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Household and familial resemblance in risk factors for type 2 diabetes and related cardiometabolic diseases in rural Uganda: a cross-sectional community sample

Jannie Nielsen, Silver K Bahendeka, Susan R Whyte, Dan W Meyrowitsch, Ib C Bygbjerg, Daniel R Witte

ABSTRACT

Objectives Prevention of type 2 diabetes (T2D) has been successfully established in randomised clinical trials. However, the best methods for the translation of this evidence into effective population-wide interventions remain unclear. To assess whether households could be a target for T2D prevention and screening, we investigated the resemblance of T2D risk factors at household level and by type of familial dyadic relationship in a rural Ugandan community.

Methods This cross-sectional household-based study included 437 individuals ≥13 years of age from 90 rural households in south-western Uganda. Resemblance in glycylated haemoglobin (HbA1c), anthropometry, blood pressure, fitness status and sitting time were analysed using a general mixed model with random effects (by household or dyad) to calculate household intraclass correlation coefficients (ICCs) and dyadic regression coefficients. Logistic regression with household as a random effect was used to calculate the ORs for individuals having a condition or risk factor if another household member had the same condition.

Results The strongest degree of household member resemblances in T2D risk factors was seen in relation to fitness status (ICC=0.24), HbA1c (ICC=0.18) and systolic blood pressure (ICC=0.11). Regarding dyadic resemblance, the highest standardised regression coefficient was seen in fitness status for spouses (0.54, 95% CI 0.32 to 0.76), parent–offspring (0.41, 95% CI 0.28 0.54) and siblings (0.41, 95% CI 0.25 to 0.57). Overall, parent–offspring and sibling pairs were the dyads with strongest resemblance, followed by spouses.

Conclusions The marked degree of resemblance in T2D risk factors at household level and between spouses, parent–offspring and sibling dyads suggest that shared behavioural and environmental factors may influence risk factor levels among cohabiting individuals, which point to the potential of the household setting for screening and prevention of T2D.

INTRODUCTION

Globally, the number of people with diabetes is increasing rapidly, and in sub-Saharan African (SSA) countries like Uganda the numbers will more than double within the next two decades. The majority (90%–95%) of all diabetes is type 2 diabetes (T2D). Prevention or postponement of the onset of T2D in high-risk individuals through a healthy diet, increased physical activity and weight loss has been successfully established in randomised clinical trials from both high-income and middle-income countries. However, it remains unclear as to the best methods for the translation of such clinical proof-of-concept evidence into low-cost effective and feasible population-wide interventions, especially in low-income countries, where access to diabetes diagnostics and treatment is often limited.
In settings where daily life is focused around the family, households may present an opportunity to target several individuals simultaneously. Most of the variation in the risk of T2D in high-income countries is explained by lifestyle and behavioural factors, or by the interaction of lifestyle factors with genetic factors. Household members are likely to share lifestyle behaviours and to some extent genes. Shared daily environment may partly explain the observed resemblance between household members such as spouses in risk factors related to the development of T2D like obesity, exercise levels, raised blood pressure and smoking. Further, spouses of a person with T2D have been shown to have higher fasting plasma glucose and higher risk of developing T2D as compared with individuals with no spousal history of T2D. For familial relations that include a genetic relationship the degree of diabetes risk concordance and resemblance in obesity, glycaemic levels, blood pressure levels and aerobic fitness status are consistently higher than for spouses or adoptees, likely due to a combination of genetic and shared environmental effects.

In SSA, the number of people with diabetes is increasing in both urban and rural areas. However, especially in the rural areas, access to diabetes diagnostics and treatment is very restricted. Thus, novel approaches to low-cost diabetes prevention in such settings are highly needed. In SSA, a family or a household often consists of multiple members and types of relationships (dyads), especially in rural areas. Yet little is known about T2D risk factor resemblance among individuals sharing daily life in a low-income country in epidemiological transition. Therefore, the objective of this study was to investigate resemblance of T2D risk factors at household level and by type of familial dyadic relationship in a rural Ugandan community.

