *et al.*, 2008:175). Overgaard *et al.* (2004:1072) found that psychological overload is associated with weight gain. Under stressful circumstances, the hypothalamus secretes corticotrophin-releasing hormone (CRH), which stimulates the release of adrenocorticotrophic hormone (ACTH) and subsequently, cortisol (Gudielka *et al.*, 2006). Bjorntorp (2001) revealed that cortisol binds to glucocorticoid receptors, which have a particularly high density in visceral fat depots – especially intra-abdominal fat depots - leading to accumulation of fat in this area. Consequently, the stress of coping with urbanisation may be linked to increased waist circumference and the progression of the metabolic syndrome (Bjorntorp, 2001).

In addition, Rose and Bond (2008:268) found in a younger cohort that both life event stress and perceived stress were consistently associated with substance abuse. Conversely, Rose and Bond (2008:268) revealed that coping ability and mastery offered some protection from substance abuse. Sjögren and Samsonowitz (1985) concur and



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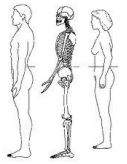
observed that alcohol abuse may be utilized as a coping strategy in the African male. Crutzen *et al.* (2010:44), in contrast, found that stress was not related to alcohol use or alcohol-related problems, and neither did drinking motives moderate the relationship between stress and alcohol use. Psychosocial stress associated with living in urban areas with high income disparity have, however, been associated with greater inter-individual tension and likelihood of inter-personal violence; both associated with increased substance use and misuse (Galea *et al.*, 2005:130).

High scores of psychic and somatic anxiety, tension, irritability and feelings of guilt, as measured by the Karolinska Scales of Personality, were found in women with resolved alcohol use disorders who had received treatment (Ostlund *et al.*, 2007:24). Shapira and Courbasson (2011) also found that depression, anxiety and low self-esteem are frequently associated with substance use disorders.

Hamer *et al.* (2011:237) demonstrated an emerging burden of disease among urban black Africans in South Africa, a phenomenon largely explained by transition from traditional African lifestyles to more westernized behaviour. In comparison with Caucasian counterparts, an excess burden of disease was seen in black Africans demonstrated by increased smoking and alcohol abuse (serum gamma glutamyl transferase) (GGT) (Hamer *et al.*, 2011:237).

Serum GGT has long been used as a conventional biomarker of liver function and a marker of excessive alcohol use (Whitfield, 2007:1, Ozer *et al.*, 2008:197) and is determined by several factors: alcohol intake, body fat content, plasma lipid/lipoproteins, glucose levels, and various medications (Grundy, 2007:4, Whitfield, 2007:1). GGT appears to be largely a reflection of ectopic liver fat or secondary hepatic inflammation (Grundy, 2007:5). Alatalo *et al.* (2008:1097) expand by adding that the effect of moderate alcohol consumption on liver enzymes (including GGT) increases with increasing BMI. Breitling *et al.* (2009:802) also found an interaction between smoking and alcohol consumption as determinants of elevated GGT levels, especially in men.

Another social consequence of living in an urban environment is an assimilation of Western cultural norms regarding body shape, in stark contrast to the traditional African



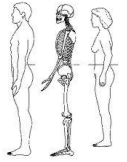
female body shape (Senekal *et al.*, 2001). Traditionally, the ideal body image among Africans has always been inclined towards a larger, fuller body shape. Also, in African religious and cultural symbolism, female body fatness is closely linked to fertility (Gordon 2001:9-10). However, a sway from the traditional African perception of body size to westernized ideals have also been reported elsewhere (Senekal *et al.*, 2001; Morris, 2008).

Stress experienced during acculturation has been identified as a causative factor for eating disorders, where acculturation is explained as the simultaneous 'push and pull' of opposing pressures to acculturate to a new culture and pressures against this acculturation from the culture of origin (Morris, 2008). Acculturation may consequently be the motivation for the rejection of traditional body size norms and accepting westernized ideals. Acculturative stress due to opposing cultures is manifested in uncertainty, anxiety and depression (Sam & Berry, 2010:473-474), although different individuals of the same culture have different approaches to acculturation.

The African notion of 'community' as embodied in the idea of 'Ubuntu', implies that a person is defined in the context of social bonds and cultural traditions rather than individual traits (Mabovula, 2011:38). Acculturative stress would surely be a factor where the African culture is in direct opposition to the westernized culture of individualism where "thin is in"; consequently emphasizing the uniqueness of the African female and possible psychological distress (Malan *et al.*, 2008:323).

## 2.6 Conclusion

Fezeu *et al.* (2007:70) investigated prevalence of the metabolic syndrome in a sub-Saharan African setting and found that central obesity may be a key determinant. In support of this hypothesis, greater visceral adiposity has been found as an increased risk for hypertension in Japanese Americans (Hayashi *et al.*, 2004:992). Hypertension was also a major risk factor in people of African descent, regardless of country of residence (Agyemang & Bhopal, 2003; Van der Merwe & Pepper, 2006; Du Plessis *et al.*, 2010, Hamer & Malan, 2010:76, Malan *et al.*, 2012:543).

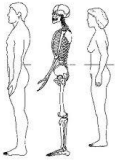


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Urbanization and accompanying psychological stress, have also been associated with increased substance use and misuse (Galea *et al.*, 2005:130). Of note, is that South Africa has one of the higher levels of consumption of alcohol, about 6 litres of alcohol per person per year, leading to very serious health and social consequences (Saxena, 2011).

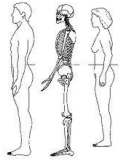
Acculturative stress in general has been manifested in uncertainty, anxiety and depression (Sam & Berry, 2010:473-474). Adding the African notion of 'Ubuntu', which is directly opposing the western individualism, acculturative stress would surely lessen the coping ability (Malan *et al.*, 2012:543) and add to the psychological distress (De Kock *et al.*, 2012) of the African people.

In conclusion, African urbanization is associated with inevitable stress and subsequent substance abuse (Opie & Seedat, 2005:3565, Malan *et al.*, 2008:323, Rose & Bond, 2008:268; Malan *et al.*, 2010:183, Hamer *et al.* 2011:237; Malan *et al.*, 2012:542). Björntorp (2001:83) proposed that stress causes central obesity, whilst evidence now indicates that central obesity plays a central role in the development of the metabolic syndrome and appears to precede the appearance of other metabolic syndrome components linked to insulin resistance (Fezeu *et al.*, 2007:70; Cameron *et al.*, 2008:2707; Després *et al.*, 2008:1039). Cardiovascular disease and co-morbidities such as diabetes, dyslipidaemia, and hypertension have been found to be associated with clearly altered carotid intima-media thickness (CIMT) (Kotliar *et al.*, 2008:61). As a result, CIMT is increasingly used for risk stratification in individuals and as an end point in intervention studies (Lorenz *et al.*, 2007:459).



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The strength of this study lies in the current void in literature where population- and country-specific guidelines for WC have not yet been appropriated for sub-Saharan Africans; also the association between obesity and mental health have not been investigated in the black South African population. Subsequently, the research questions that was aimed to answer by means of this study are: (1) Can new population-specific WC cut off points be established in a cohort of urban African teachers? (2) Which WC cut off point (JSC or new proposed)(NPM) is best associated with structural vascular disease, as indicated by carotid intima-media thickness (CIMT)? and (3) Which of the two WC cut offs will be best associated with perception of own health in this African cohort?



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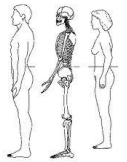
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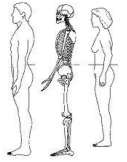
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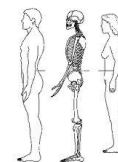
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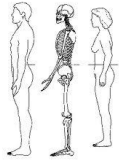
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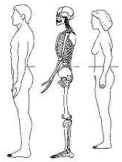
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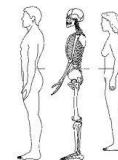
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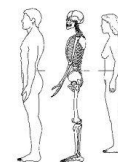
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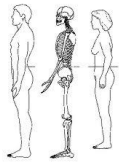
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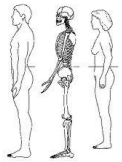
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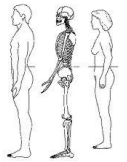
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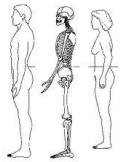
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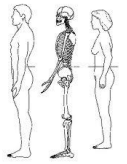
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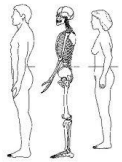
Chapter  
3**Determining the Waist Circumference Cut  
off which Best Predicts the Metabolic  
Syndrome Components in Urban Africans:  
The SABPA Study**

J. Prinsloo, L. Malan, J.H. de Ridder, J. Potgieter and H.S. Steyn  
*Experimental and Clinical Endocrinology and Diabetes*, 199:599-603.

**Abstract:**

Various studies have shown that the relationship between waist circumference (WC) and abdominal obesity is age, gender as well as ethnicity dependent. WC criteria for Sub Saharan Africans have not been defined by the International Diabetes Federation (IDF). The aim was to determine which WC cut off best predicted Metabolic Syndrome (MetS) in a group of urban African teachers (80 males and 93 females). We determined sphygmomanometer blood pressure, WC, glucose, high density lipoprotein cholesterol (HDL) and triglyceride (TRIG) values. The males' MetS profile was less favourable as their glucose, TRIG and blood pressure levels were higher than the proposed cut off for MetS. The females could be classified as obese, based on their mean BMI ( $32.78 \pm 6.36$ ) and WC ( $93.48 \pm 15.68$ ). Receiver operating characteristic (ROC) WC cut off points of 90, 91, 94 and 96 cm for the respective MetS components in males (blood pressure, HDL, glucose and TRIG) were suggested. In the females, cut off points of 92, 98, 94 and 94 cm for TRIG, blood pressure, HDL and glucose respectively, were put forward. Odds ratios revealed that increased blood pressure best predicted ROC WC in both males (OR 9.59; 95 % CI 3.14–29.32) and females (OR 3.11; 95 % CI 1.30–7.42) irrespective of age. We suggest that the optimal cut off point for the males be set at 90 cm, as opposed to the current 94 cm; whilst the female cut off be set at 98 cm as opposed to the existing cut off of 80 cm. Larger sample groups are recommended to justify our data.

**Keywords:** [metabolic syndrome; central obesity; Setswana; black Africans; ROC]



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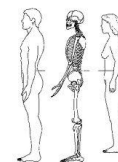
## Introduction

Body fat distribution is an important risk factor for obesity-related diseases and Zhang et al. (2008) observed that anthropometric measures of abdominal adiposity were strongly and positively associated with all-cause, cardiovascular disease (CVD), and cancer mortality independently of body mass index (BMI).

A cluster of risk factors for CVD and type 2 diabetes have become known as the metabolic syndrome (MetS) (Alberti et al., 2009). The 2009 Joint Statement Consensus (JSC) risk factors include raised blood pressure (systolic BP  $\geq 130$  or diastolic BP  $\geq 85$  mm Hg), raised triglycerides (Trig) ( $\geq 1.7$  mmol/L) and lowered high-density lipoprotein cholesterol (HDL) ( $< 1.03$  mmol/L in males and  $< 1.29$  mmol/L in females), raised fasting glucose ( $\geq 5.6$  mmol/L), and central obesity (Alberti et al., 2009). Alberti et al. (2009) decided that a single set of cut points would be used for all components except waist circumference, for which further work is required. Population-specific and country-specific definitions for waist circumference (WC) are recommended, although the International Diabetes Federation (IDF) cut points are to be used for non-Europeans until more data are available (Alberti et al., 2009). Several studies have proposed new population-specific cut points (Okusun et al., 2000; Berber et al., 2001; Sumner et al., 2008).

Conversely, evidence now indicates that central obesity plays a central role in the development of the MetS and appears to precede the appearance of other metabolic syndrome components linked to insulin resistance (Cameron et al., 2008; Després et al., 2008).

However, various studies have shown that the relationship of WC to abdominal obesity is age-, gender, as well as ethnicity-dependent (Han et al., 1997; Després et al., 2000; Misra et al., 2005). Due to a distinct lack of information for Sub-Saharan Africans we aimed to investigate new population specific WC cut off points best correlating with different MetS components in a cohort of urban African teachers.



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## Materials and Methods

Our sub-study formed part of the Sympathetic Activity and Ambulatory Blood Pressure in Africans (SABPA) study. The study was a multidisciplinary target-population study, conducted from February to May 2008 avoiding seasonal changes. Permission to participate was granted by the North-West Department of Education and the South African Teachers Union, as well as the Ethics Committee of the North-West University (project nr: NWU-00036-07S6). The study conformed to the ethical guidelines of the World Medical Association Declaration of Helsinki (revised 2004).

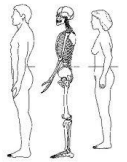
## Study population

Urban black African (hereafter referred to as African) teachers, (101 males and 99 females, aged 25-65 years) were recruited and included as participants from schools in the Dr. Kenneth Kaunda District in North-West Province.

Exclusion criteria for this study included ear temperature  $>37^{\circ}\text{C}$ , alpha- or beta blockers usage, donated blood or vaccinated in the previous 3 months. We also excluded users of diabetic (N=7) and anti-depressant medication (N=1) as well as HIV positive status (N=19) participants in our sub-study.

## Experimental procedure

Each morning of the working week the ambulatory blood pressure (Cardiotens®, Meditech, Budapest, Hungary) as well as Actical® accelerometers (Montréal, Québec) apparatuses were fitted to four participants. Suitable blood pressure (BP) cuff sizes were fastened to each subjects' non-dominant arm. The procedure was explained to the participants in order to obtain successful inflation rates. The Cardiotens® apparatus is validated by the British Hypertension Society (BHS) and measures ambulatory BP oscillometrically at intervals of 30 minutes during the day and every 60 minutes at night (Kohara et al., 1995). The Cardiotens® and Actical® software programmes were activated and participants resumed their daily activities. The participants also received an ambulatory diary card in which they reported any symptoms that may have occurred during the measuring period such as dizziness, fatigue, light physical activity, visual disturbances and stress.



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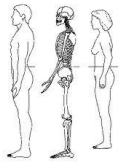
At 16:30 they were transported to the Metabolic Unit Research Facility of the North-West University for an overnight stay, where participants were welcomed and introduced to the experimental setup to lessen anticipation for the stress test. At 18:00 they received a standardized dinner. Last beverages were given at 20:30.

After an overnight fast and the last blood pressure reading at 06:00, the Cardiotens® and Actical apparatuses were disconnected. The ambulatory BP data were downloaded and analysed with the Cardiovisions 1.15 software program (Meditech, Budapest, Hungary). The anthropometric measurements were taken in triplicate by qualified anthropometrists according to International Society for the Advancement of Kinanthropometry (ISAK) standards (Marfell-Jones et al., 2006). Participants rested for 5 minutes in a semi-recumbent position before the first measurement was taken. BP was measured with a sphygmomanometer using the Riva-Rocci/Korotkoff method on the non-dominant arm. Two duplicate measures were taken with a 3 minute resting period between each measurement and the last measurement was used for the non-parametric receiver operating characteristic (ROC) curve (Zweig and Campbell, 1993). Fasting blood samples (70 ml) were obtained by a registered nurse.

### **Assessment of anthropometric and biological variables**

All anthropometric measurements were performed by ISAK level 2 accredited anthropometrists, with subjects in minimal clothing and without shoes. The waist circumference was taken in triplicate at the end of normal expiration at the narrowest point of the abdomen between the lower costal (10<sup>th</sup> rib) border and the top of the iliac crest, perpendicular to the long axis of the trunk (Marfell-Jones et al., 2006). The subject assumed a relaxed standing position with the arms folded across the thorax (Marfell-Jones et al., 2006).

BMI (kg/m<sup>2</sup>) was calculated. The body weight measured by a KRUPS scale with the participant wearing minimal clothes and with the weight evenly distributed, to the nearest 0.1 kg (Marfell-Jones et al., 2006). Height was measured with a stadiometer to the nearest 0.1cm while the participant's head was in the Frankfort plane, the heels together and the buttocks and upper back touching the stadiometer (Marfell-Jones et al., 2006).



Fasting blood samples were obtained with a winged infusion set from the brachial vein branches from the right arm by a registered nurse. Serum glucose, HDL and Trig were handled according to standardised procedures and stored at  $-80^{\circ}\text{C}$ . Analyses were done with the Konelab<sup>TM</sup> 20i sequential multiple analyser computer (SMAC) (Thermo Scientific, Vantaa, Finland) and Unicel DXC 800 (Beckman and Coulter, Germany) at accredited independent laboratories.

### Statistical analyses

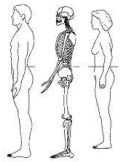
Data analyses were performed with SPSS, v17 for Windows. Firstly, gender-specific descriptive statistics (including mean and standard deviation) were done for the Africans. Thereafter, a non-parametric receiver operating characteristic (ROC) curve (Zweig and Campbell, 1993), was computed together with the area under this curve (AUC) to explore the association between waist circumference and the different MetS components using different cut off points. Sensitivity and specificity values were computed to determine the cut off point that would maximize the sum of the number of true positive and true negative predictions. Subjects were grouped according to their suggested cut off points for all MetS components (glucose, HDL, triglycerides and blood pressure). Sensitivity and specificity values and also odds ratios (OR's) with 95% Confidence Intervals (CI's) were calculated to highlight the odds of each component predicting pathological waist circumference.

### Sensitivity analyses

Sensitivity analyses, addressing age as a known important covariate for WC and BP (Alberti et al., 2009), was calculated and tested whether age influences the discriminatory power of the ROC WC cut off point (Faraggi, 2003).

### Results

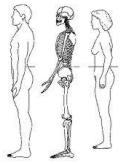
Initially, gender-specific descriptive analyses were done for the Africans (including mean and standard deviation) (**Table 1**). The mean ambulatory blood pressure successful inflation rate for males in the 22-23h period was 82.75 % ( $\pm 9.7$ ) and for females 74.59 % ( $\pm 14.88$ ). These values are in accordance to the European Hypertension Society required level of > 70% in a 24h period.