METHODS

Study design and setting
This cross-sectional study was part of a larger study examining households with and without a member with previously diagnosed T2D. Data were collected between December 2012 and March 2013 in Kasese District, Uganda. The district is mountainous and agrarian, though substantial parts may not be cultivated because they are national forest, national park or water bodies. The majority of the approximately 770,000 inhabitants (75.3%) live in rural areas and around 80% is involved in crop production, with small-scale farming being the main occupation for the villagers. The main crops include cassava, sweet potatoes, maize and matoke (plantain), which are also the primary staple foods, and cash crops like coffee. The majority of people live in houses made of mud or sun-dried bricks with an iron sheet roof, no electricity and no piped water. Average household size is 5.3 individuals. Kasese District has three hospitals—one public general hospital (Bwera District Hospital) and two private-not-for-profit hospitals. Diabetes and hypertension diagnostics and treatment are mainly available at hospital level and only free of charge in public facilities. In 2012, the health services were severely understaffed, with only 405 out of 933 positions filled. The doctor-to-patient ratio was 1:43037 and the nurse-to-patient ratio was 1:12 662 as compared with the overall national ratios of 1:24 725 for doctor-to-patients and 1:11 000 for nurse-to-patients.

One hundred households were approached and ninety agreed to participate. Reasons for non-participation were lack of time. Of the 90 households, half included a person diagnosed with T2D, selected from diabetes patient records at diabetes clinic at the nearby hospital. Households without diagnosed T2D were selected using a random sampling plan. To be included in the study, the household should consist of at least two generations, have at least three individuals aged ≥13 years and no member with diagnosed HIV/AIDS, type 1 diabetes or active tuberculosis. Households were defined as people living together and sharing food on a daily basis. All members aged 13 years or above, who had lived in the household for more than 3 months prior to the visit by the survey team were invited to participate (response rate 97.5%). Details of sampling, inclusion and exclusion criteria are described elsewhere.

Ethics
Prior to data collection, the households were visited, the overall aim of the project was verbally explained and an information leaflet was handed out. On the day of data collection, verbal information about the project was given again and the participants were given time to ask questions. Verbal and written consent was obtained from all participants who still agreed to participate. For participants below 18 years of age, written consent was obtained from the caretaker. The study was approved by the Uganda National Council of Science and Technology (ADM 154/212/01), Makerere University School of Medicine Research and Ethics Committee (REC-REF 2012–183), St Francis Hospital Nsambya and Kagando Hospital.

Procedures
After the initial presentation of the study, a household profile was developed, detailing family structure, members, dyads (relationship between every pair of members) and age. Dwelling elevation (metres above sea level) was measured using a Garmin Trex10 (Garmin, Southampton, UK). Haemoglobin A1c (HbA1c) (%) was measured using an Afinion AS100 Analyzer (Axis Shield PoC, Oslo, Norway); values were presented as percentage and converted to mmol/mol. Dysglycaemia was defined as HbA1c ≥42 mmol/mol (≥6%). Blood pressure was measured three times in sitting position after at least 10 min of rest (Omron M6 HEM7211E, Kyoto, Japan). Hypertension was defined as a systolic blood pressure ≥140 mm Hg or a diastolic blood pressure ≥90 mm Hg.
Hg averaged over the last two blood pressure readings. Body weight measured using a flat scale (model 876, SECA, Birmingham, UK) and height measured using a portable stadiometer (model 213, SECA, UK) were used to calculate body mass index (BMI) as weight (kg)/height (m²). Underweight, normal weight, overweight and obesity were defined according to the WHO classifications for adults and for adolescents aged from 13 to 19 years according to WHO Child Growth Standards.

For dyads where one member could be below 19 years of age (parent–offspring, siblings and grandparent–grandchild), a Z-score of height-for-age was calculated and used instead of height (cm) for both dyad members. The Z-score was calculated according to de Onis et al. and individuals ≥19 years of age were handled as the oldest category in the WHO Child Growth Reference.