**Table 1: Baseline characteristics of African males and females.**

|   | <b>Males (N = 81)</b> | <b>Females (N = 90)</b> |
|---|-----------------------|-------------------------|
| Age, yrs                                      | 42.64 ± 8.34          | 45.32 ± 8.10            |
| <b>Lifestyle and Anthropometric variables</b> |                       |                         |
| BMI, kg/m <sup>2</sup>                        | 27.67 ± 5.94          | 32.78 ± 7.29            |
| Physical activity, kcal/h                     | 2723.28 ± 823.31      | 2665.83 ± 800.83        |
| GGT, U/L                                      | 76.28 ± 72.51         | 48.06 ± 69.75           |
| Cotinine, ng/ml                               | 24.06 ± 47.60         | 18.06 ± 55.25           |
| <b>Metabolic Syndrome Components</b>          |                       |                         |
| Glucose, mmol/L                               | 6.01 ± 2.12           | 5.03 ± 1.10             |
| HDL, mmol/L                                   | 1.08 ± 0.38           | 1.21 ± 0.31             |
| Triglycerides, mmol/L                         | 1.81 ± 1.70           | 0.99 ± 0.54             |
| Ambulatory SBP, mmHg                          | 138.00 ± 16.91        | 128.60 ± 15.17          |
| Ambulatory DBP, mmHg                          | 87.94 ± 11.43         | 78.87 ± 8.98            |
| Sphygmomanometer SBP, mmHg                    | 140 ± 21              | 130 ± 17                |
| Sphygmomanometer DBP, mmHg                    | 94 ± 15               | 84 ± 10                 |
| Waist circumference, cm                       | 93.49 ± 16.41         | 93.48 ± 15.68           |

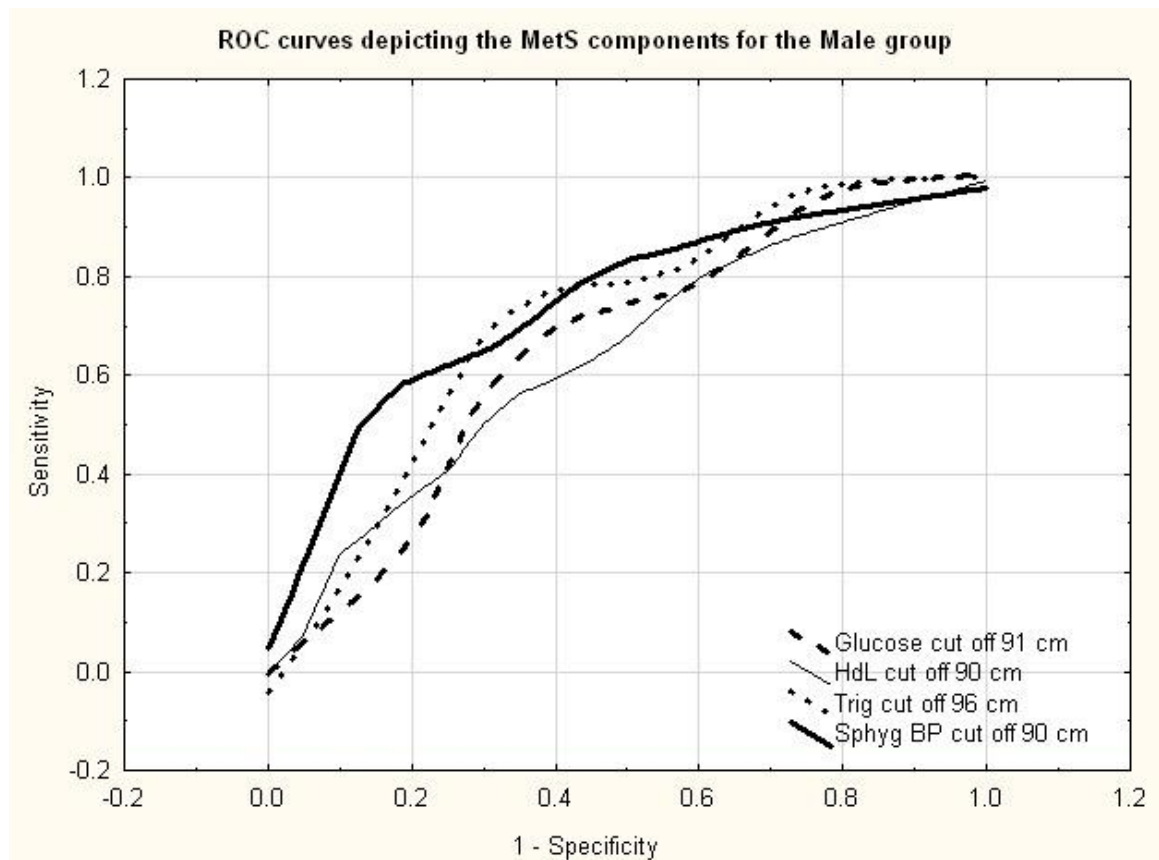
**Table 1 caption:**

Values are arithmetic mean ± SD. Where; BMI, body mass index; GGT, Gamma Glutamyl Transferase; HDL, High Density Lipoprotein; SBP, Systolic Blood Pressure; DBP, Diastolic Blood Pressure.

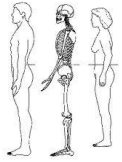


In **Table 1**, the males' MetS profile was less favourable as their glucose levels, Trig, **ambulatory and sphygmomanometer blood pressure** were higher than the proposed cut off for MetS. The females could be classified as obese, based on their mean body mass index ( $32.78 \pm 6.36$ ) and WC ( $93.48 \pm 15.68$ ). Despite their level of obesity, Trig, **ambulatory and sphygmomanometer blood pressure** and plasma glucose were within the normal range. Even though the current WC cut off for females is set at 80cm (Alberti et al., 2009), this apparently healthy group of females had a mean WC of 93cm.

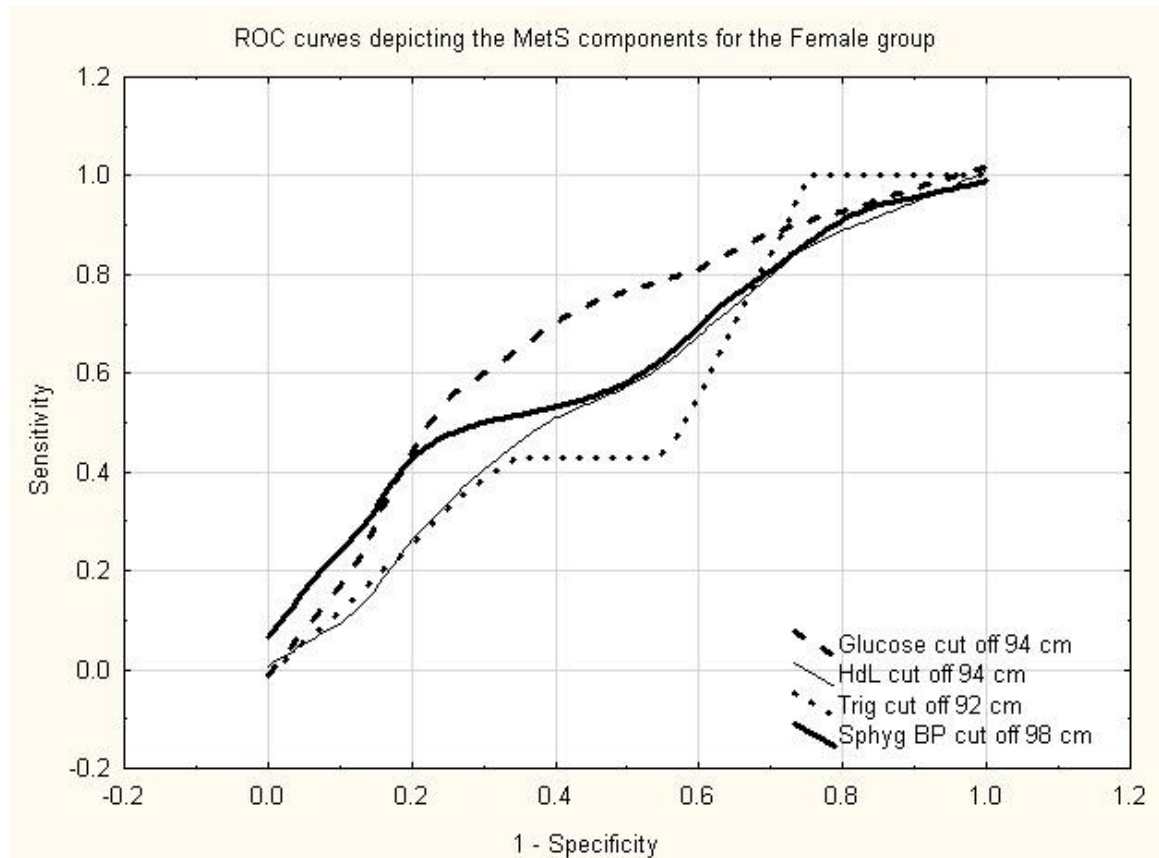
ROC curves were used to determine the suggested cut off value for the WC by using increased glucose, HDL, Trig and hypertension as reference of morbidity in the African male (**Figure 1**) and female (**Figure 2**) group. The respective ROC cut off values, yielding maximum sensitivity and specificity ranged between 90 - 96cm in males and between 92 - 98 cm in females.



**Figure 1: ROC curves depicting the MetS components for the Male group: Glucose, HDL, Trig and BP in predicting pathological WC. The area under the curve (AUC)**

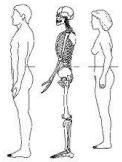


(95%CI) was 0.66 (0.54; 0.78) for Glucose, 0.64 (0.50; 0.78) for HDL, 0.71 (0.60; 0.83) for Trig and 0.75(0.62; 0.88) for Sphyg BP.



**Figure 2: ROC curves depicting the MetS components for the Female group: Glucose, HDL, Trig and BP.** The area under the curve (AUC) (95%CI) was 0.68 (0.53; 0.83) for Glucose, 0.57 (0.44; 0.70) for HDL, 0.55 (0.35; 0.75) for Trig and 0.62 (0.51; 0.74) for Sphyg BP.

Odds ratios were calculated in **Table 2** in order to obtain the best MetS predictor for pathological WC. It was found that increased blood pressure best predicted WC pathology (OR of 9.50 with 95% CI 3.14 to 29.32) according to the suggested ROC cut off. This means that the odds of the high WC group to the low WC group within the high blood pressure group is 9.5 times larger than that obtained in the low blood pressure group, but can be as low as 3.23 and as high as 29.32 with high probability. Glucose (OR of 3.23 with 95% CI 1.27 to 8.24) and Trig (OR of 4.41 with 95% CI of 1.68 to 11.92) predicted pathological WC in males. In females, only increased blood pressure had a



higher odds of 3.11 (95% CI 1.30 to 7.42) predicting WC pathology. Sensitivity analyses demonstrated weak associations between age and WC for the ROC BP cut off points in all gender groups.

**Table 2: Odds ratios with WC ROC cut off as dependent variable for each of the Metabolic Syndrome components.**

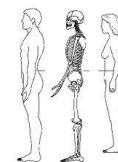
|                         | Waist circumference |             |      |                  |            |      |
|-------------------------|---------------------|-------------|------|------------------|------------|------|
|                         | Males (N = 81)      |             |      | Females (N = 90) |            |      |
|                         | OR                  | ± 95 CI     | p    | OR               | ± 95 CI    | p    |
| <b>Glucose</b>          | 3.23                | 1.27; 8.24  | 0.01 | 2.80             | 0.84; 9.39 | 0.10 |
| <b>HDL</b>              | 1.70                | 0.64; 4.87  | 0.28 | 1.61             | 0.66; 3.93 | 0.30 |
| <b>Triglycerides</b>    | 4.41                | 1.68; 11.92 | 0.00 | 0.63             | 0.13; 3.00 | 0.56 |
| <b>Sphygmomanometer</b> | 9.50                | 3.14; 29.32 | 0.00 | 3.11             | 1.30; 7.42 | 0.01 |
| <b>BP</b>               |                     |             |      |                  |            |      |

**Table 2 caption:**

Data presented as odds ratio (OR) with 95% Confidence Interval and p-values for significance of OR. Where; HDL, High Density Lipoprotein.

**Discussion**

The aim of this study was to determine a WC cut off point which will best predict the MetS components in a cohort of urban African male and females. We have demonstrated that the suggested ROC WC cut off for the particular MetS components (blood pressure, HDL, glucose and Trig) in males varied between 90-96 cm whereas the females' cut offs varied between 92-98 cm. Currently, we carefully want to suggest a cut off for males at 90 cm, and recommend a WC cut off for females at 98 cm. More research however is needed to verify our findings as we obtained data from a small sample group comprising 101 African males and 99 females.



Even though the current WC cut off for females is set at 80 cm (Alberti et al., 2009), this apparently healthy group of females has a mean WC of 93 cm. This might put forward the hypothesis of metabolically healthy obese females as first put forward by Walker et al. (1989). Björntorp (1997) established that centralisation of body fat is associated with the complications of obesity. Central obesity has been identified as the culprit that contributes to the development of glucose intolerance and hyperinsulinaemia in obesity (Després et al., 2001). Van der Merwe and Pepper (2006) found that, when compared to white urban women, matched for BMI and body composition, black women have less central obesity, which may account for the proposed higher WC cut off of 98 cm in this group of females.

The OR's highlighted the raised BP values as the variable with the highest odds of presenting with pathological WC in both the males ( $9.50 \pm 3.14$ ; 29.32) and the females ( $3.11 \pm 1.30$ ; 7.42). Blood pressure is a major risk factor in people of African descent, regardless of country of residence (Cappuccio et al., 1997; Wild and Mckeigue, 1997; Agyemang and Bhopal, 2003). Other South African studies corroborated these findings (Van der Merwe and Pepper, 2006; Vorster, 2002; Du Plessis et al., 2010; Sliwa et al., 2008). Due to the impact of blood pressure on disease risk in this ethnic group, we suggest that the WC cut off be considered to correspond with 90 cm in males and 98 cm in females.

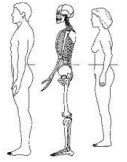
Glucose, Trig and blood pressure levels in the males were all higher than those proposed by the Joint Statement (Alberti et al., 2009). Calculated odds ratio strengthen these findings and glucose ( $3.23 \pm 1.27$ ; 8.24 ), as well as Trig ( $4.41 \pm 1.68$ ; 11.92) should both support blood pressure as contributors presenting with pathological WC in males. Conversely, raised glucose, Trig and blood pressure, together with substance abuse (both alcohol and smoking), present in the males (Table 1), are major risk factors for cardiovascular disease and stroke (Kannel and Mcgee, 1979; Ezzati et al., 2003; Connor et al., 2005; Kudielka et al., 2006). Duckrow et al. (1987) found that both chronic and acute raised glucose levels, are associated with decreased regional cerebral blood flow and the mechanism for this effect does not appear to adapt to chronic hyperglycemia.



Mostly, stroke incidence in developing countries, such as South Africa, is likely accounted for by the adoption of a more urbanised lifestyle (Vorster et al., 2005; Brainin et al., 2007; Astrup et al., 2008). Coping with a stressful environment revealed increased abdominal obesity, hypertension prevalence, vascular responsiveness and glucose levels in African males (Vorster et al., 2000; Malan et al., 2006; Malan et al., 2008). The burden of chronic stress and accompanying changes in personal behaviours (smoking, over-eating, drinking, disturbed sleeping patterns; otherwise referred to as “lifestyle”) have been associated with the concept of allostatic overload (McEwen, 2008). Chronic allostatic load produce a chronic wear and tear on the cardiovascular system that can result, over time, in disorders such as stroke and heart attacks (McEwen, 2008). Under stressful circumstances, the hypothalamus secretes corticotrophin-releasing hormone (CRH), which stimulates the release of adrenocorticotrophic hormone (ACTH) and subsequently, cortisol (Gudielka et al., 2006). Bjorntorp (2001) revealed that cortisol binds to glucocorticoid receptors, which have a particularly high density in visceral fat depots – especially intra-abdominal fat depots - leading to accumulation of fat in this area. Consequently, the stress of coping with urbanisation may be linked to increased waist circumference and the progression of the metabolic syndrome (Bjorntorp, 2001).

One limitation of our study was the use of cross-sectional data and therefore causality cannot be inferred. Another limitation was that the study sample was not selected from the whole African population. The participants included Setswana-speaking Africans, only one of the eleven ethnic groups residing in South Africa and it is recommended that the findings be verified in other African communities. Increasing sample sizes will strengthen recommended WC cut off points for age, gender- and ethnicity specific groups.

In conclusion, population-specific and country-specific definitions for waist circumference (WC) are recommended. Currently, it is suggested that the optimal cut off point for the males be set at 90 cm, as opposed to the current 94 cm. In addition, it is proposed that the female cut off be set at 98 cm as opposed to the existing cut off of 80 cm. If taken into account the less favourable MetS profile of the African males and the high OR of



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blood pressure, glucose and triglycerides predicting WC, earlier screening could lessen cardiometabolic morbidity and mortality.

### **Acknowledgements**

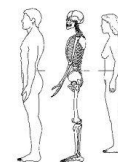
The authors gratefully acknowledge the assistance of all members of the SABPA research team, especially C Lessing (RN) and S Péter (MD), as well as the participants.

### **Disclosure**

No conflict of interest

### **Author Contribution**

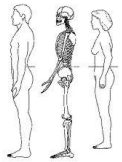
Prof L Malan (principal investigator); Me J Prinsloo, Prof JH de Ridder, HS Steyn and Prof JC Potgieter (staff members) (Potchefstroom). JP planned, analysed the data, wrote and edited the manuscript; LM planned, analysed the data and edited the manuscript; JHdR and JCP edited the manuscript; HSS analysed the data and edited the manuscript.



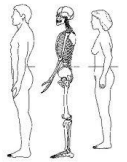
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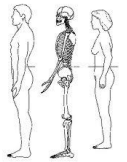
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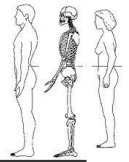
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**Chapter**  
**4****Comparing performances of two central obesity models to predict structural vascular disease by using ROC analyses, Logistic Regression and Neural Networks: the SABPA study.**

J. Botha <sup>a</sup>, J.H. de Ridder., J.C. Potgieter, H.S. Steyn, L. Malan

**Abstract:**

*Objectives:* A new proposed model (NPM) for waist circumference (WC) cut points, driven by increased blood pressure, was recently demonstrated. This study aimed to demonstrate whether the NPM can be used in an African population by comparing the NPM and the Alberti Joint Statement Consensus (JSC) model via ROC analyses, Logistic Regression (LR) and Neural Networks (NN).

*Methods:* Urban African gender groups (N=171) were stratified into the JSC model and the NPM. Ultrasound carotid intima media thickness (CIMT), blood pressure (BP) and fasting bloods (glucose, high density lipoprotein (HDL) and triglycerides) were obtained in a well-controlled setting.

*Results:* The NPM male model predicted structural vascular disease (ROC AUC: 0.65, LR ROC AUC: 0.71, NN ROC AUC: 0.71) equal to the JSC model (0.65, 0.71, 0.69). Similarly, the female JSC model (0.61, 0.82, 0.81) and NPM model (0.62, 0.84, 0.82) equally predicted structural vascular disease. In both the male NPM and the JSC (WC > 90-94 cm) the CIMT's were >1 mm, with BP in the hypertensive range.