As a measure of aerobic fitness status, an 8 min step test was conducted to estimate aerobic capacity (maximal oxygen uptake, VO₂ max (mLO₂/min/kg body weight)) and managed according to the Cambridge Protocol. Fifty individuals did not perform/complete at least 4 min of the step test. In data analyses using fitness status as a continuous variable, these individuals were excluded, whereas in data analyses where fitness status was used as a dichotomous variable, the 50 individuals were coded as unfit with the exception of those who had recently given birth or had an acute illness (n=5).

Household socioeconomic status (SES), and individual educational level, age, sex, disease status and smoking were assessed using questionnaires. Daily sitting time was assessed using a locally adapted version of the International Physical Activity Questionnaire.

Statistical analysis

The amount of resemblance in T2D risk factors in individuals living within the same household was assessed calculating intraclass correlation coefficients (ICCs) with general mixed models with household as a random effect, adjusting for sex, age, SES and household size.

Dyadic relationships were restricted to spouses, parent–offspring, grandparent–grandchild and sibling dyads and analysed as distinguishable members based on sex for spousal dyads (husband dyad number 1 and wife dyad number 2), birth order for sibling dyads (oldest sibling dyad number 1) and age for parent–offspring and grandparent–grandchild dyads (parent and grandparents as dyad number 1 respectively). As non-independence was assumed, a mixed model was used to analyse the dyadic resemblance between the same risk factor in the two dyad members. Our primary analyses modelled the risk factors Hba1c, blood pressure, height, BMI, fitness status and sitting time, separately, in dyad member 2 as a function of the same risk factor in dyad member 1. Random effects were dyad member 1 (to account, eg, for a parent having more than one child) or household (to account for more than one of the same type of dyad occurring per household). For dyadic relationships, regression coefficient estimates were reported with 95% confidence intervals (CIs). Logistic regression with household as a random effect was used to calculate the OR of an individual having a condition if someone else in the household had the same condition. ORs are reported with 95% CIs. Explanatory variables were introduced sequentially: individual level (sex, age); dyad level (age difference between the dyad members) and household level (SES, elevation of the dwelling, household size). Statistical significance was set as p<0.05.

For analyses including Hba1c, individuals with diagnosed T2D (n=45) were excluded and for analyses including blood pressure measures, individuals with diagnosed hypertension (n=32) were excluded as medication may have influenced these values. All statistical analyses were performed using Stata V.14.1 SE (StataCorp).

RESULTS

From the 90 households, we identified a total of 947 dyads of which 91 (9.6%) were spouses, 283 (29.8%) were parent–offspring dyads, 97 (10.2%) were grandparent–grandchild dyads and 148 (15.6%) were sibling dyads. The remaining 330 dyads were primarily in-laws and uncle/aunt–nephew/niece dyads (not included in this analysis). General characteristics and cardiometabolic risk factors at household level and by dyadic relationship are summarised in table 1. In 84 (93.3%) households, all meals were eaten within the household. Median dwelling elevation was 1177 m above sea level (range 951–1742 m above sea level).

Household resemblance in T2D risk factors

At household level, ICCs showed statistically significant household member resemblance for four risk factors. After adjustment for age and sex, ICCs were statistically significant for fitness status (ICC=0.24, p<0.001), Hba1c (ICC=0.18, p<0.001), BMI (ICC=0.08, p=0.010) and systolic blood pressure (ICC=0.11, p=0.003), while only a tendency was observed for diastolic blood pressure (ICC=0.06, p=0.06). Additional adjustment for SES, household size or dwelling elevation did not change the ICCs.

Dyad resemblance

Dyad resemblance in T2D risk factors is shown as regression coefficients adjusted for age difference and sex in table 2. Sibling and parent–offspring dyads both had five statistically associated risk factors. Siblings were associated in measures of Hba1c, systolic blood pressure, diastolic blood pressure, height and fitness status, while parent–offspring dyads were associated with in Hba1c, systolic blood pressure, height, fitness status and sitting time. Spouses were statistically significantly associated in systolic blood pressure and fitness status, while grandparent–grandchild dyads were only associated with regard to diastolic blood pressure. None of the four dyad types had a statically significant association for BMI.
### Table 1  General characteristics and cardiometabolic risk factors at household level and by dyadic relationships