*Conclusion:* The results indicated that overall, validation of the NPM model was comparable with the current JSC model in their prediction of CIMT, an indicator for structural vascular disease.

**Keywords:** [central obesity; anthropometry; ethnicity; ROC; Logistic Regression; Neural Networks]



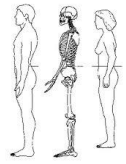
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## 1. Introduction

Obesity affects virtually all organ systems and results in a wide range of metabolic and endocrine abnormalities [1]. Obesity and insulin resistance have been implicated in the pathogenesis of the metabolic syndrome [2]. The metabolic syndrome, according to the 2009 Joint Statement Consensus (JSC) [3], includes several risk factors: raised blood pressure (systolic BP  $\geq 130$  or diastolic BP  $\geq 85$  mm Hg), raised triglycerides (Trig) ( $\geq 1.7$  mmol/L) and lowered high-density lipoprotein cholesterol (HDL) ( $< 1.03$  mmol/L in males and  $< 1.29$  mmol/L in females), raised fasting glucose ( $\geq 5.6$  mmol/L), and central obesity [3]. Alberti et al. [3] further determined that a single set of cut points would be used for all components except waist circumference. Population-specific and country-specific definitions for waist circumference (WC) have been recommended, although the International Diabetes Federation (IDF) cut points are to be used for non-Europeans until more data are available [3].

Prinsloo et al. [4] recently suggested a new population-specific cut off for a group of urban African males ( $\geq 90$  cm) and females ( $\geq 98$  cm), namely the new proposed model (NPM). The NPM recommended cut offs, especially those for the females', differed considerably from those proposed by Alberti et al [3] (males  $\geq 94$  cm; females  $\geq 80$  cm). Prinsloo et al. [4] further demonstrated that sphygmomanometer blood pressure (BP) was the variable with the highest odds of presenting with pathological WC in black African males and females alike. Kotliar et al. [5] and Redon et al. [2] suggested that hypertension is a systemic, chronic, inflammatory disease which may induce structural and functional vessel impairment with a high risk for subclinical structural vascular disease. Ultrasound carotid intima-media thickness (CIMT) is a non-invasive marker of early structural vascular disease and indicates a profile caused by multiple risk factors over time on arterial walls [6]. Hypertension, impaired fasting glucose, smoking and obesity may be important factors for atherosclerosis [5,7].

The impact of metabolic syndrome on CIMT has not yet been determined. Ethnicity-specific WC cut off points, as part of the metabolic syndrome may impact on CIMT. Therefore we needed to demonstrate whether the NPM can be utilized in an African population through valid computational analyses. The aims were to compare



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performances of the two models (JSC and NPM) via receiver operating characteristic (ROC), logistic regression analyses and multilayer perceptron neural networks to predict structural vascular disease. We propose, taking the high prevalence of hypertension among people of African descent into account, and considering that the ethnicity-specific proposed WC (NPM) is based on hypertension, that the NPM will better predict structural vascular disease in this African cohort.

## 2. Methods

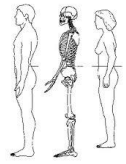
This sub-study formed part of the Sympathetic activity and Ambulatory Blood Pressure in Africans (SABPA) cohort study, conducted from February to May 2008. Permission to participate was granted by the North-West Department of Education and the South African Teachers Union, as well as the Ethics Committee of the North-West University (project nr: NWU-00036-07S6). The study conformed to the ethical guidelines of the World Medical Association Declaration of Helsinki (revised 2004). All participants signed an informed consent form.

### 2.1 *Study population and sample*

Male and female urban black African (hereafter referred to as African) school teachers, (N=200, aged 25-65 years) were recruited and included as participants from governmental organizations in the Dr. Kenneth Kaunda District in North-West Province. Exclusion criteria for this study included ear temperature  $>37.5^{\circ}\text{C}$ , alpha- or beta blockers usage, donated blood or vaccinated in the previous 3 months. Clinically confirmed diabetics (n=7), use of anti-depressant medication (n=1) and HIV-infected (n=13) participants were also excluded. The final participant sample comprised 171 Africans (n=81 males and females n=90).

### 2.2 *Experimental procedure*

On every working day of the week, for the extent of the project, the Actical® physical activity monitors (Minimitter, Quebec, Canada) apparatus were fitted at 07:00, to four



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participants for 24 hours. Software programmes were activated and participants resumed their daily activities.

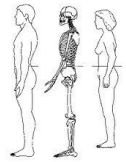
At 16:30 the participants were transported to the Metabolic Unit Research Facility of the North-West University for an overnight stay, where they were welcomed and introduced to the experimental setup. After an overnight fast, the Actical® apparatuses were disconnected at 06:00. Anthropometric measurements were taken in triplicate by anthropometrists according to the standard of the International Society for the Advancement of Kinanthropometry (ISAK) [8].

Two mercury sphygmomanometer blood pressure readings using Korotkoff IV or V for blood pressure (BP) followed while the participants rested for five minutes in a semi-recumbent position, with a three-minute rest between measurements. The second measurement was used for statistical analyses. BP measurements were taken by a medical doctor and registered nurse. Fasting blood samples were obtained by a registered nurse.

### 2.3 *Anthropometric measurements*

All anthropometric measurements were performed by ISAK level 2 accredited anthropometrists, with subjects in minimal clothing and without shoes. The waist circumference was taken in triplicate at the end of normal expiration at the narrowest point of the abdomen between the lower costal (10<sup>th</sup> rib) border and the top of the iliac crest, perpendicular to the long axis of the trunk [8]. The subject assumed a relaxed standing position with the arms folded across the thorax [8].

Body mass index (BMI) was calculated. The body weight measured via a KRUPS scale with the participant wearing minimal clothes and with the weight evenly distributed, to the nearest 0.1 kg [8]. Height was measured with a stadiometer to the nearest 0.1cm while the participant's head was in the Frankfort plane, the heels together and the buttocks and upper back touching the stadiometer [8].



#### 2.4 *Biochemical analysis*

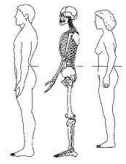
Fasting blood samples were obtained with a winged infusion set from the brachial vein branches from the dominant arm by a registered nurse. Serum gamma glutamyl transferase and cotinine were used as markers of alcohol abuse and smoking-, respectively. Sodium fluoride glucose, serum HDL and triglycerides were handled according to standardized procedures and were stored at  $-80^{\circ}\text{C}$ . Analyses were done using the Konelab<sup>TM</sup> 20i sequential multiple analyzer computer (SMAC) (Thermo Scientific, Vantaa, Finland) and Unicel DXC 800 (Beckman & Coulter, Germany) at accredited laboratories.

#### 2.5 *Carotid Intima-media thickness (CIMT)*

High resolution ultrasound carotid intima-media scan [9] determined structural vascular disease. CIMT images, from at least two optimum angles of the left and the right common carotid artery segments were acquired using Sonosite Micromaxx ultrasound system (Sonosite Inc., Bothell, WA, USA) and 6-13 MHz linear array transducer, using the Rudy Meijer protocol. The images were digitized and imported into the Artery Measurement Systems (AMS) automated software for dedicated analysis of CIMT. A maximal 10 mm segment with good image quality was chosen for analysis. The program automatically identifies the borders of the CIMT of the near and far wall. Intra-observer variability was 0.04 mm between 2 measurements made four weeks apart on 10 participants. A mean CIMT of  $>1.0$  mm was used as an estimate of high risk.

#### 2.6 *Statistical Analyses*

Data analyses were performed with Statistica version 10 [10]. In the African cohort, gender-specific descriptive statistics (including mean and standard deviation) were done in both the JSC and NPM models for the at risk (high) and normal (low) groups. Next, each model's high and low WC groups were compared by means of t-tests. Thereafter, a non-parametric receiver operating characteristic (ROC) curve [11], was computed together with the area under this curve (AUC) to explore the association between structural vascular disease, i.e., CIMT and each of the 2 WC models, separate for each



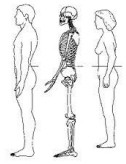
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gender group. The ROC analyses were utilized as a sensitivity analysis without confounders.

The area under the ROC curve (AUC) is an overall summary of diagnostic accuracy [12]. It is a measure of the ability to discriminate between CIMT distributions for the normal and at risk groups, using the JSC or NPM WC models. Larger values of AUC indicate a greater discriminatory ability. An AUC value of 0.5 and smaller indicates that a model is unable to distinguish between the two groups, while a value of 1.0 is obtained for complete discrimination, i.e. for perfect accuracy [12].

In ROC analyses, sensitivity is defined as True Positives / [True Positives + False Negatives], ranging from one, when all cases at risk are read as abnormal (no False Negatives), to zero, when all are called normal (no True Positives) [13]. Thus, high sensitivity corresponds to high negative predictive value [14]. Specificity on the other hand, defined as True Negatives / [True Negatives + False Positives] counters by moving from zero (all cases not at risk are normal, no False Positives) to one (all cases at risk are normal, no True Negatives) [13]. High specificity corresponds to high positive predictive value [14].

Logistic Regression (LR) analyses were then performed as it is useful for situations in which you would want to be able to predict the outcome based on values of confounders or independent variables which can be continuous, categorical or both [15]. Hosmer and Lemeshow [16] and Kleinbaum [17] add that LR assumes that measures of dependent variables are independently and randomly sampled and all independent variables in the model are relevant. Categorical CIMT, where mean CIMT  $\geq 1$  mm was classified as atherosclerosis and  $< 1$  mm as no atherosclerosis, was used in the LR analyses as dependent variable for structural vascular disease [18-19]. For each WC model (JSC and NPM), gender separate, used dichotomized WC, and a priori confounders age, BMI and physical activity were entered as independent variables. The predicted probabilities from these LR models are depicted as ROC curves, with subsequent AUC, sensitivity and -specificity values.



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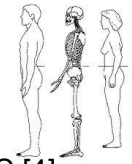
Next, Neural Networks (NN) analyses were performed in order to validate the results obtained with the logistic regression analyses. NN resemble biological neuronal systems and are used to calculate an output variable on the basis of independent input variables [20]. It learns how to transform data into a desired response, so they are widely used for pattern classification. With one or two hidden layers they can approximate virtually any input-output map, with back propagation computing the sensitivity, and update each weight proportional to the sensitivity [15].

Unlike classic statistical linear models and correlative methods, NN consists of multiple indirect interconnections between input and output variables and employ non-linear mathematical equations and statistical techniques to successively minimize the variance between actual and predicted outputs [20]. This eventually yields a model from which the predictions can be entered into ROC analyses in order to obtain AUC values, sensitivity and -specificity.

Data entered into the NN for each WC model (JSC and NPM), gender separate, used dichotomized WC and a priori confounders (age, BMI and physical activity) as independent variables, with the dichotomized CIMT as dependent variable. The data of the confounders were scaled by dividing each value by the group mean and this value was logarithmically transformed. A multilayer perceptron (MLP) model was used.

### 3. Results

Tables 1 (male) and 2 (female) describes the characteristics for the total African cohort. Lifestyle risk factors, metabolic syndrome components and mean CIMT are included in each model's respective high and low WC groups.

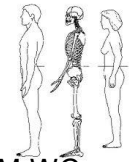


**Table 1:** Comparing males' high and low JSC [3] WC cut off and the high and low NPM WC [4].

|                                      | 2009 JSC<br>Males (N=43)<br>< 94 cm | 2009 JSC<br>Males (N=38)<br>≥ 94 cm | NPM<br>Males (N=36)<br>< 90 cm | NPM<br>Males (N=45)<br>≥ 90 cm |
|--------------------------------------|-------------------------------------|-------------------------------------|--------------------------------|--------------------------------|
| Age (years)                          | 41.37 ± 9.33                        | 44.08 ± 6.90                        | 40.50 ± 9.79                   | 44.36 ± 6.60*                  |
| <b>Lifestyle variables</b>           |                                     |                                     |                                |                                |
| Physical Activity (kCal)             | 2350.24 ± 654.08                    | 3145.41 ± 797.02*                   | 2341.32 ± 705.58               | 3028.85 ± 788.40*              |
| Gamma-glutamyl transferase (μ/L)     | 66.72 ± 86.10                       | 87.10 ± 52.21                       | 71.08 ± 93.41                  | 80.45 ± 50.78                  |
| Cotinine, ng/ml                      | 25.42 ± 48.84                       | 22.53 ± 46.76                       | 25.81 ± 46.94                  | 22.67 ± 48.61                  |
| <b>Physiological variables</b>       |                                     |                                     |                                |                                |
| Body Mass Index (kg/m <sup>2</sup> ) | 23.52 ± 3.15                        | 32.37 ± 4.73*                       | 23.06 ± 3.10                   | 31.36 ± 5.03*                  |
| Glucose (mmol/L)                     | 5.48 ± 1.38                         | 6.61 ± 2.61*                        | 5.38 ± 1.24                    | 6.52 ± 2.52*                   |
| High Density Lipoprotein (mmol/L)    | 1.21 ± 0.40                         | 0.94 ± 0.31*                        | 1.22 ± 0.42                    | 0.97 ± 0.32*                   |
| Triglycerides (mmol/L)               | 1.61 ± 2.04                         | 2.03 ± 1.17                         | 1.64 ± 2.20                    | 1.94 ± 1.16                    |
| Systolic Blood Pressure (mmHg)       | 133 ± 18                            | 147 ± 23*                           | 132 ± 16                       | 146 ± 23*                      |
| Diastolic Blood Pressure (mmHg)      | 89 ± 13                             | 99 ± 16*                            | 89 ± 12                        | 97 ± 16*                       |
| CIMT mean                            | 0.98 ± 0.17                         | 1.13 ± 0.25*                        | 0.97 ± 0.17                    | 1.11 ± 0.24*                   |

All data indicated as arithmetic mean ± standard deviation. Statistically significant difference indicated with \*,  $p \leq 0.05$

From Table 1 it is evident that for both lifestyle and physiological variables, the male high WC groups in both models demonstrated increased physical activity and CIMT, as well as higher increases in metabolic syndrome components compared to the low WC groups (glucose, high density lipoprotein, blood pressure).

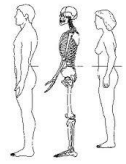


**Table 2:** Comparing females' high and low JSC [3] WC cut off and the high and low NPM WC [4].

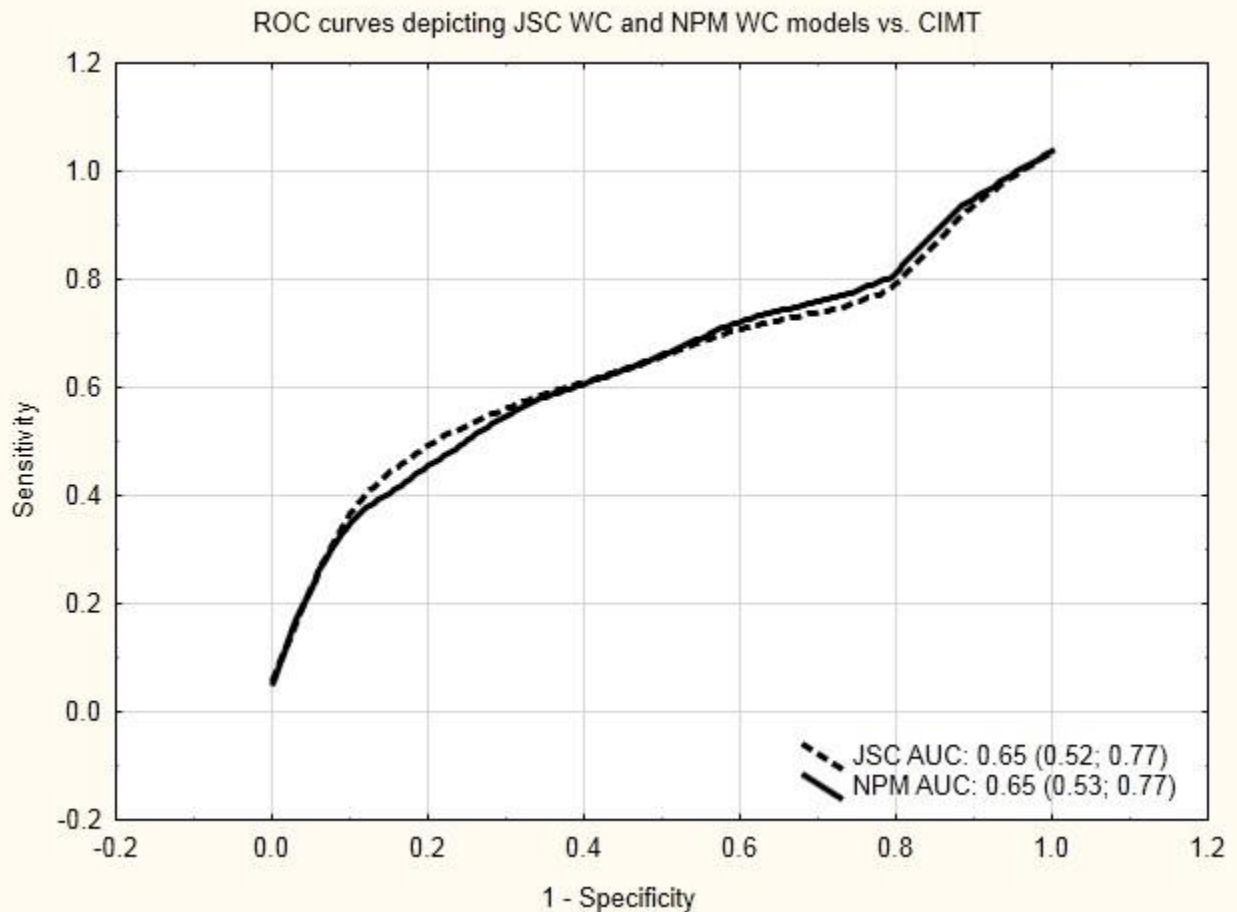
|                                      | <b>2009 JSC<br/>Females<br/>(N=17)<br/>&lt; 80 cm</b> | <b>2009 JSC<br/>Females<br/>(N=73)<br/>≥ 80 cm</b> | <b>NPM<br/>Females<br/>(N=58)<br/>&lt; 98 cm</b> | <b>NPM<br/>Females (N=32)<br/>≥ 98 cm</b> |
|--------------------------------------|---|--|--|---|
| Age (years)                          | 42.77 ± 6.54  | 45.92 ± 8.35                                       | 44.31 ± 7.28                                     | 47.16 ± 9.25                              |
| <b>Lifestyle variables</b>           |   |  |  |   |
| Physical Activity (kCal)             | 1957.67 ± 383.15                                      | 2830.74 ± 783.68*                                  | 2325.34 ± 487.42                                 | 3282.97 ± 891.26*                         |
| Gamma-glutamyl transferase (μ/L)     | 39.99 ± 44.65   | 49.71 ± 73.99                                      | 39.00 ± 35.02                                    | 63.91 ± 105.22                            |
| Cotinine, ng/ml                      | 20.31 ± 48.72   | 17.56 ± 56.90                                      | 18.26 ± 61.36                                    | 17.68 ± 42.72                             |
| <b>Physiological variables</b>       |   |  |  |   |
| Body Mass Index (kg/m <sup>2</sup> ) | 25.70 ± 3.50  | 34.43 ± 6.95*                                      | 29.26 ± 4.53                                     | 39.16 ± 7.04*                             |
| Glucose (mmol/L)                     | 4.26 ± 0.57   | 5.20 ± 1.12*                                       | 4.79 ± 0.67                                      | 5.47 ± 1.54*                              |
| High Density Lipoprotein (mmol/L)    | 1.34 ± 0.23   | 1.12 ± 0.32  | 1.25 ± 0.30                                      | 1.16 ± 0.34                               |
| Triglycerides (mmol/L)               | 0.72 ± 0.26   | 1.05 ± 0.56*                                       | 0.90 ± 0.41                                      | 1.16 ± 0.67*                              |
| Systolic Blood Pressure (mmHg)       | 121 ± 11  | 132 ± 18*  | 124 ± 14   | 140 ± 19*                                 |
| Diastolic Blood Pressure (mmHg)      | 80 ± 7  | 85 ± 11  | 81 ± 9   | 89 ± 10*                                  |
| CIMT mean                            | 0.93 ± 0.18   | 1.01 ± 0.15  | 0.98 ± 0.16                                      | 1.04 ± 0.15                               |

All data indicated as arrhythmic mean ± standard deviation. Statistically significant difference indicated with \*,  $p \leq 0.05$

Table 2 illustrates that, even though the two cut offs differ widely, each female model's high WC groups demonstrated higher physical activity and CIMT, and increased metabolic syndrome components compared to the low WC groups (glucose, triglycerides, blood pressure). The NPM distinguishes better between non-pathological and pathological diastolic blood pressure measurements than the JSC model.

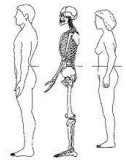


ROC curves were used as a sensitivity analysis to assess the associations between each of the 2 WC models separately and CIMT (categorical), yielding an area under the curve (AUC) with accompanying  $\pm$  95% CI, sensitivity and -specificity in the African male (Figure 1a) and female (Figure 1b) groups.

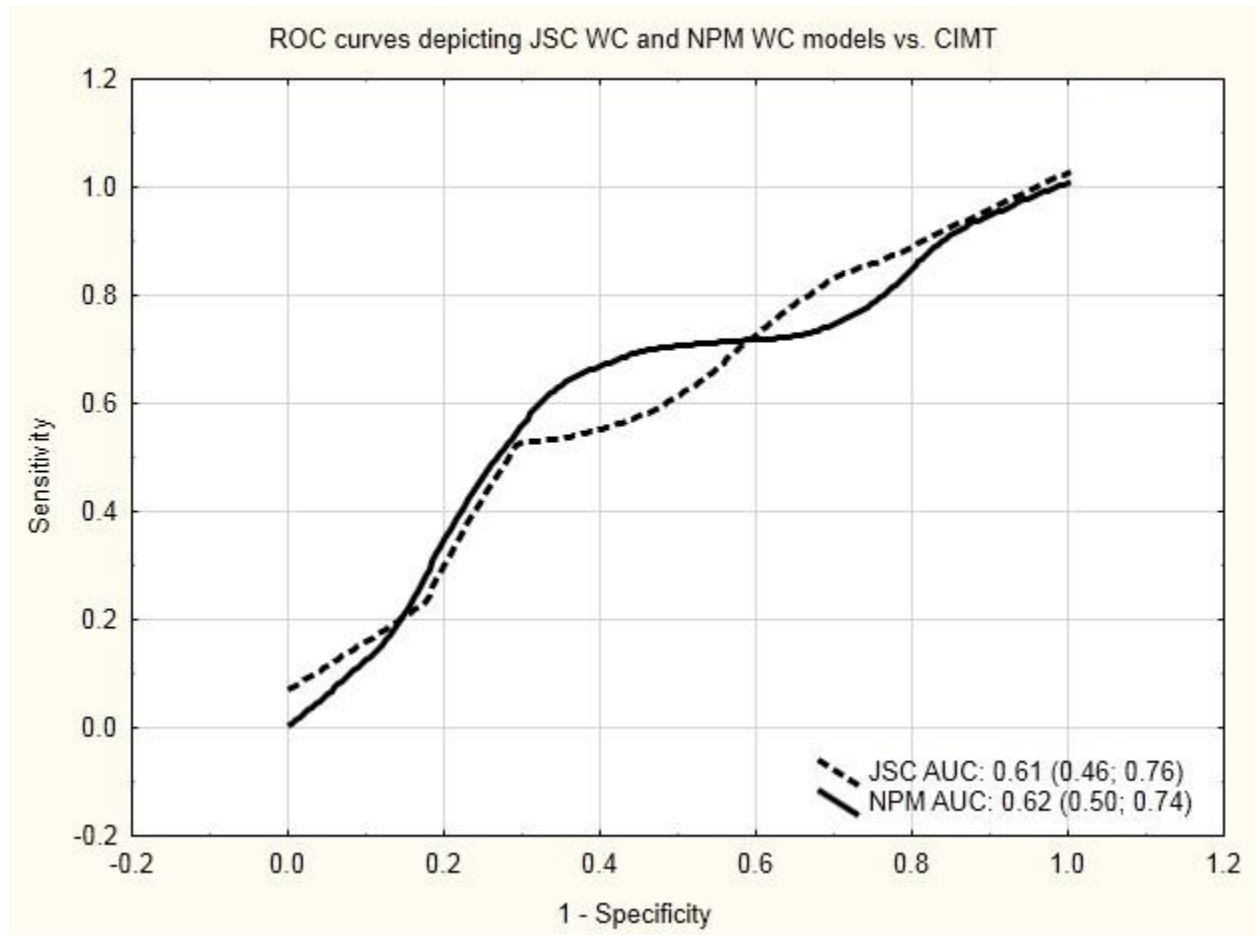


**Figure 1a:** ROC curves depicting the association between the 2 WC models (JSC and NPM) vs. CIMT for the Male group.

**Legend Figure 1a:** ROC curves depicting the 2 WC models (JSC and NPM) vs. CIMT for the Male group, AUC ( $\pm$  95% CI). The JSC model yielded a sensitivity and -specificity of 0.47 and 0.88, whilst the NPM model yielded sensitivity and -specificity values of 0.60 and 0.71.



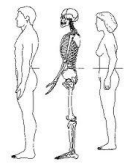
In Figure 1a, both models (both JSC and NPM) seem to equally predict structural vascular disease.



**Figure 1b:** ROC curves depicting the association between the 2 WC models (JSC and NPM) vs. CIMT for the Female group.

**Legend Figure 1b:** ROC curves depicting the association between the 2 WC models (JSC and NPM) vs. CIMT for the Female group, AUC ( $\pm$  95% CI). The JSC model yielded a sensitivity and -specificity of 0.52 and 0.77, whilst the NPM model yielded sensitivity and -specificity values of 0.69 and 0.66.

Figure 1b demonstrates that both the JSC and the NPM models equally discriminate between structural vascular disease. However, the lower limits of the CI's smaller or equal to 0.5 suggest the possibility of no distinction between the groups for both models.



Logistic regression (LR) analyses were performed for each of the 2 WC models. The analyses were done separately for each gender group and the associations between each of the 2 WC models (JSC and NPM) and categorized CIMT were assessed. Each WC model (with cut offs) and a priori confounders, age, BMI and physical activity (scaled), were entered into the model, with the dichotomized CIMT as dependent variable. The MLP NN analyses, with the same confounders as in the LR analyses, were also done for each of the models with CIMT as dependent variable. Details of the different models and their predictive validity analyses are given in the appendix.

Consequently, 1000 unique stratified samples of size 16 (8 from each group) for males and 18 (9 from each group) for females were selected and excluded from the data. The relevant model was built for each of the 1000 samples using the remaining cases and the group membership of the withheld cases (about 30% of all cases) were predicted. The sensitivity and -specificity as well as the percentage of miss-classified cases were recorded for each sample and averaged to find the cross-validation estimates of sensitivity and -specificity (Table 3).

**Table 3:** Comparison of the performance of LR and NN models.

| Model          | Sensitivity | Specificity | % Miss-classified |
|----------------|-------------|-------------|-------------------|
| LR, male JSC   | 53.6 ± 16.9 | 67.0 ± 17.2 | 39.7 ± 11.0       |
| LR, male NPM   | 53.2 ± 17.9 | 66.9 ± 17.3 | 40.0 ± 10.8       |
| LR, female JSC | 73.2 ± 15.1 | 72.2 ± 14.4 | 27.3 ± 9.7        |
| LR, female NPM | 72.6 ± 14.9 | 75.4 ± 14.7 | 26.0 ± 9.6        |
| NN, male JSC   | 59.7 ± 25.0 | 53.0 ± 25.1 | 43.7 ± 11.1       |
| NN, male NPM   | 59.3 ± 23.6 | 52.7 ± 24.1 | 44.0 ± 10.5       |
| NN, female JSC | 70.5 ± 15.7 | 72.3 ± 17.6 | 28.6 ± 10.1       |
| NN, female NPM | 67.6 ± 17.3 | 66.2 ± 17.1 | 33.1 ± 9.6        |

**Table 3 caption:** Variables indicated as mean ± Standard Error of Mean, LR, Logistic Regression, NN, Neural Networks

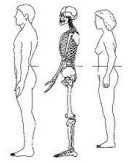
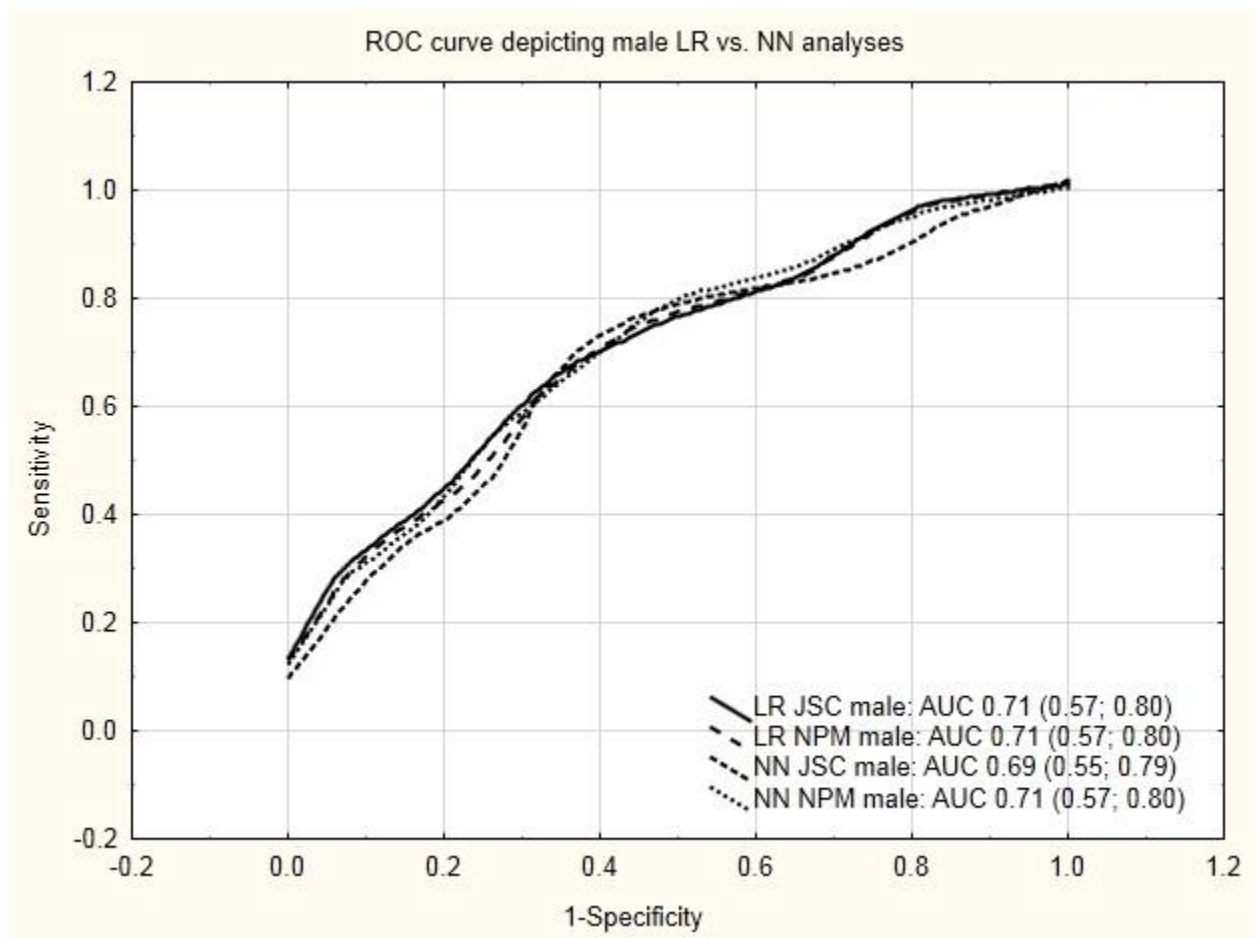
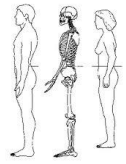


Table 3 compares the LR and NN analyses with regard to sensitivity, -specificity and percent miss-classified. All the analyses, taking note of the means  $\pm$  SEM, were comparable in predicting structural vascular disease.

In Figure 2a (male) and Figure 2b (female) LR analyses revealed comparable AUC values (JSC males:0.71; NPM males:0.71; JSC females:0.82; NPM females:0.84) with the NN analyses (JSC males:0.69; NPM males:0.71; JSC females:0.81; NPM females:0.82).

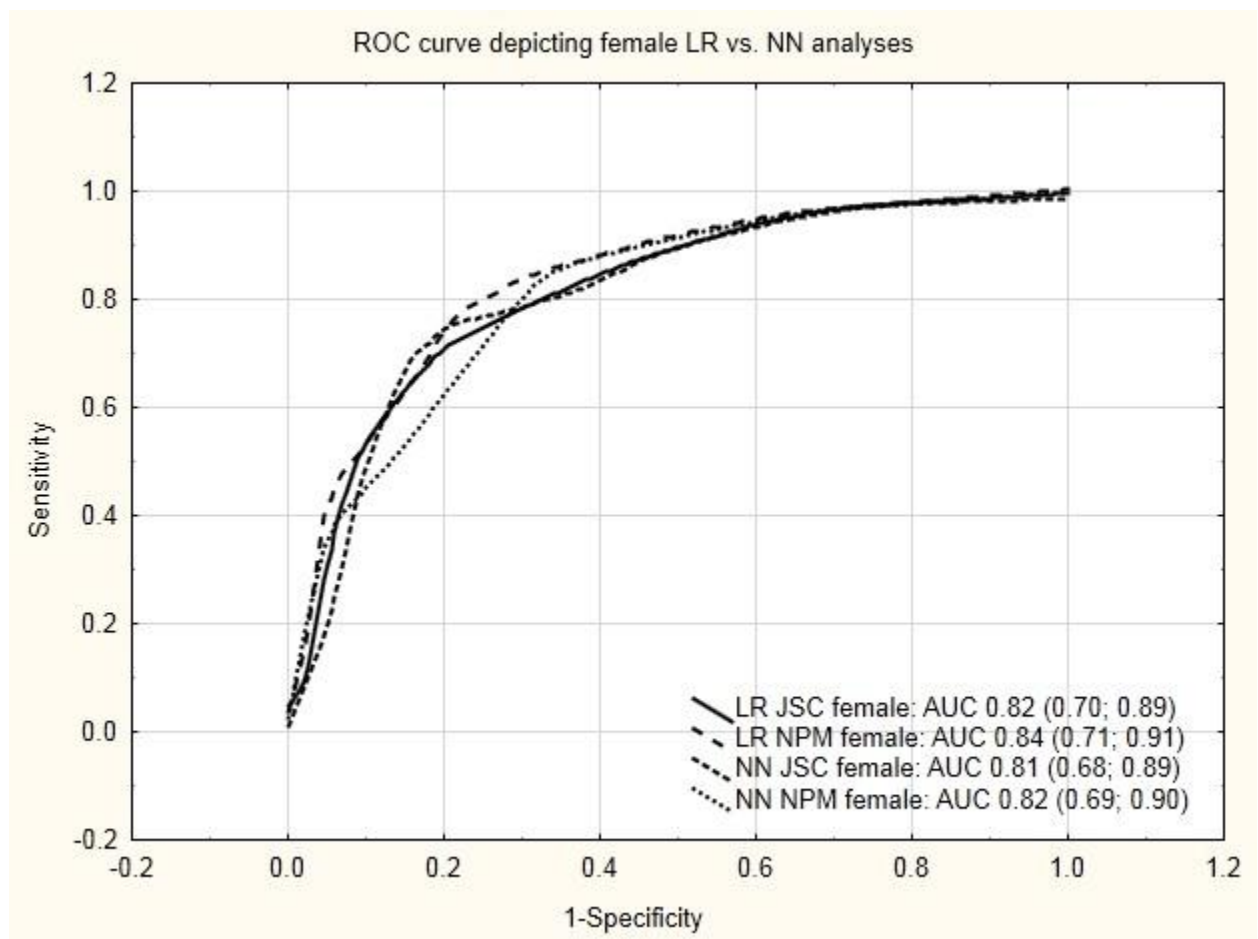


**Figure 2a:** LR and NN analyses ROC curves depicting the association between the 2 WC models (JSC and NPM) and CIMT ( $\geq 1$  mm) for the Male group.

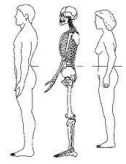


**Legend Figure 2a:** LR and NN analyses ROC curves depicting the 2 WC models (JSC and NPM) vs. CIMT for the Male group, AUC ( $\pm$  95% CI). The LR JSC model yielded a sensitivity and -specificity of 0.61 and 0.74, whilst the LR NPM model yielded sensitivity and -specificity values of 0.66 and 0.68. The NN JSC model, on the other hand yielded a sensitivity and -specificity of 0.44 and 0.90, whilst the NPM model yielded sensitivity and -specificity values of 0.54 and 0.80.

In Figure 2a, within both statistical methods and models (both JSC and NPM) the results were comparable.