<table>
<thead>
<tr>
<th>Members (n unique)</th>
<th>Households (90)</th>
<th>Dyads by type (n)*</th>
<th>Parents–offspring (283)</th>
<th>Offspring (164)</th>
<th>Grandparents–grandchildren (97)</th>
<th>Grandparents (64)</th>
<th>Grandchildren (64)</th>
<th>Siblings (148)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Individuals in household</td>
<td>5 (range 3–10)</td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Age (years)†</td>
<td>38.7 (33.0; 43.1)</td>
<td>51.0 (42.7; 57.4)</td>
<td>56.1 (49.6; 67.4)</td>
<td>53.7 (47.6; 63.1)</td>
<td>60.4 (52.6; 68.4)</td>
<td>21.3 (15.8; 30.7)</td>
<td>19.7 (16.0; 26.7)</td>
<td>67.5 (56.6; 72.7)</td>
</tr>
<tr>
<td>Age difference (years)‡</td>
<td>46.0 (38.7; 4.0)</td>
<td>6.1 (3.0; 11.0)</td>
<td>32.6 (27.0; 38.2)</td>
<td>50.2 (41.7; 56.0)</td>
<td>4.8 (3.0; 8.0)</td>
<td></td>
<td></td>
<td></td>
</tr>
<tr>
<td>Diagnosed T2D, n (%)</td>
<td>45 (50)</td>
<td>9 (9.9)</td>
<td>29 (32.6)</td>
<td>12 (13.6)</td>
<td>23 (37.1)</td>
<td>2 (2.5)</td>
<td>1 (1.2)</td>
<td>8 (19.5)</td>
</tr>
<tr>
<td>Dysglycaemia, n (%)</td>
<td>22 (24.4)</td>
<td>12 (13.2)</td>
<td>5 (5.6)</td>
<td>11 (12.5)</td>
<td>4 (6.5)</td>
<td>5 (6.2)</td>
<td>1 (1.2)</td>
<td>6 (14.6)</td>
</tr>
<tr>
<td>Diagnosed hypertension, n (%)</td>
<td>28 (31.1)</td>
<td>10 (11.0)</td>
<td>14 (15.7)</td>
<td>12 (13.6)</td>
<td>10 (16.1)</td>
<td>0 (0)</td>
<td>0 (0)</td>
<td>12 (29.3)</td>
</tr>
<tr>
<td>Undiagnosed hypertension, n (%)</td>
<td>67 (74.4)</td>
<td>16 (17.6)</td>
<td>23 (26.1)</td>
<td>17 (19.5)</td>
<td>20 (32.8)</td>
<td>3 (3.7)</td>
<td>3 (3.6)</td>
<td>8 (19.5)</td>
</tr>
<tr>
<td>HbA1c (%) (mmol/mol)§</td>
<td>5.4 (0.3)</td>
<td>35.4 (3.2)</td>
<td>36.9 (5.5)</td>
<td>35.8 (5.9)</td>
<td>37.5 (5.6)</td>
<td>36.4 (8.5)</td>
<td>35.0 (4.4)</td>
<td>34.6 (5.7)</td>
</tr>
<tr>
<td>Systolic blood pressure (mm Hg)†</td>
<td>128 (123; 137)</td>
<td>134 (117; 149)</td>
<td>139 (127; 164)</td>
<td>137 (118; 155)</td>
<td>151 (132; 170)</td>
<td>118 (112; 129)</td>
<td>124 (117; 131)</td>
<td>152 (134; 167)</td>
</tr>
<tr>
<td>Diastolic blood pressure (mm Hg)†</td>
<td></td>
<td>78 (74; 82)</td>
<td>80 (74; 89)</td>
<td>84 (76; 92)</td>
<td>84 (79; 91)</td>
<td>87 (79; 95)</td>
<td>75 (71; 81)</td>
<td>72 (69; 80)</td>
</tr>
<tr>
<td>Height (cm)</td>
<td>155.2 (3.8)</td>
<td>152.4 (5.4)</td>
<td>161.3 (6.3)</td>
<td>151.5 (6.1)</td>
<td>160.6 (5.3)</td>
<td>153.5 (6.6)</td>
<td>158.4 (8.4)</td>
<td>150.4 (5.2)</td>
</tr>
<tr>
<td>Short stature, n (%)**††</td>
<td>78 (86.7)</td>
<td>30 (32.8)</td>
<td>49 (55.7)</td>
<td>33 (37.5)</td>
<td>38 (61.3)</td>
<td>19 (23.5)</td>
<td>45 (64.2)</td>
<td>22 (53.7)</td>
</tr>
<tr>
<td>BMI††</td>
<td>21.7 (20.7; 23.2)</td>
<td>23.1 (21.0; 25.3)</td>
<td>21.5 (19.6; 25.0)</td>
<td>22.4 (19.7; 24.6)</td>
<td>22.1 (19.6; 25.4)</td>
<td>22.4 (20.2; 24.4)</td>
<td>20.4 (19.0; 21.5)</td>
<td>21.2 (18.3; 24.7)</td>
</tr>
<tr>
<td>Underweight, n (%)</td>
<td>27 (30.0)</td>
<td>9 (9.9)</td>
<td>8 (9.1)</td>
<td>13 (14.8)</td>
<td>6 (9.7)</td>
<td>5 (6.2)</td>
<td>2 (2.4)</td>
<td>11 (26.8)</td>
</tr>
<tr>
<td>Overweight, n (%)</td>
<td>52 (57.8)</td>
<td>25 (27.5)</td>
<td>24 (27.3)</td>
<td>19 (21.6)</td>
<td>20 (32.3)</td>
<td>19 (23.5)</td>
<td>5 (6.0)</td>
<td>10 (24.4)</td>
</tr>
<tr>
<td>Fitness status (VO₂-max: mL O₂/min/ kg)††</td>
<td>38.5 (4.7)</td>
<td>34.3 (6.8)</td>
<td>33.5 (8.8)</td>
<td>32.0 (6.7)</td>
<td>32.7 (7.2)</td>
<td>37.3 (6.6)</td>
<td>44.6 (6.8)</td>
<td>29.1 (6.8)</td>
</tr>
<tr>
<td>Unfit, n (%)§§</td>
<td>72 (80.0)</td>
<td>35 (40.2)</td>
<td>39 (44.3)</td>
<td>43 (49.4)</td>
<td>25 (41.0)</td>
<td>24 (29.6)</td>
<td>30 (36.1)</td>
<td>30 (75.0)</td>
</tr>
</tbody>
</table>

Continued
<table>
<thead>
<tr>
<th>Members (n unique)</th>
<th>Housesolds (90)</th>
<th>Spouses (91)</th>
<th>Parents–offspring (283)</th>
<th>Offspring (164)</th>
<th>Grandparents–grandchildren (97)</th>
<th>Grandparents (64)</th>
<th>Grandchildren (64)</th>
<th>Siblings (148)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Sitting per day (min)†</td>
<td>275.4 (225; 310)</td>
<td>208 (169; 279)</td>
<td>274 (189; 351)</td>
<td>231 (189; 334)</td>
<td>283 (197; 373)</td>
<td>261 (204; 326)</td>
<td>240 (189; 369)</td>
<td>274 (197; 380)</td>
</tr>
<tr>
<td>Smoking status, n (%)</td>
<td></td>
<td></td>
<td></td>
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<td></td>
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<td></td>
<td></td>
</tr>
<tr>
<td>Never smoked</td>
<td>52 (57.8)</td>
<td>71 (78.0)</td>
<td>59 (66.3)</td>
<td>57 (64.8)</td>
<td>39 (62.9)</td>
<td>77 (95.1)</td>
<td>68 (81.9)</td>
<td>22 (53.7)</td>
</tr>
<tr>
<td>Former smoker</td>
<td>38 (42.2)</td>
<td>11 (12.1)</td>
<td>21 (23.6)</td>
<td>19 (21.6)</td>
<td>17 (27.4)</td>
<td>2 (2.5)</td>
<td>9 (10.8)</td>
<td>13 (31.7)</td>
</tr>
<tr>
<td>Current smoker</td>
<td>21 (23.3)</td>
<td>9 (9.9)</td>
<td>9 (10.1)</td>
<td>12 (13.6)</td>
<td>6 (9.7)</td>
<td>2 (2.5)</td>
<td>6 (7.2)</td>
<td>6 (14.6)</td>
</tr>
<tr>
<td>Years of education†</td>
<td>5.3 (2.2)</td>
<td>2 (0.6)</td>
<td>6 (3.7)</td>
<td>2 (0.4)</td>
<td>5 (1.7)</td>
<td>6 (5.10)</td>
<td>7 (5.10)</td>
<td>0 (0.2)</td>
</tr>
</tbody>
</table>