**Figure 2b:** LR and NN analyses ROC curves depicting the 2 WC models (JSC and NPM) and CIMT ( $\geq 1$  mm) for the Female group.



**Legend Figure 2b:** LR and NN analyses ROC curves depicting the 2 WC models (JSC and NPM) vs. CIMT for the Female group, AUC ( $\pm$  95% CI). LR JSC model revealed a sensitivity value of 0.71 and -specificity value of 0.82, whilst the LR NPM model showed a sensitivity value of 0.78 and -specificity value of 0.82. The NN JSC model demonstrated sensitivity and -specificity values of 0.78 and 0.82, whilst the NN NPM model showed sensitivity and -specificity values of 0.85 and 0.80.

The LR statistics in Figure 2a reveal that both female models were comparable in predicting CIMT pathology in this group, indicating that the NPM has equal success in differentiating structural vascular disease.

#### 4. Discussion

The main purpose of this study was to apply a receiver operating curve, logistic regression and neural networks in order to compare a new, population-specific, proposed waist circumference cut off model (NPM) with the current IDF cut off points (JSC) [3] by weighing both models against structural vascular disease, as indicated by CIMT.

Our main findings demonstrated that both male and female WC cut off models are comparable in predicting structural vascular disease. ROC analyses, where no adjustments were made for confounders, illustrated that both the JSC and NPM models for both genders were comparable. Cross-validation estimates, where 1000 unique stratified samples were selected and excluded from the data, revealed that the JSC and NPM models were comparable. Consequently, the LR and NN analyses, which were adjusted for confounders (age, BMI and physical activity), demonstrated that both the JSC and NPM models were comparable in predicting structural vascular disease as demonstrated by CIMT.

In another study, Kurt et al. [15] compared LR and NN and demonstrated that multilayer perceptron NN surpassed LR in order to compare the analyses' prediction of coronary artery disease. Our study had smaller sample groups of 81 and 90 participants. In



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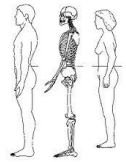
comparison, Kurt et al. [15] had a sample group of 1245, which may account for the different statistical outcomes.

Cameron et al. [21] suggested that central obesity plays a central role in the development of the metabolic syndrome and appears to precede the appearance of the other metabolic syndrome components. Furthermore, the metabolic syndrome is associated with an increased risk of cardiovascular outcomes [22], where carotid intima-media thickness (CIMT) is a strong predictor of future cardiovascular events [23]. In both the male high JSC and NPM models it was clear that CIMT  $\geq 1$  mm as well as hypertensive blood pressure values were increased.

Conversely, as part of the metabolic syndrome criteria, population-specific and country-specific definitions for waist circumference (WC) are recommended [3]. A recent investigation of an African cohort found sphygmomanometer blood pressure (BP) values to be the variable with the highest odds of presenting with pathological WC in both males and females [4]. Therefore a newly proposed optimal WC cut off model (NPM) was suggested where the cut off point for males be set at 90 cm, as opposed to the current 94 cm and the female WC cut off measurement be set at 98 cm as opposed to the existing 80 cm cut off [4].

The female NPM cut off differentiated between the diastolic blood pressure in the descriptive statistics, whereas the JSC failed to do so. This may be expected as the population-specific WC cut off is based on blood pressure. Previous studies found that the African population is a high-risk group regarding the prevalence of hypertension, consequently having higher blood pressure compared to the Caucasian population [24-28], adding to the significance of the ethnicity and population-specific proposed WC cut off [4].

The results of the study must be viewed in light of its limitations. The study sample was not representative of the entire African population. Larger sample sizes may also yield better results when comparing statistical methods. The participants included Setswana-speaking Africans, only one of the eleven ethnic groups residing in South Africa, and we recommend that the findings be verified in other African communities.



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In conclusion, we applied logistic regression and neural networks to compare two proposed waist circumference cut off models in a group of urban African males and females. The results indicate that overall, validation of the NPM model was comparable to the current JSC model, based on logistic regression and neural networks statistical analyses. Both WC models were compared with CIMT, a marker for structural vascular disease.

### **Acknowledgements**

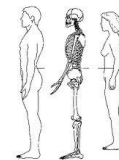
The authors gratefully acknowledge the assistance of Dr G Koekemoer, statistical consultation services of the North-West University for performing the validity analyses. Also, the participation of all members of the SABPA research team, especially C Lessing (RN) and the participants is greatly appreciated.

### **Disclosure**

No conflict of interest

### **Author Contribution**

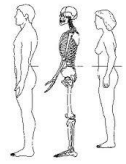
Prof L Malan (principal investigator); Me J Botha, Prof JH de Ridder, Prof JC Potgieter and Prof HS Steyn (staff members) (Potchefstroom). JB and LM planned the manuscript; JB analysed the data and HSS assisted and reviewed the analyses of the data; JB wrote the manuscript; JHdR, LM, JCP, HSS edited the manuscript.



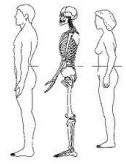
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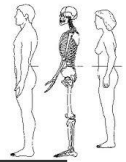
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Chapter  
5**Association of waist circumference with perception of own health in a group of urban African males and females: the Sympathetic Activity and Ambulatory Blood Pressure in Africans (SABPA) study.**

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Johannes. H. de Ridder

**Abstract:**

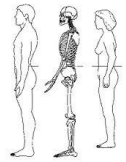
**Objectives:** Current waist circumference (WC) cut off points of the Joint Statement Consensus (JSC) (male  $\geq 94$  cm, female  $\geq 80$  cm) were compared with a newly proposed WC cut point model (NPM) (male  $\geq 90$  cm, female  $\geq 94$  cm). In this study we aimed to compare the two models in terms of its predictive value with regards to the perception of own health.

**Method:** We determined BP and fasting bloods (glucose, high density lipoprotein (HDL) and triglycerides) as metabolic syndrome markers for 171 urban teachers. Perception of own health was determined via the General Health Questionnaire-28 (GHQ-28).

**Results:** NPM was an improved discriminator between the WC groups regarding perceived mental health as reflected in the GHQ-subscores. In the male group, higher scores were obtained by the NPM high WC ( $\geq 90$  cm) compared to the low WC groups ( $< 90$  cm) for somatic symptoms, social dysfunction and GHQ total score. Compared to the high WC NPM ( $\geq 98$  cm) females, the low WC ( $< 98$  cm) obtained significantly higher anxiety and sleeplessness subscale scores.

**Conclusion:** Results suggest that in this African cohort, when adding perception of health as a discriminatory variable between models, the NPM distinguished better between WC groups based on their perception of own health than the JSC model.

**Keywords:** [waist circumference; anthropometry; perception of health; ethnicity]



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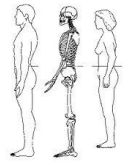
## 1. Introduction

A cluster of risk factors for cardiovascular disease and type 2 diabetes mellitus has become known as the metabolic syndrome. The recent Joint Statement Consensus (JSC) model for metabolic syndrome factors includes raised blood pressure, dyslipidaemia (raised triglycerides and lowered high-density lipoprotein cholesterol), raised fasting glucose, and central obesity.<sup>1</sup> Currently, the International Diabetes Federation (IDF) cut points are to be used for non-Europeans until more data are available. The IDF suggested a cut off point of  $\geq 94$  cm for males and  $\geq 80$  cm for females. Alberti et al.<sup>1</sup>, however, proposed that population- and country-specific recommended waist circumference thresholds for abdominal obesity be used. A recent investigation of an African cohort found sphygmomanometer blood pressure (BP) values to be the variable with the highest odds of presenting with pathological WC in both males and females.<sup>2</sup> Therefore a newly proposed optimal WC cut off model (NPM) was suggested where the cut off point for males be set at 90 cm, as opposed to the current 94 cm, and the female WC cut off measurement be set at 98 cm as opposed to the existing 80 cm cut off.<sup>2</sup>

Obesity is a well-known cause of cardiovascular disease burden and premature death, but associations with psychological morbidity remain uncertain.<sup>3</sup> Several studies showed that poor mental health was associated with obesity.<sup>4-7</sup> Au contraire, other studies found no association between mental health and obesity.<sup>3,8</sup>

Van der Merwe<sup>9</sup> stated that psychological co-morbidity in patients with obesity is associated with a variety of medical and dietary problems as well as demographic, social and cognitive risk factors. Björntorp and Rosmond<sup>10</sup> proposed that psychosocial and socioeconomic handicaps could be expected to be followed by frequent stress reactions and an increase in alcohol- and smoking habits, and often show robust associations with depressive and anxiety traits. In the current South African context, the rapid rate of urbanization with its accompanying insecurities and disruption in social relationships, could contribute to psychological distress and a perception of poorer own health.<sup>11</sup>

Urbanization is a critical factor influencing the traditional ideal body image among Africans, who have always been inclined towards a larger, fuller body shape.<sup>12</sup> In this



“Big is beautiful” group, obesity was traditionally associated with dignity, health, wealth and respect.<sup>13</sup> However, in view of massive socio-economic changes and increased social integration taking place, black young South Africans are rapidly being exposed to different belief systems. This may alter their value systems regarding body size<sup>14</sup>, which may in turn influence their perception of own health.

Consequently we aimed to compare two models of central obesity, JSC<sup>1</sup> and the NPM<sup>2</sup>, and its associations with perception of own health in a cohort of urban Africans.

## 2. Methods

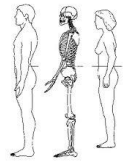
This sub-study formed part of the Sympathetic activity and Ambulatory Blood Pressure in Africans (SABPA) prospective cohort study, conducted from February to May 2008. Permission to participate was granted by the North-West Department of Education and the South African Teachers Union, as well as the Ethics Committee of the North-West University (project nr: NWU-00036-07S6). The study conformed to the ethical guidelines of the World Medical Association Declaration of Helsinki (revised 2004). All participants signed an informed consent form.

### 2.1 Study population and sample

Male and female urban black African (hereafter referred to as African) school teachers, (N=200, aged 25-65 years) were recruited and included as participants from governmental organizations in the Dr. Kenneth Kaunda District in North-West Province. Exclusion criteria for this study included ear temperature  $>37,5^{\circ}\text{C}$ , alpha- or beta blockers usage, donated blood or vaccinated in the previous 3 months. Clinically confirmed diabetics (n=7), users of anti-depressant medication (n=1) and HIV-infected (n=13) participants were also excluded. The final participant sample comprised 171 Africans (n=81 males and females n=90).

### 2.2 Experimental procedure

On every working day of the week, for the extent of the project, the physical activity monitors (Actical®) apparatuses were fitted at 07:00, to four participants for 24 hours. Software programmes were activated and participants resumed their daily activities.



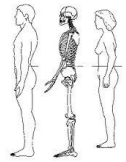
At 16:30 they were transported to the Metabolic Unit Research Facility of the North-West University for an overnight stay where participants were welcomed and introduced to the experimental setup. The first half of a psychosocial battery was completed under supervision of trained field workers and at 18:00 participants received a standardized dinner. After dinner the last part of the psychosocial battery questionnaires were completed and last beverages were given at 20:30.

After an overnight fast, the Actical® apparatuses were disconnected. Anthropometric measurements were taken in triplicate by anthropometrists according to the standard of the International Society for the Advancement of Kinanthropometry (ISAK).<sup>15</sup> Two mercury sphygmomanometer blood pressure readings using Korotkoff IV or V for blood pressure (BP) followed while the participants rested for five minutes in a semi-recumbent position, with a three-minute rest between measurements. The second measurement was used for statistical analyses. BP measurements were taken by a registered nurse. Fasting blood samples were obtained by a registered nurse.

### 2.3 Anthropometric measurements

All anthropometric measurements were performed by ISAK level 2 accredited anthropometrists, with subjects in minimal clothing and without shoes. The waist circumference was taken in triplicate at the end of normal expiration at the narrowest point of the abdomen between the lower costal (10<sup>th</sup> rib) border and the top of the iliac crest, perpendicular to the long axis of the trunk.<sup>15</sup> The subject assumed a relaxed standing position with the arms folded across the thorax.<sup>15</sup>

Body mass index (BMI) was calculated by the  $\text{kg/m}^2$  formula. The body weight measured by means of a KRUPS scale with the participant wearing minimal clothes and with the weight evenly distributed, to the nearest 0.1 kg.<sup>15</sup> Height was measured with a stadiometer to the nearest 0.1 cm while the participant's head was in the Frankfort plane, the heels together and the buttocks and upper back touching the stadiometer.<sup>15</sup>



## 2.4 General Health Questionnaire-28 (GHQ-28)<sup>16</sup>

The General Health Questionnaire (GHQ-28) is a 28-item self-report measure aimed at distinguishing people with some form of psychological disturbance from those who are relatively healthy.<sup>16</sup> The GHQ has four subscales (Somatic Symptoms, Anxiety and Insomnia, Social Dysfunction, and Severe Depression), each of which consists of seven items. Subjects reported on their own perception of health during the past few weeks. Each item was scored via bimodal scoring on a 0-0-1-1 scale.<sup>16</sup> The value of response possibilities one (1) and two (2) are equal to nil (0) and three (3) and four (4) equal to one (1). The sum of these scale scores yields a single score that ranges from 0 (for no symptoms) to 28 (severe pathology), for which threshold scores of  $\geq 4$  (bimodal scoring) would indicate probable psychological distress and/or psychiatric disorder, hereafter described as caseness.<sup>17</sup>

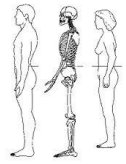
The median specificity of the GHQ-28 is 0.82 and the median sensitivity is 0.86.<sup>18</sup> Cronbach alpha-reliability coefficients ranged between 0.78 and 0.85 for different subscales in this study.

## 2.5 Biochemical analysis

Fasting blood samples were obtained by a registered nurse with a winged infusion set from the brachial vein branches from the dominant arm. Serum gamma glutamyl transferase and cotinine were used as markers of alcohol abuse and smoking respectively. Sodium fluoride glucose, serum HDL and triglycerides were handled according to standardized procedures and were stored at  $-80^{\circ}\text{C}$ . Analyses were done using the Konelab<sup>TM</sup> 20i sequential multiple analyzer computer (SMAC) (Thermo Scientific, Vantaa, Finland) and Unicel DXC 800 (Beckman & Coulter, Germany) at accredited laboratories.

## 2.6 Statistical Analyses

Data analyses were performed with Statistica version 10.<sup>19</sup> Descriptive statistics and prevalence were obtained for the entire black African sample. A 2 X 2 analysis of covariance (ANCOVA) was executed to test the interaction between WC (JSC<sup>1</sup> as well as NPM<sup>2</sup>) and gender with GHQ-caseness and metabolic syndrome markers as outcome variables. Subsequent ANCOVAs followed independent of priori confounders age, BMI

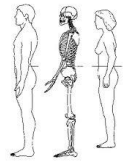


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and physical activity. Standardized differences in means as effect sizes ( $d$ ) were also determined as a measure of practical significance, where practical significance can be understood to be a large enough difference to have an effect in practice.<sup>20</sup> As such, an effect size of 0.2 is described as small and not visible. Likewise, an effect size of 0.5 is described as medium and visible to a researcher. An effect size of 0.8 is practically significant and can be understood as a large enough difference to have an effect in practice. Odds Ratio analyses were performed to assess the odds of each component predicting psychological distress in high WC gender groups. Independent covariates included age, BMI, physical activity.

### 3. Results

In Table I the Africans demonstrated levels of obesity, high normal glucose and blood pressure values with 74.27% reporting a perception of own poorer health. Interaction on main effects (NPM  $\times$  gender) demonstrated significance for somatic symptoms ( $F(1,171), 5.28$ ;  $P = 0.02$ ) and GHQ-caseness ( $F(1,171), 4.62$ ;  $P = 0.02$ ). Subsequently, groups were stratified into high and low WC groups for (1) JSC and (2) NPM. Separate analysis of covariance (ANCOVA) independent of confounders (age, BMI and physical activity) were done, firstly for the males (Table II), and thereafter for the females (Table III) to determine significant differences.

**Table I: Descriptive statistics for the entire African group.**

| <b>Total African group<br/>N = 171</b>                       |                  |
|--|------------------|
| Males (N)  | 81               |
| Females (N)  | 90               |
| Age (years)  | 44.05 ± 8.30     |
| <b>Lifestyle variables</b>                                   |                  |
| Physical Activity (kCal)                                     | 2693.04 ± 809.67 |
| Gamma-glutamyl transferase (μ/L)                             | 62.59 ± 72.27    |
| Cotinine (ng/ml)   | 20.94 ± 51.66    |
| <b>Physiological and anthropometric variables</b>            |                  |
| Body Mass Index (kg/m <sup>2</sup> )                         | 30.36 ± 7.14     |
| Glucose (mmol/L)   | 5.50 ± 1.73      |
| High Density Lipoprotein (mmol/L)                            | 1.15 ± 0.35      |
| Triglycerides (mmol/L)                                       | 1.38 ± 1.30      |
| Systolic Blood Pressure (mmHg)                               | 134 ± 20         |
| Diastolic Blood Pressure (mmHg)                              | 89 ± 13          |
| <b>Psychological variables: General Health Questionnaire</b> |                  |
| Somatic Symptoms   | 2.54 ± 2.20      |
| Anxiety & Sleeplessness                                      | 2.64 ± 2.46      |
| Social Dysfunction   | 2.01 ± 2.12      |
| Depressive Symptoms  | 1.16 ± 1.93      |
| GHQ_Total  | 8.37 ± 6.61      |
| GHQ caseness N (%)   | 127 (74.27%)     |
| <b>Medication usage</b>                                      |                  |
| Hypertension N (%)   | 34 (19.88%)      |

Variables indicated as arithmetic mean ± standard deviation (SD).