Data are presented as mean (SD). *Please note that for the different types of dyads there can be a different number of dyad member 1 and dyad member 2 as for example, one husband had two wives or a mother can have more than one child. †Median (p25; p75). ††A household level, age difference is between the oldest and youngest individual in the household. §Individuals with diagnosed diabetes are excluded. ¶Individuals with diagnosed hypertension are excluded. **Short stature is defined as age (months) Z-score below –2SD or final height for males below 161.9 cm and for females below 150.1 cm. †††Missing value on one man. §§Data missing on six individuals because of pregnancy, recent delivery, sickness or technical error. BMI, body mass index; HbA1c, glycosylated haemoglobin; T2D, type 2 diabetes.
Table 2  Dyad regression coefficients for type 2 diabetes risk factors (adjusted for age difference and sex)

<table>
<thead>
<tr>
<th>(n)</th>
<th>Spouses (91)</th>
<th>Parents–offspring (283)</th>
<th>Grandparents–grandchildren (97)</th>
<th>Siblings (148)</th>
</tr>
</thead>
<tbody>
<tr>
<td>HbA1c (%)†</td>
<td>0.18 (−0.09 to 0.45)</td>
<td>0.16* (0.02 to 0.29)</td>
<td>0.07 (−0.8 to 0.22)</td>
<td>0.28* (0.13 to 0.44)</td>
</tr>
<tr>
<td>Systolic blood pressure (mm Hg)‡</td>
<td>0.27* (0.01 to 0.53)</td>
<td>0.10* (0.04 to 0.16)</td>
<td>0.08 (−0.02 to 0.19)</td>
<td>0.18* (0.01 to 0.36)</td>
</tr>
<tr>
<td>Diastolic blood pressure (mm Hg)‡</td>
<td>0.10 (−0.13 to 0.34)</td>
<td>0.02 (−0.07 to 0.10)</td>
<td>0.14* (0.02 to 0.27)</td>
<td>0.16* (0.01 to 0.32)</td>
</tr>
<tr>
<td>Height (cm or SD)§</td>
<td>0.07 (−0.13 to 0.26)</td>
<td>0.35* (0.19 to 0.52)</td>
<td>0.10 (−0.17 to 0.38)</td>
<td>0.26* (0.09 to 0.42)</td>
</tr>
<tr>
<td>BMI (kg/m²)</td>
<td>0.19 (−0.04 to 0.42)</td>
<td>0.02 (−0.07 to 0.12)</td>
<td>−0.01 (−0.14 to 0.13)</td>
<td>0.11 (−0.06 to 0.29)</td>
</tr>
<tr>
<td>Fitness status (mLO2/min/kg)¶</td>
<td>0.42* (0.25 to 0.59)</td>
<td>0.46* (0.31 to 0.60)</td>
<td>−0.08 (−0.37 to 0.20)</td>
<td>0.38* (0.22 to 0.53)</td>
</tr>
<tr>
<td>Daily sitting time (minutes)</td>
<td>0.09 (−0.05 to 0.24)</td>
<td>0.15* (0.04 to 0.27)</td>
<td>0.10 (−0.07 to 0.27)</td>
<td>0.09 (−0.08 to 0.27)</td>
</tr>
</tbody>
</table>

Values are presented as regression coefficients (95% CI). Coefficients express the difference in each risk factor in dyad member 2 per unit difference in that same risk factor in dyad member 1.