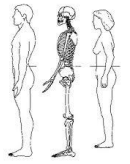
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**Table II: Comparing males' JSC<sup>1</sup> WC cut off and the NPM WC<sup>2</sup>, independent of covariates (age, BMI and physical activity).**

|   | 2009 JSC<br>Males (N=43)<br><94cm | 2009 JSC<br>Males (N=38)<br>≥94cm | Effect<br>size<br>(d) | NPM<br>Males (N=36)<br><90cm | NPM<br>Males (N=45)<br>≥90cm | Effect<br>size<br>(d) |
|---|-----------------------------------|-----------------------------------|-----------------------|------------------------------|------------------------------|-----------------------|
| Age (years)                                       | 41.37 ± 9.33                      | 44.08 ± 6.90                      |                       | 40.50 ± 9.79                 | 44.36 ± 6.60*                | 0.39                  |
| <b>Lifestyle variables</b>                        |                                   |                                   |                       |                              |                              |                       |
| Physical Activity (kCal)                          | 2350.24 ± 654.08                  | 3145.41 ± 797.02*                 | 1.00                  | 2341.32 ± 705.58             | 3028.85 ± 788.40*            | 0.87                  |
| Gamma-glutamyl transferase (µ/L)                  | 66.72 ± 86.10                     | 87.10 ± 52.21                     |                       | 71.08 ± 93.41                | 80.45 ± 50.78                |                       |
| Cotinine (ng/ml)                                  | 25.42 ± 48.84                     | 22.53 ± 46.76                     |                       | 25.81 ± 46.94                | 22.67 ± 48.61                |                       |
| <b>Physiological and anthropometric variables</b> |                                   |                                   |                       |                              |                              |                       |
| Body Mass Index (kg/m <sup>2</sup> )              | 23.52 ± 3.15                      | 32.37 ± 4.73*                     | 1.87                  | 23.06 ± 3.10                 | 31.36 ± 5.03*                | 1.65                  |
| Glucose (mmol/L)                                  | 5.19 (4.41; 5.96)                 | 6.94 (6.10; 7.79)*                | 0.67                  | 5.21 (4.37; 6.05)            | 6.66 (5.93; 7.38)*           | 0.57                  |
| High Density Lipoprotein (mmol/L)                 | 1.16 (1.02; 1.29)                 | 1.0 (0.85; 1.15)                  |                       | 1.15 (1.01; 1.30)            | 1.02 (0.90; 1.15)            |                       |
| Triglycerides (mmol/L)                            | 1.48 (0.82; 2.13)                 | 2.18 (1.46; 2.89)                 |                       | 1.63 (0.92; 2.34)            | 1.95 (1.33; 2.56)            |                       |
| Systolic Blood Pressure (mmHg)                    | 132 (125; 139)                    | 149 (141; 157)*                   | 0.67                  | 131 (123; 139)               | 146 (140; 153)*              | 0.64                  |
| Diastolic Blood Pressure (mmHg)                   | 89 (84; 95)                       | 99 (93; 104)*                     | 0.52                  | 91 (85; 97)                  | 96 (91; 101)                 |                       |
| Hypertension medication N (%)                     | 4 (9.30 %)                        | 9 (23.68 %)                       |                       | 4 (11.11 %)                  | 9 (20 %)                     |                       |
| <b>Psychological variables</b>                    |                                   |                                   |                       |                              |                              |                       |
| <b>General Health Questionnaire</b>               |                                   |                                   |                       |                              |                              |                       |
| Somatic Symptoms                                  | 1.88 (1.12; 2.64)                 | 3.01 (2.18; 3.84)                 |                       | 1.58 (0.77; 2.38)            | 3.08 (2.39; 3.77)*           | 0.62                  |
| Anxiety & Sleeplessness                           | 2.59 (1.65; 3.53)                 | 2.44 (1.42; 3.46)                 |                       | 2.26 (1.25; 3.26)            | 2.73 (1.86; 3.60)            |                       |
| Social Dysfunction                                | 1.60 (0.78; 2.41)                 | 2.07 (1.18; 2.95)                 |                       | 1.01 (0.16; 1.86)            | 2.46 (1.73; 3.20)*           | 0.57                  |
| Depressive Symptoms                               | 0.54 (-0.07; 1.15)                | 1.36 (0.69; 2.02)                 |                       | 0.56 (-0.10; 1.22)           | 1.21 (0.65; 1.78)            |                       |
| GHQ_Total   | 6.62 (4.12; 9.12)                 | 8.89 (6.17; 11.61)                |                       | 5.42 (2.79; 8.05)            | 9.49 (7.22; 11.77)*          | 0.51                  |
| GHQ caseness N (%)                                | 26 (60.47 %)                      | 28 (73.68 %)                      |                       | 21 (58.33 %)                 | 33 (73.33 %)                 |                       |

Variables indicated as arithmetic mean ± standard deviation (SD); subsequent data presented as geometric mean ± 95% confidence interval (95%CI). Statistical difference indicated with \*, p ≤ 0.05



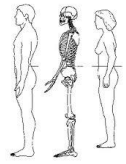
In Table II, for both models (JSC and NPM), the lifestyle and physiological variables were less favourable for the high WC cut off male groups. The outcomes of the GHQ and its subscales demonstrated increases in somatic symptoms, social dysfunction and total poorer perception of own health in the high WC groups of the NPM only.



**Table III: Comparing female’s JSC<sup>1</sup> WC cut off and the NPM WC cut off<sup>2</sup>, independent of covariates (age, BMI and physical activity).**

|   | 2009 JSC<br>Females (N=17)<br><80cm | 2009 JSC<br>Females (N=73)<br>≥80cm | Effect<br>size<br>(d) | NPM<br>Females (N=58)<br><98cm | NPM<br>Females (N=32)<br>≥98cm | Effect<br>size<br>(d) |
|---|-------------------------------------|-------------------------------------|-----------------------|--------------------------------|--------------------------------|-----------------------|
| Age (years)                                       | 42.77 ± 6.54                        | 45.92 ± 8.35                        |                       | 44.31 ± 7.28                   | 47.16 ± 9.25                   |                       |
| <b>Lifestyle variables</b>                        |                                     |                                     |                       |                                |                                |                       |
| Physical Activity (kCal)                          | 1957.67 ± 383.15                    | 2830.74 ± 783.68*                   | 1.11                  | 2325.34 ± 487.42               | 3282.97 ± 891.26*              | 1.07                  |
| Gamma-glutamyl transferase (µ/L)                  | 39.99 ± 44.65                       | 49.71 ± 73.99                       |                       | 39.00 ± 35.02                  | 63.91 ± 105.22                 |                       |
| Cotinine (ng/ml)                                  | 20.31 ± 48.72                       | 17.56 ± 56.90                       |                       | 18.26 ± 61.36                  | 17.68 ± 42.72                  |                       |
| <b>Physiological and anthropometric variables</b> |                                     |                                     |                       |                                |                                |                       |
| Body Mass Index (kg/m <sup>2</sup> )              | 25.70 ± 3.50                        | 34.43 ± 6.95*                       | 1.26                  | 29.26 ± 4.53                   | 39.16 ± 7.04*                  | 1.41                  |
| Glucose (mmol/L)                                  | 4.25 (3.65; 4.85)                   | 5.20 (4.95; 5.45)*                  | 0.79                  | 4.75 (4.42; 5.07)              | 5.54 (5.05; 6.02)*             | 0.58                  |
| High Density Lipoprotein (mmol/L)                 | 1.32 (1.14; 1.50)                   | 1.20 (1.12; 1.26)                   |                       | 1.23 (1.14; 1.33)              | 1.18 (1.04; 1.32)              |                       |
| Triglycerides (mmol/L)                            | 0.78 (0.48; 1.08)                   | 1.04 (0.91; 1.16)                   |                       | 0.90 (0.75; 1.06)              | 1.15 (0.92; 1.38)              |                       |
| Systolic Blood Pressure (mmHg)                    | 128 (119; 136)                      | 130 (126; 134)                      |                       | 125 (120; 129)                 | 138 (131; 145)*                | 0.69                  |
| Diastolic Blood Pressure (mmHg)                   | 83 (77; 88)                         | 85 (82; 87)                         |                       | 82 (79; 84)                    | 89 (85; 94)*                   | 0.62                  |
| Hypertension medication N (%)                     | 1 (5.88 %)                          | 20 (27.40 %)                        |                       | 7 (22.58 %)                    | 14 (24.56 %)                   |                       |
| <b>Psychological variables</b>                    |                                     |                                     |                       |                                |                                |                       |
| <b>General Health Questionnaire</b>               |                                     |                                     |                       |                                |                                |                       |
| Somatic Symptoms                                  | 2.92 (1.62; 4.22)                   | 2.61 (2.03; 3.17)                   |                       | 2.97 (2.26; 3.68)              | 2.10 (1.04; 3.16)              |                       |
| Anxiety & Sleeplessness                           | 2.23 (0.84; 3.62)                   | 2.88 (2.27; 3.48)                   |                       | 3.32 (2.57; 4.07)              | 1.74 (0.63; 2.85)*             | -0.44                 |
| Social Dysfunction                                | 1.55 (0.36; 2.74)                   | 2.34 (1.82; 2.86)                   |                       | 2.10 (1.44; 2.76)              | 2.36 (1.38; 3.404)             |                       |
| Depressive Symptoms                               | 1.57 (0.37; 2.77)                   | 1.32 (0.80; 1.84)                   |                       | 1.58 (0.92; 2.23)              | 0.99 (0.01; 1.96)              |                       |
| GHQ_Total   | 8.37 (4.68; 11.07)                  | 9.13 (7.51; 10.74)                  |                       | 9.98 (7.96; 11.99)             | 7.19 (4.20; 10.19)             |                       |
| GHQ caseness N (%)                                | 13 (76.47 %)                        | 60 (82.19 %)                        |                       | 28 (87.5 %)                    | 45 (77.59 %)                   |                       |

Variables indicated as arithmetic mean ± standard deviation (SD); subsequent data presented as geometric mean ± 95% confidence interval (95%CI). Statistical difference indicated with \*, p ≤ 0.05



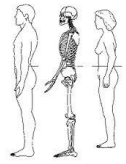
In Table III, a marked difference between the two models (JSC and NPM) is evident among female participants. The NPM demonstrated that the low WC group ( $< 98$  cm) showed more anxiety and sleeplessness even though the high NPM group ( $\geq 98$  cm) revealed a higher mean blood pressure. In line with this it is noteworthy that the low WC NPM group showed a trend toward more Somatic Symptoms, Depressive Symptoms and a higher GHQ Total Score than the high WC group.

Odds Ratios were performed and demonstrated that alcohol abuse best predicted psychological distress in both male high WC JSC and RPWC groups although clinically not indicating risk with an OR of  $> 1$ . In the female group the Odds Ratio revealed an odds of alcohol associated with psychological distress, although not clinically indicating risk in the high WC female JSC group. Interestingly, the female high WC RPWC cut off showed no odds of any of the variables predicting psychological distress.

#### 4. Discussion

The main purpose of this study was to compare two proposed models of central obesity, the JSC<sup>1</sup> and an ethnic-specific newly proposed model (NPM).<sup>2</sup> The association between each model and perception of own mental health was assessed in a group of urban African males and females.

In the male group, it was found that both models were good discriminators between high WC and low WC groups for lifestyle and physiological variables. The NPM<sup>2</sup> though, revealed statistically significant differences between high and low WC groups pertaining to psychological distress. The NPM male high WC group demonstrated a less favourable metabolic profile, and also reported significantly poorer perception of own mental health as represented by somatic symptoms and social dysfunction. This is in line with findings by Bodenlos et al.<sup>7</sup> who reported that the relationship between mood disorders and obesity had a trend towards significance in African Americans. The prominence of somatic symptoms in this group also seemed to confirm the findings of Kirmayer and Young<sup>21</sup>, who indicated that ethno-cultural groups in the same urban milieu, with equal access to healthcare services, almost exclusively demonstrate somatic symptoms rather than psychosocial distress.

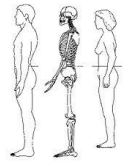


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The female cut offs differed extensively between the two models ( $\geq 80$  cm vs.  $\geq 98$  cm). In an attempt to suggest population-specific cut offs for this population, Prinsloo et al.<sup>2</sup> did receiver operating characteristic (ROC) analyses for each of the Metabolic Syndrome components. The blood pressure cut off, used in our current sub-study, was the most significant predictor of pathology. Blood pressure is mostly a major risk factor in people of African descent, regardless of country of residence.<sup>22-26</sup>

Focusing on the female's perception of own health, the NPM distinguished between low and high groups with regard to the Anxiety and Sleeplessness sub-scale, indicating that the low group ( $< 98$  cm) had a higher occurrence of anxiety and sleeplessness even though the high group ( $\geq 98$  cm) demonstrated higher mean blood pressure. Katzman and Lee<sup>27</sup> theorized that women juggling two cultural worlds exposed to “Western Ideals” in their home countries show an increase in disordered eating. The results of this study might also offer the hypothesis of metabolically healthy obese females as first put forward by Walker et al.<sup>28</sup> Kruger et al.<sup>29</sup> found that WC was associated with the risk for non-communicable disease in black South African women. In line with this, the current study shows that the high WC group is not metabolically healthy, but healthy by their own perception, whereas the low WC group is metabolically healthy with a very high occurrence of psychological distress. Healthy obesity in this group does not seem to refer to metabolic health but rather to improved perception of own health. This trend seems to be particularly prominent when the NPM cut offs are used.

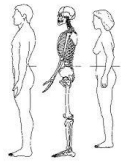
Dallman et al.<sup>30</sup> proposed that people eat comfort food in an attempt to reduce the activity of the chronic stress-response network. In line with this, Kivimäki et al.<sup>4</sup> suggested that internalization of negative obesity-related stereotypes, negative self-body image and unsuccessful weight control by dieting were related to decreased mental health among individuals who are obese. Oswald and Powdthavee<sup>31</sup> demonstrated that a 10-point increase in body mass index (BMI) was associated with a drop in mental health of approximately 0.3 points on the General Health Questionnaire<sup>17</sup>. Whether these findings hold true for the African population is not certain, especially in line with our findings.



No differences in GGT levels in the low vs. the high JSC and NPM cut off groups may rule out the fact that obesity is responsible for the increased GGT levels. Higher levels of especially GGT in both the high WC cut off JSC and NPM male groups were evident although odds ratio's revealed that alcohol abuse showed odds with low clinical significance for psychological distress in the high WC JSC males (OR 0.97,  $\pm 95\%CI$  0.93; 1.00;  $P = 0.05$ ) and in high WC male NPM (0.98, 0.95;1.00;  $P = 0.08$ ). Malan et al.<sup>32</sup> stated that urban gender groups reported higher alcohol consumption, maybe as a manner of coping. Conversely, Hamer et al.<sup>33</sup> confirmed that objectively measured alcohol intake, as indicated by GGT, exceeded normal cut off values and were associated with structural vascular disease (OR 3.9; 95% CI 1.0-15.5). Saxena<sup>34</sup> and the World Health Organization (WHO)<sup>35</sup> reported Africa to be one of the two regions with the most rapid rise in alcohol consumption, where the consumption of absolute alcohol per person per year is as much as six litres. Also, levels of alcohol consumption is an indication of the burden of disease for the country, leading to serious health and social consequences in our country<sup>35</sup>. Of note is the absence of the high WC female NPM group's odds of presenting with increased alcohol use, possibly indicating their own perception of own good health and not needed as a coping strategy. Further research is needed on this topic.

The results of the study must be viewed in light of its limitations. Given the cross-sectional nature of this study, causal relationships between central obesity and perception of health could currently not be determined. Another limitation was that the study sample was not representative of the entire African population. The participants included Setswana-speaking Africans, only one of the eleven ethnic groups residing in South Africa and we recommend that the findings be verified in other African communities.

In conclusion we compared two models of central obesity and its associations with perception of health in a group of urban African males and females: The model proposed by Prinsloo et al.<sup>2</sup> appeared to be a clearer discriminate for the perception of own health in both African males and females, with WC cut off points set at 90 cm and 98 cm, respectively.



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**Acknowledgements**

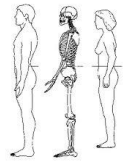
The authors gratefully acknowledge the assistance of all members of the SABPA research team, especially C Lessing (RN) and the participants.

**Disclosure**

No conflict of interest

**Author Contribution**

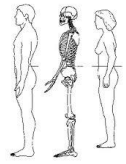
Prof L Malan (principal investigator); Me J Prinsloo, Prof JC Potgieter, Prof HS Steyn, Prof JH de Ridder (staff members) (Potchefstroom). JP planned, analysed the data, wrote and edited the manuscript; LM planned, analysed the data and edited the manuscript; JCP edited the manuscript; HSS analysed the data and edited the manuscript, JHdR edited the manuscript.



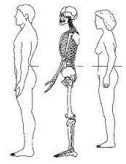
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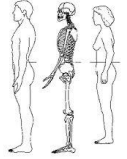
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## Conclusions and Recommendations

- 6.1 Summary
- 6.2 Conclusions
- 6.3 Recommendations

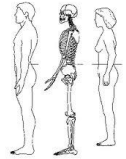
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### 6.1 Summary

The aim of this study was firstly to determine which waist circumference best predicted metabolic syndrome in a group of urban African teachers. The second aim was to compare two WC models, namely the Joint Statement Consensus (JSC) and a new ethnicity-specific proposed model (NPM), and its contribution towards structural vascular disease as indicated by carotid intima-media thickness (CIMT). Validation of the NPM via logistic regression and neural networks illustrated that the NPM was comparable to the JSC cut offs currently in use. Thirdly, the aim of the study was to assess whether any of the two models (JSC or NPM) and markers of metabolic syndrome correlated with perception of own health.

This dissertation was presented in four main parts, namely an introduction (Chapter 1), a literature review (Chapter 2), and three research articles (Chapters 3, 4 and 5). The article format of the dissertation is approved by the North-West University (Potchefstroom Campus), and the three research articles have been submitted to appropriate and accredited journals.

Chapter 1 introduced the problem, and stated the aims and hypotheses of this study. The literature (Chapter 2) review focused on the relationship between central obesity and psychological distress.



Chapter 3 took the form of an article: **Determining the waist circumference cut off which best predicts the metabolic syndrome components in urban Africans: the SABPA study.** This article was accepted for publication by *Experimental and Clinical Endocrinology and Diabetes* (Prinsloo, J., Malan, L., De Ridder, J.H., Potgieter, J.C. & Steyn, H.S. 2011. Determining the waist circumference cut off which best predicts the metabolic syndrome components in urban Africans: the SABPA study. *Experimental and Clinical Endocrinology and Diabetes*, 119:599-603).

The second article, Chapter 4, consists of the second research article titled: **Comparing performances of two central obesity models to predict structural vascular disease by using ROC analyses, Logistic Regression and Neural networks: the SABPA study.** This article has been submitted to *Atherosclerosis*, a journal with an impact factor of 4.1.

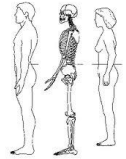
The final article, Chapter 5, was titled **Association of waist circumference with perception of own health in a group of urban African males and females: the Sympathetic Activity and Ambulatory Blood Pressure in Africans (SABPA) study.** This article has been submitted to the *Journal of Endocrinology, Metabolism and Diabetes of South Africa* and is currently in rebuttal.

## 6.2 Conclusions

The conclusions drawn from this research project are presented in accordance with the set hypotheses (Chapter 1).

- **Hypothesis 1: New population-specific WC cut off points in a cohort of urban Africans would differ from those proposed by the JSC.**

The first hypothesis is accepted, based on the research findings that, via ROC analyses, suggested new population- and ethnicity-specific cut off points that differed from the currently proposed JSC model, proposed by the International Diabetes Federation. This study suggest that the optimal cut off point for the males be set at 90 cm, as opposed to the current 94 cm; whilst the female cut off be set at 98 cm as opposed to the existing



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cut off of 80 cm based on blood pressure, confirming the necessity of early diagnosis, especially in view of the high risk of hypertension in people of African descent.

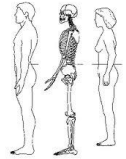
- **Hypothesis 2: The NPM would predict structural vascular disease (CIMT) in a cohort of urban Africans.**

The second hypothesis is accepted based on the research findings that overall, validation of the NPM model was comparable with the current JSC model in predicting structural vascular disease in a group of urban African teachers.

- **Hypothesis 3: The NPM would predict a poorer perception of own health in a cohort of urban Africans.**

The third hypothesis is accepted, based on the research findings that in this African cohort, when adding perception of health as a discriminatory variable between models, the NPM distinguished better between WC groups, based on their perception of own health, than the JSC model.

The Joint Statement Consensus, the most recent definition of the metabolic syndrome, declared set cut offs for all components except waist circumference, for which further ethnicity- and country-specific work is required (Alberti *et al.*, 2009:1640). A new population and ethnicity-specific WC cut off was recommended (NPM) for a group of urban Africans and subsequently validated via Logistic Regression and Neural Networks statistical analyses. The NPM was comparable with the JSC cut offs which are currently in use in predicting structural vascular disease with CIMT as marker. It is therefore proposed that the NPM cut offs be used in this population due to the strong association between blood pressure and the proposed WC cut offs, as blood pressure is a major risk factor in people of African descent, regardless of country of residence (Cappuccio *et al.*, 1997; Wild and Mckeigue, 1997; Agyemang and Bhopal, 2003; Van der Merwe and Pepper, 2006; Vorster, 2002; Du Plessis *et al.*, 2010; Sliwa *et al.*, 2008). Furthermore, associations were demonstrated between the NPM and perception of own health in a group of urban Africans, adding weight to the NPM as a new ethnicity-specific cut off in this group.



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### 6.3 Recommendations

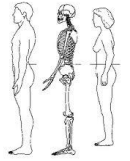
The results from this study contributed to the lack of data available on South African ethnicity- and population-specific WC cut offs that predict pathology. The findings of this study give support to the possibility of a population-specific WC cut off that differs substantially from the currently proposed WC cut offs in use. It is proposed that the NPM cut offs be used in this population due to the strong association between blood pressure and the proposed WC cut offs, validated by Logistic Regression and Neural Networks statistical analyses. As the NPM was associated with the GHQ, it is recommended that the ethnicity-specific cut off be used and that cultural differences could mask subtle changes.

Certain shortcomings regarding this study can, however, be indicated as there are some important factors that may have affected the results and could have caused weaknesses in this study and, therefore, might have influenced the different outcomes.

First, analyses used cross-sectional data. Therefore, our results do not establish a causal relationship between perception of own health and waist circumference cut offs (Hypothesis 3). A longitudinal follow-up of the subjects and changes in perception of own health status would provide stronger evidence for a causal association between perception of own health and body composition.

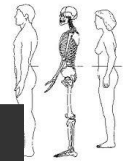
Secondly, the study consisted of a small number of subjects. As mentioned above, a longitudinal study design with larger numbers of participants would strengthen recommended WC cut off points for age, gender- and ethnicity-specific groups.

Lastly, the participants included Setswana-speaking Africans, only one of the eleven ethnic groups residing in South Africa, and we recommend that the findings be verified in other African communities.



In conclusion:

A new population and ethnicity-specific WC cut off was recommended (NPM) and subsequently validated with Logistic Regression and Neural Networks statistical analyses. The NPM was comparable with the JSC cut offs which are currently in use in predicting structural vascular disease via CIMT. It is therefore proposed that the NPM cut offs be used in this population due to the strong association between blood pressure and the proposed WC cut offs, validated by Logistic Regression and Neural Networks statistical analyses. Furthermore, associations were demonstrated between the NPM and perception of own health in a group of urban Africans.



# Appendices

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ℓ Logistic Regression: JSC males

Test of all effects

| CIMT atherosclerosis or not - Test of all effects (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |                  |            |          |
|--|------------------|------------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |                  |            |          |
| Modeled probability that CIMT atherosclerosis or not = 1   |                  |            |          |
| Include condition: v2=1  |                  |            |          |
| Effect   | Degr. of Freedom | Wald Stat. | p        |
| Intercept  | 1                | 4.463965   | 0.034617 |
| AGE  | 1                | 7.122030   | 0.007614 |
| BMI  | 1                | 0.536494   | 0.463889 |
| METS (kcal)  | 1                | 0.180470   | 0.670970 |
| Alberti WC code  | 1                | 0.037260   | 0.846937 |

Parameter estimates

| CIMT atherosclerosis or not - Parameter estimates (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |                 |        |          |                |            |                |                |          |
|--|-----------------|--------|----------|----------------|------------|----------------|----------------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |                 |        |          |                |            |                |                |          |
| Modeled probability that CIMT atherosclerosis or not = 1   |                 |        |          |                |            |                |                |          |
| Include condition: v2=1  |                 |        |          |                |            |                |                |          |
| Effect   | Level of Effect | Column | Estimate | Standard Error | Wald Stat. | Lower CL 95. % | Upper CL 95. % | p        |
| Intercept  |                 | 1      | -5.08334 | 2.405962       | 4.463965   | -9.79894       | -0.367741      | 0.034617 |
| AGE  |                 | 2      | 0.09435  | 0.035354       | 7.122030   | 0.02506        | 0.163642       | 0.007614 |
| BMI  |                 | 3      | 0.05966  | 0.081454       | 0.536494   | -0.09998       | 0.219308       | 0.463889 |
| METS (kcal)  |                 | 4      | -0.00018 | 0.000431       | 0.180470   | -0.00103       | 0.000662       | 0.670970 |
| Alberti WC code  | 1               | 5      | 0.07075  | 0.366533       | 0.037260   | -0.64764       | 0.789143       | 0.846937 |
| Scale  |                 |        | 1.00000  | 0.000000       |            | 1.00000        | 1.000000       |          |

Goodness of fit: Hosmer-Lemeshow Test

| CIMT atherosclerosis or not - Goodness of Fit: Hosmer-Lemeshow Test (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |        |        |        |        |        |        |        |        |        |         |          |
|--|--------|--------|--------|--------|--------|--------|--------|--------|--------|---------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |        |        |        |        |        |        |        |        |        |         |          |
| Hosmer Lemeshow = 10.485, p value = 0.2325   |        |        |        |        |        |        |        |        |        |         |          |
| Include condition: v2=1  |        |        |        |        |        |        |        |        |        |         |          |
| Response   | Group1 | Group2 | Group3 | Group4 | Group5 | Group6 | Group7 | Group8 | Group9 | Group10 | Row Tot. |
| 0: Observed  | 6.00   | 4.00   | 3.00   | 6.00   | 4.00   | 4.00   | 1.00   | 4.00   | 4.00   | 2.0     | 38.0     |
| Expected   | 5.80   | 4.95   | 4.64   | 4.18   | 3.69   | 3.26   | 3.02   | 2.69   | 2.39   | 3.4     |          |
| 1: Observed  | 1.00   | 3.00   | 4.00   | 1.00   | 3.00   | 3.00   | 6.00   | 3.00   | 3.00   | 14.0    | 41.0     |
| Expected   | 1.20   | 2.05   | 2.36   | 2.82   | 3.31   | 3.74   | 3.98   | 4.31   | 4.61   | 12.6    |          |
| All Groups   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 16.0    | 79.0     |



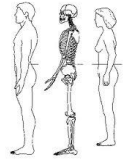
ℓ **Neural Networks Statistics: JSC males**

Summary of active networks

| Summary of active networks (CIMT_NN_data.sta) |           |                |            |                  |                    |                |                   |                   |
|---|-----------|----------------|------------|------------------|--------------------|----------------|-------------------|-------------------|
| Index   | Net. name | Training perf. | Test perf. | Validation perf. | Training algorithm | Error function | Hidden activation | Output activation |
| 8   | MLP 5-5-2 | 71.92982       | 36.36364   | 72.72727         | BFGS 0             | Entropy        | Tanh              | Softmax           |

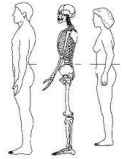
Classification summary

|             | CIMT atherosclerosis or not (Classification summary) (CIMT_NN_data.sta) Samples: Train, Test, Validation |                               |                               |                                 |
|-------------|--|-------------------------------|-------------------------------|---------------------------------|
|             |  | CIMT atherosclerosis or not-1 | CIMT atherosclerosis or not-2 | CIMT atherosclerosis or not-All |
| 8.MLP 5-5-2 | Total  | 41.00000                      | 38.00000                      | 79.00000                        |
|             | Correct  | 28.00000                      | 25.00000                      | 53.00000                        |
|             | Incorrect  | 13.00000                      | 13.00000                      | 26.00000                        |
|             | Correct (%)  | 68.29268                      | 65.78947                      | 67.08861                        |
|             | Incorrect (%)  | 31.70732                      | 34.21053                      | 32.91139                        |



## Network weights

| Weight ID | Network weights (CIMT_NN_data.sta)        |                              |
|-----------|---|------------------------------|
|           | Connections<br>8.MLP 5-5-2                | Weight values<br>8.MLP 5-5-2 |
| 1         | Manne LOG Age --> hidden neuron 1         | -1.04653                     |
| 2         | Manne LOG BMI --> hidden neuron 1         | -0.83081                     |
| 3         | Manne LOG METS (kcal) --> hidden neuron 1 | -0.26479                     |
| 4         | Alberti WC code(1) --> hidden neuron 1    | 0.51339                      |
| 5         | Alberti WC code(2) --> hidden neuron 1    | 0.26076                      |
| 6         | Manne LOG Age --> hidden neuron 2         | 2.19015                      |
| 7         | Manne LOG BMI --> hidden neuron 2         | 0.88337                      |
| 8         | Manne LOG METS (kcal) --> hidden neuron 2 | -0.26245                     |
| 9         | Alberti WC code(1) --> hidden neuron 2    | 0.59248                      |
| 10        | Alberti WC code(2) --> hidden neuron 2    | -0.62524                     |
| 11        | Manne LOG Age --> hidden neuron 3         | -0.93239                     |
| 12        | Manne LOG BMI --> hidden neuron 3         | -0.43454                     |
| 13        | Manne LOG METS (kcal) --> hidden neuron 3 | 0.02257                      |
| 14        | Alberti WC code(1) --> hidden neuron 3    | -0.11548                     |
| 15        | Alberti WC code(2) --> hidden neuron 3    | 0.58931                      |
| 16        | Manne LOG Age --> hidden neuron 4         | -0.91793                     |
| 17        | Manne LOG BMI --> hidden neuron 4         | -0.55536                     |
| 18        | Manne LOG METS (kcal) --> hidden neuron 4 | -0.39002                     |
| 19        | Alberti WC code(1) --> hidden neuron 4    | -0.77518                     |
| 20        | Alberti WC code(2) --> hidden neuron 4    | 0.91287                      |
| 21        | Manne LOG Age --> hidden neuron 5         | -0.04149                     |
| 22        | Manne LOG BMI --> hidden neuron 5         | -0.01497                     |
| 23        | Manne LOG METS (kcal) --> hidden neuron 5 | 0.09211                      |
| 24        | Alberti WC code(1) --> hidden neuron 5    | 0.17856                      |
| 25        | Alberti WC code(2) --> hidden neuron 5    | 0.25215                      |



|    |  |          |
|----|--|----------|
| 26 | input bias --> hidden neuron 1                     | 0.86992  |
| 27 | input bias --> hidden neuron 2                     | -0.03395 |
| 28 | input bias --> hidden neuron 3                     | 0.41882  |
| 29 | input bias --> hidden neuron 4                     | 0.13544  |
| 30 | input bias --> hidden neuron 5                     | 0.36291  |
| 31 | hidden neuron 1 --> CIMT atherosclerosis or not(1) | -1.68511 |
| 32 | hidden neuron 2 --> CIMT atherosclerosis or not(1) | 1.47265  |
| 33 | hidden neuron 3 --> CIMT atherosclerosis or not(1) | -0.87211 |
| 34 | hidden neuron 4 --> CIMT atherosclerosis or not(1) | 1.05080  |
| 35 | hidden neuron 5 --> CIMT atherosclerosis or not(1) | -0.81870 |
| 36 | hidden neuron 1 --> CIMT atherosclerosis or not(2) | 1.68755  |
| 37 | hidden neuron 2 --> CIMT atherosclerosis or not(2) | -1.35631 |
| 38 | hidden neuron 3 --> CIMT atherosclerosis or not(2) | 0.88307  |
| 39 | hidden neuron 4 --> CIMT atherosclerosis or not(2) | -1.06426 |
| 40 | hidden neuron 5 --> CIMT atherosclerosis or not(2) | 0.78536  |
| 41 | hidden bias --> CIMT atherosclerosis or not(1)     | -0.38666 |
| 42 | hidden bias --> CIMT atherosclerosis or not(2)     | 0.37702  |



ℓ Logistic Regression: NPM males

Test of all effects

|   |                  | CIMT atherosclerosis or not - Test of all effects (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |          |  |
|---|------------------|--|----------|--|
|   |                  | Distribution : BINOMIAL, Link function: LOGIT  |          |  |
|   |                  | Modeled probability that CIMT atherosclerosis or not = 1   |          |  |
|   |                  | Include condition: v2=1  |          |  |
| Effect                                  | Degr. of Freedom | Wald Stat.   | p        |  |
| Intercept                               | 1                | 4.808051   | 0.028327 |  |
| AGE                                     | 1                | 6.895966   | 0.008639 |  |
| BMI                                     | 1                | 0.639256   | 0.423980 |  |
| METS (kcal)                             | 1                | 0.175672   | 0.675119 |  |
| Sphyg BP ROC male >=90cm en fem >= 98cm | 1                | 0.021298   | 0.883970 |  |

Parameter estimates

|   |                 | CIMT atherosclerosis or not - Parameter estimates (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |          |                |            |                |                |          |
|---|-----------------|--|----------|----------------|------------|----------------|----------------|----------|
|   |                 | Distribution : BINOMIAL, Link function: LOGIT  |          |                |            |                |                |          |
|   |                 | Modeled probability that CIMT atherosclerosis or not = 1   |          |                |            |                |                |          |
|   |                 | Include condition: v2=1  |          |                |            |                |                |          |
| Effect                                  | Level of Effect | Column   | Estimate | Standard Error | Wald Stat. | Lower CL 95. % | Upper CL 95. % | p        |
| Intercept                               |                 | 1  | -5.16527 | 2.355639       | 4.808051   | -9.78224       | -0.548305      | 0.028327 |
| AGE                                     |                 | 2  | 0.09384  | 0.035734       | 6.895966   | 0.02380        | 0.163874       | 0.008639 |
| BMI                                     |                 | 3  | 0.06280  | 0.078540       | 0.639256   | -0.09114       | 0.216730       | 0.423980 |
| METS (kcal)                             |                 | 4  | -0.00018 | 0.000432       | 0.175672   | -0.00103       | 0.000665       | 0.675119 |
| Sphyg BP ROC male >=90cm en fem >= 98cm | 1               | 5  | 0.05103  | 0.349649       | 0.021298   | -0.63427       | 0.736326       | 0.883970 |
| Scale                                   |                 |  | 1.00000  | 0.000000       |            | 1.00000        | 1.000000       |          |

Goodness of fit: Hosmer-Lemeshow Test

|             |        | CIMT atherosclerosis or not - Goodness of Fit: Hosmer-Lemeshow Test (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |        |        |        |        |        |        |        |         |          |  |
|-------------|--------|--|--------|--------|--------|--------|--------|--------|--------|---------|----------|--|
|             |        | Distribution : BINOMIAL, Link function: LOGIT  |        |        |        |        |        |        |        |         |          |  |
|             |        | Hosmer Lemeshow = 6.7748, p value = 0.5611   |        |        |        |        |        |        |        |         |          |  |
|             |        | Include condition: v2=1  |        |        |        |        |        |        |        |         |          |  |
| Response    | Group1 | Group2   | Group3 | Group4 | Group5 | Group6 | Group7 | Group8 | Group9 | Group10 | Row Tot. |  |
| 0: Observed | 6.00   | 4.00   | 3.00   | 6.00   | 4.00   | 3.00   | 2.00   | 3.00   | 4.00   | 3.00    | 38.0     |  |
| Expected    | 5.81   | 4.95   | 4.64   | 4.19   | 3.66   | 3.28   | 3.01   | 2.67   | 2.40   | 3.4     |          |  |
| 1: Observed | 1.00   | 3.00   | 4.00   | 1.00   | 3.00   | 4.00   | 5.00   | 4.00   | 3.00   | 13.0    | 41.0     |  |
| Expected    | 1.19   | 2.05   | 2.36   | 2.81   | 3.34   | 3.72   | 3.99   | 4.33   | 4.60   | 12.6    |          |  |
| All Groups  | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 7.00   | 16.0    | 79.0     |  |



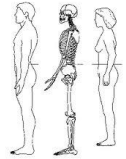
ℓ **Neural Networks Statistics: NPM males**

Summary of active networks

| Summary of active networks (CIMT_NN_data.sta) |           |                |            |                  |                    |                |                   |                   |
|---|-----------|----------------|------------|------------------|--------------------|----------------|-------------------|-------------------|
| Index   | Net. name | Training perf. | Test perf. | Validation perf. | Training algorithm | Error function | Hidden activation | Output activation |
| 3   | MLP 5-3-2 | 64.91228       | 36.36364   | 81.81818         | BFGS 0             | Entropy        | Tanh              | Softmax           |

Classification summary

|             | CIMT atherosclerosis or not (Classification summary) (CIMT_NN_data.sta) Samples: Train, Test, Validation |                               |                               |                                 |
|-------------|--|-------------------------------|-------------------------------|---------------------------------|
|             |  | CIMT atherosclerosis or not-1 | CIMT atherosclerosis or not-2 | CIMT atherosclerosis or not-All |
| 3.MLP 5-3-2 | Total  | 41.00000                      | 38.00000                      | 79.00000                        |
|             | Correct  | 27.00000                      | 23.00000                      | 50.00000                        |
|             | Incorrect  | 14.00000                      | 15.00000                      | 29.00000                        |
|             | Correct (%)  | 65.85366                      | 60.52632                      | 63.29114                        |
|             | Incorrect (%)  | 34.14634                      | 39.47368                      | 36.70886                        |



## Network weights

| Weight ID | Network weights (CIMT_NN_data.sta)                             |                              |
|-----------|--|------------------------------|
|           | Connections<br>3.MLP 5-3-2                                     | Weight values<br>3.MLP 5-3-2 |
| 1         | Manne LOG Age --> hidden neuron 1                              | -1.97776                     |
| 2         | Manne LOG BMI --> hidden neuron 1                              | -0.23588                     |
| 3         | Manne LOG METS (kcal) --> hidden neuron 1                      | -0.02218                     |
| 4         | Sphyg BP ROC male >=90cm en fem >= 98cm(1) --> hidden neuron 1 | -1.03776                     |
| 5         | Sphyg BP ROC male >=90cm en fem >= 98cm(2) --> hidden neuron 1 | 1.61465                      |
| 6         | Manne LOG Age --> hidden neuron 2                              | -0.87175                     |
| 7         | Manne LOG BMI --> hidden neuron 2                              | -0.25618                     |
| 8         | Manne LOG METS (kcal) --> hidden neuron 2                      | -0.03557                     |
| 9         | Sphyg BP ROC male >=90cm en fem >= 98cm(1) --> hidden neuron 2 | 0.42780                      |
| 10        | Sphyg BP ROC male >=90cm en fem >= 98cm(2) --> hidden neuron 2 | -0.13289                     |
| 11        | Manne LOG Age --> hidden neuron 3                              | -0.91978                     |
| 12        | Manne LOG BMI --> hidden neuron 3                              | -0.05704                     |
| 13        | Manne LOG METS (kcal) --> hidden neuron 3                      | 0.38179                      |
| 14        | Sphyg BP ROC male >=90cm en fem >= 98cm(1) --> hidden neuron 3 | -0.26287                     |
| 15        | Sphyg BP ROC male >=90cm en fem >= 98cm(2) --> hidden neuron 3 | 0.60379                      |
| 16        | input bias --> hidden neuron 1                                 | 0.59519                      |
| 17        | input bias --> hidden neuron 2                                 | 0.44966                      |
| 18        | input bias --> hidden neuron 3                                 | 0.29552                      |
| 19        | hidden neuron 1 --> CIMT atherosclerosis or not(1)             | -1.07939                     |
| 20        | hidden neuron 2 --> CIMT atherosclerosis or not(1)             | -2.77320                     |
| 21        | hidden neuron 3 --> CIMT atherosclerosis or not(1)             | 0.12390                      |
| 22        | hidden neuron 1 --> CIMT atherosclerosis or not(2)             | 0.94065                      |
| 23        | hidden neuron 2 --> CIMT atherosclerosis or not(2)             | 2.74530                      |
| 24        | hidden neuron 3 --> CIMT atherosclerosis or not(2)             | -0.03638                     |
| 25        | hidden bias --> CIMT atherosclerosis or not(1)                 | -0.39581                     |
| 26        | hidden bias --> CIMT atherosclerosis or not(2)                 | 0.47721                      |



ℓ Logistic Regression: JSC females

Test of all effects

| CIMT atherosclerosis or not - Test of all effects (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |                  |            |          |
|--|------------------|------------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |                  |            |          |
| Modeled probability that CIMT atherosclerosis or not = 1   |                  |            |          |
| Include condition: v2=2  |                  |            |          |
| Effect   | Degr. of Freedom | Wald Stat. | p        |
| Intercept  | 1                | 15.64418   | 0.000076 |
| AGE  | 1                | 20.62929   | 0.000006 |
| BMI  | 1                | 0.05629    | 0.812466 |
| METS (kcal)  | 1                | 1.11534    | 0.290925 |
| Alberti WC code  | 1                | 0.14351    | 0.704815 |

Parameter estimates

| CIMT atherosclerosis or not - Parameter estimates (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |                 |        |          |                |            |                |                |          |
|--|-----------------|--------|----------|----------------|------------|----------------|----------------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |                 |        |          |                |            |                |                |          |
| Modeled probability that CIMT atherosclerosis or not = 1   |                 |        |          |                |            |                |                |          |
| Include condition: v2=2  |                 |        |          |                |            |                |                |          |
| Effect   | Level of Effect | Column | Estimate | Standard Error | Wald Stat. | Lower CL 95. % | Upper CL 95. % | p        |
| Intercept  |                 | 1      | -9.03896 | 2.285293       | 15.64418   | -13.5180       | -4.55986       | 0.000076 |
| AGE  |                 | 2      | 0.17923  | 0.039461       | 20.62929   | 0.1019         | 0.25657        | 0.000006 |
| BMI  |                 | 3      | -0.01224 | 0.051604       | 0.05629    | -0.1134        | 0.08890        | 0.812466 |
| METS (kcal)  |                 | 4      | 0.00057  | 0.000536       | 1.11534    | -0.0005        | 0.00162        | 0.290925 |
| Alberti WC code  | 1               | 5      | -0.14339 | 0.378497       | 0.14351    | -0.8852        | 0.59846        | 0.704815 |
| Scale  |                 |        | 1.00000  | 0.000000       |            | 1.0000         | 1.00000        |          |

Goodness of fit: Hosmer-Lemeshow Test

| CIMT atherosclerosis or not - Goodness of Fit: Hosmer-Lemeshow Test (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |        |        |        |        |        |        |        |        |        |         |          |
|--|--------|--------|--------|--------|--------|--------|--------|--------|--------|---------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |        |        |        |        |        |        |        |        |        |         |          |
| Hosmer Lemeshow = 5.6976, p value = 0.6810   |        |        |        |        |        |        |        |        |        |         |          |
| Include condition: v2=2  |        |        |        |        |        |        |        |        |        |         |          |
| Response   | Group1 | Group2 | Group3 | Group4 | Group5 | Group6 | Group7 | Group8 | Group9 | Group10 | Row Tot. |
| 0: Observed  | 8.00   | 8.00   | 6.00   | 6.00   | 6.00   | 3.00   | 3.00   | 2.00   | 0.00   | 2.00    | 44.0     |
| Expected   | 7.95   | 7.31   | 6.71   | 5.98   | 5.20   | 4.07   | 3.05   | 1.85   | 1.21   | 0.67    |          |
| 1: Observed  | 1.00   | 1.00   | 3.00   | 3.00   | 3.00   | 6.00   | 6.00   | 7.00   | 9.00   | 7.00    | 46.0     |
| Expected   | 1.05   | 1.69   | 2.29   | 3.02   | 3.80   | 4.93   | 5.95   | 7.15   | 7.79   | 8.33    |          |
| All Groups   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00    | 90.0     |



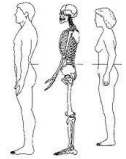
## ℓ Neural Networks Statistics: JSC females

Summary of active networks

| Summary of active networks (CIMT_NN_data.sta) |           |                |            |                  |                    |                |                   |                   |
|---|-----------|----------------|------------|------------------|--------------------|----------------|-------------------|-------------------|
| Index   | Net. name | Training perf. | Test perf. | Validation perf. | Training algorithm | Error function | Hidden activation | Output activation |
| 1   | MLP 5-3-2 | 71.87500       | 92.30769   | 92.30769         | BFGS 22            | Entropy        | Tanh              | Softmax           |

Classification summary

|             | CIMT atherosclerosis or not (Classification summary) (CIMT_NN_data.sta) Samples: Train, Test, Validation |                               |                               |                                 |
|-------------|--|-------------------------------|-------------------------------|---------------------------------|
|             |  | CIMT atherosclerosis or not-1 | CIMT atherosclerosis or not-2 | CIMT atherosclerosis or not-All |
| 1.MLP 5-3-2 | Total  | 46.00000                      | 44.00000                      | 90.00000                        |
|             | Correct  | 33.00000                      | 37.00000                      | 70.00000                        |
|             | Incorrect  | 13.00000                      | 7.00000                       | 20.00000                        |
|             | Correct (%)  | 71.73913                      | 84.09091                      | 77.77778                        |
|             | Incorrect (%)  | 28.26087                      | 15.90909                      | 22.22222                        |



## Network weights

| Weight ID | Network weights (CIMT_NN_data.sta)                 |                              |
|-----------|--|------------------------------|
|           | Connections<br>1.MLP 5-3-2                         | Weight values<br>1.MLP 5-3-2 |
| 1         | Vroue LOG Age --> hidden neuron 1                  | -0.193950                    |
| 2         | Vroue LOG BMI --> hidden neuron 1                  | -0.076115                    |
| 3         | Vroue LOG METS (kcal) --> hidden neuron 1          | -0.093386                    |
| 4         | Alberti WC code(1) --> hidden neuron 1             | -0.087266                    |
| 5         | Alberti WC code(2) --> hidden neuron 1             | -0.036088                    |
| 6         | Vroue LOG Age --> hidden neuron 2                  | 0.485818                     |
| 7         | Vroue LOG BMI --> hidden neuron 2                  | 0.010753                     |
| 8         | Vroue LOG METS (kcal) --> hidden neuron 2          | 0.012111                     |
| 9         | Alberti WC code(1) --> hidden neuron 2             | -0.070367                    |
| 10        | Alberti WC code(2) --> hidden neuron 2             | -0.014241                    |
| 11        | Vroue LOG Age --> hidden neuron 3                  | -0.060991                    |
| 12        | Vroue LOG BMI --> hidden neuron 3                  | -0.020891                    |
| 13        | Vroue LOG METS (kcal) --> hidden neuron 3          | -0.059849                    |
| 14        | Alberti WC code(1) --> hidden neuron 3             | 0.038860                     |
| 15        | Alberti WC code(2) --> hidden neuron 3             | 0.351678                     |
| 16        | input bias --> hidden neuron 1                     | -0.184003                    |
| 17        | input bias --> hidden neuron 2                     | -0.146762                    |
| 18        | input bias --> hidden neuron 3                     | 0.381945                     |
| 19        | hidden neuron 1 --> CIMT atherosclerosis or not(1) | 0.173825                     |
| 20        | hidden neuron 2 --> CIMT atherosclerosis or not(1) | 0.447800                     |
| 21        | hidden neuron 3 --> CIMT atherosclerosis or not(1) | 0.042662                     |
| 22        | hidden neuron 1 --> CIMT atherosclerosis or not(2) | -0.081742                    |
| 23        | hidden neuron 2 --> CIMT atherosclerosis or not(2) | -0.599712                    |
| 24        | hidden neuron 3 --> CIMT atherosclerosis or not(2) | 0.085807                     |
| 25        | hidden bias --> CIMT atherosclerosis or not(1)     | -0.030297                    |
| 26        | hidden bias --> CIMT atherosclerosis or not(2)     | 0.014487                     |



ℓ Logistic Regression: NPM females

Test of all effects

| CIMT atherosclerosis or not - Test of all effects (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |                  |            |          |
|--|------------------|------------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |                  |            |          |
| Modeled probability that CIMT atherosclerosis or not = 1   |                  |            |          |
| Include condition: v2=2  |                  |            |          |
| Effect   | Degr. of Freedom | Wald Stat. | p        |
| Intercept  | 1                | 5.93580    | 0.014836 |
| AGE  | 1                | 18.10659   | 0.000021 |
| BMI  | 1                | 1.38870    | 0.238625 |
| METS (kcal)  | 1                | 0.54544    | 0.460188 |
| Sphyg BP ROC male >=90cm en fem >= 98cm  | 1                | 3.17588    | 0.074733 |

Parameter estimates

| CIMT atherosclerosis or not - Parameter estimates (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |                 |        |          |                |            |                |                |          |
|--|-----------------|--------|----------|----------------|------------|----------------|----------------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |                 |        |          |                |            |                |                |          |
| Modeled probability that CIMT atherosclerosis or not = 1   |                 |        |          |                |            |                |                |          |
| Include condition: v2=2  |                 |        |          |                |            |                |                |          |
| Effect   | Level of Effect | Column | Estimate | Standard Error | Wald Stat. | Lower CL 95. % | Upper CL 95. % | p        |
| Intercept  |                 | 1      | -6.19732 | 2.543689       | 5.93580    | -11.1829       | -1.21178       | 0.014836 |
| AGE  |                 | 2      | 0.17421  | 0.040941       | 18.10659   | 0.0940         | 0.25445        | 0.000021 |
| BMI  |                 | 3      | -0.07361 | 0.062468       | 1.38870    | -0.1960        | 0.04882        | 0.238625 |
| METS (kcal)  |                 | 4      | 0.00040  | 0.000541       | 0.54544    | -0.0007        | 0.00146        | 0.460188 |
| Sphyg BP ROC male >=90cm en fem >= 98cm  | 1               | 5      | 0.71011  | 0.398467       | 3.17588    | -0.0709        | 1.49109        | 0.074733 |
| Scale  |                 |        | 1.00000  | 0.000000       |            | 1.0000         | 1.00000        |          |

Goodness of fit: Hosmer-Lemeshow Test

| CIMT atherosclerosis or not - Goodness of Fit: Hosmer-Lemeshow Test (SABPA 2008 FINAL_2010 - 1 JUNE met standardize_2.sta) |        |        |        |        |        |        |        |        |        |         |          |
|--|--------|--------|--------|--------|--------|--------|--------|--------|--------|---------|----------|
| Distribution : BINOMIAL, Link function: LOGIT  |        |        |        |        |        |        |        |        |        |         |          |
| Hosmer Lemeshow = 4.5649, p value = 0.8029   |        |        |        |        |        |        |        |        |        |         |          |
| Include condition: v2=2  |        |        |        |        |        |        |        |        |        |         |          |
| Response   | Group1 | Group2 | Group3 | Group4 | Group5 | Group6 | Group7 | Group8 | Group9 | Group10 | Row Tot. |
| 0: Observed  | 8.00   | 8.00   | 7.00   | 6.00   | 6.00   | 2.00   | 4.00   | 1.00   | 1.00   | 1.00    | 44.0     |
| Expected   | 8.12   | 7.42   | 6.90   | 5.98   | 4.87   | 4.13   | 3.32   | 1.87   | 1.00   | 0.40    |          |
| 1: Observed  | 1.00   | 1.00   | 2.00   | 3.00   | 3.00   | 7.00   | 5.00   | 8.00   | 8.00   | 8.00    | 46.0     |
| Expected   | 0.88   | 1.58   | 2.10   | 3.02   | 4.13   | 4.87   | 5.68   | 7.13   | 8.00   | 8.60    |          |
| All Groups   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00   | 9.00    | 90.0     |



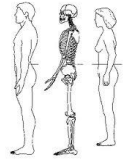
ℓ **Neural Networks Statistics: NPM females**

Summary of active networks

| Summary of active networks (CIMT_NN_data.sta) |           |                |            |                  |                    |                |                   |                   |
|---|-----------|----------------|------------|------------------|--------------------|----------------|-------------------|-------------------|
| Index   | Net. name | Training perf. | Test perf. | Validation perf. | Training algorithm | Error function | Hidden activation | Output activation |
| 7   | MLP 5-3-2 | 75.00000       | 76.92308   | 84.61538         | BFGS 18            | Entropy        | Exponential       | Softmax           |

Classification summary

|             | CIMT atherosclerosis or not (Classification summary) (CIMT_NN_data.sta) Samples: Train, Test, Validation |                               |                               |                                 |
|-------------|--|-------------------------------|-------------------------------|---------------------------------|
|             |  | CIMT atherosclerosis or not-1 | CIMT atherosclerosis or not-2 | CIMT atherosclerosis or not-All |
| 7.MLP 5-3-2 | Total  | 46.00000                      | 44.00000                      | 90.00000                        |
|             | Correct  | 40.00000                      | 29.00000                      | 69.00000                        |
|             | Incorrect  | 6.00000                       | 15.00000                      | 21.00000                        |
|             | Correct (%)  | 86.95652                      | 65.90909                      | 76.66667                        |
|             | Incorrect (%)  | 13.04348                      | 34.09091                      | 23.33333                        |



## Network weights

| Weight ID | Network weights (CIMT_NN_data.sta)                             |                              |
|-----------|--|------------------------------|
|           | Connections<br>7.MLP 5-3-2                                     | Weight values<br>7.MLP 5-3-2 |
| 1         | Vroue LOG Age --> hidden neuron 1                              | -0.27177                     |
| 2         | Vroue LOG BMI --> hidden neuron 1                              | 0.11326                      |
| 3         | Vroue LOG METS (kcal) --> hidden neuron 1                      | 0.52767                      |
| 4         | Sphyg BP ROC male >=90cm en fem >= 98cm(1) --> hidden neuron 1 | -0.88151                     |
| 5         | Sphyg BP ROC male >=90cm en fem >= 98cm(2) --> hidden neuron 1 | 1.10550                      |
| 6         | Vroue LOG Age --> hidden neuron 2                              | 0.56304                      |
| 7         | Vroue LOG BMI --> hidden neuron 2                              | 0.36253                      |
| 8         | Vroue LOG METS (kcal) --> hidden neuron 2                      | 0.13273                      |
| 9         | Sphyg BP ROC male >=90cm en fem >= 98cm(1) --> hidden neuron 2 | 0.63340                      |
| 10        | Sphyg BP ROC male >=90cm en fem >= 98cm(2) --> hidden neuron 2 | 0.29321                      |
| 11        | Vroue LOG Age --> hidden neuron 3                              | -1.35763                     |
| 12        | Vroue LOG BMI --> hidden neuron 3                              | 0.34480                      |
| 13        | Vroue LOG METS (kcal) --> hidden neuron 3                      | -0.04387                     |
| 14        | Sphyg BP ROC male >=90cm en fem >= 98cm(1) --> hidden neuron 3 | -0.35386                     |
| 15        | Sphyg BP ROC male >=90cm en fem >= 98cm(2) --> hidden neuron 3 | 0.82574                      |
| 16        | input bias --> hidden neuron 1                                 | 0.05921                      |
| 17        | input bias --> hidden neuron 2                                 | 0.90131                      |
| 18        | input bias --> hidden neuron 3                                 | 0.46831                      |
| 19        | hidden neuron 1 --> CIMT atherosclerosis or not(1)             | 0.46393                      |
| 20        | hidden neuron 2 --> CIMT atherosclerosis or not(1)             | 0.07446                      |
| 21        | hidden neuron 3 --> CIMT atherosclerosis or not(1)             | -1.90609                     |
| 22        | hidden neuron 1 --> CIMT atherosclerosis or not(2)             | -0.36036                     |
| 23        | hidden neuron 2 --> CIMT atherosclerosis or not(2)             | 0.01802                      |
| 24        | hidden neuron 3 --> CIMT atherosclerosis or not(2)             | 1.79037                      |
| 25        | hidden bias --> CIMT atherosclerosis or not(1)                 | 1.34403                      |
| 26        | hidden bias --> CIMT atherosclerosis or not(2)                 | -1.15977                     |