* p<0.05.
† Individuals with diagnosed diabetes were excluded.
‡ Individuals with diagnosed hypertension were excluded.
§ For spouses, height (cm) is used while for parents–offspring, grandparents–grandchildren and siblings, height for age is used and not adjusted for age difference or sex.
¶ In 15% of the dyads, one member did not complete the step test.
BMI, body mass index; HbA1c, haemoglobin A1c.

Standardised regression coefficients are shown in table 3. For spouses, parent–offspring and sibling dyads, the standardised regression coefficients were highest for fitness status.

Concordance in risk factors
The results of the logistic regression models are shown in table 4. At household level, effect estimates showed that if one member in the household had dysglycaemia, the OR of another household member having the same status was increased almost 20 times. Having diagnosed hypertension in the household increased the odds of another member having diagnosed or undiagnosed hypertension 2.6 times, whereas undiagnosed hypertension increased the odds of diagnosed or undiagnosed hypertension in another member 4.8 times. The ORs of being overweight or obese, underweight, unfit, smoker or former smoker were all statistically significantly higher if another member of the household had the same status as compared with if no one in the household had the same status (table 4).

DISCUSSION
The results of the present study indicate that individuals living in the same household in rural Uganda share risk factors for T2D and cardiometabolic diseases. We showed that, in particular for systolic blood pressure and fitness status, the spousal association was at least as strong as the association between siblings or parent–offspring pairs.

Table 3  Standardised regression coefficients for type 2 diabetes risk factors (adjusted for age difference and sex)

<table>
<thead>
<tr>
<th></th>
<th>Spouses (91)</th>
<th>Parents–offspring (283)</th>
<th>Grandparents–grandchildren (97)</th>
<th>Siblings (148)</th>
</tr>
</thead>
<tbody>
<tr>
<td>HbA1c†</td>
<td>0.19 (−0.11 to 0.50)</td>
<td>0.21* (0.02 to 0.40)</td>
<td>0.12 (−0.13 to 0.37)</td>
<td>0.26* (0.11 to 0.42)</td>
</tr>
<tr>
<td>Systolic blood pressure‡</td>
<td>0.28* (0.01 to 0.54)</td>
<td>0.20* (0.08 to 0.33)</td>
<td>0.22 (−0.06 to 0.50)</td>
<td>0.20* (0.01 to 0.39)</td>
</tr>
<tr>
<td>Diastolic blood pressure‡</td>
<td>0.10 (−0.14 to 0.35)</td>
<td>0.02 (−0.11 to 0.15)</td>
<td>0.27* (0.03 to 0.05)</td>
<td>0.20* (0.01 to 0.39)</td>
</tr>
<tr>
<td>Height for age§</td>
<td>0.07 (−0.13 to 0.28)</td>
<td>0.26* (0.14 to 0.37)</td>
<td>0.08 (−0.13 to 0.31)</td>
<td>0.26* (0.09 to 0.42)</td>
</tr>
<tr>
<td>BMI</td>
<td>0.16 (−0.04 to 0.37)</td>
<td>0.02 (−0.11 to 0.16)</td>
<td>0.02 (−0.20 to 0.24)</td>
<td>0.14 (−0.03 to 0.31)</td>
</tr>
<tr>
<td>VO₂-max¶</td>
<td>0.54* (0.32 to 0.76)</td>
<td>0.41* (0.28 to 0.54)</td>
<td>−0.09 (−0.38 to 0.21)</td>
<td>0.41* (0.25 to 0.57)</td>
</tr>
<tr>
<td>Daily sitting time</td>
<td>0.11 (−0.09 to 0.31)</td>
<td>0.17* (0.04 to 0.32)</td>
<td>0.11 (−0.09 to 0.32)</td>
<td>0.09 (−0.10 to 0.27)</td>
</tr>
</tbody>
</table>

Values are presented as standardised regression coefficients (95% CI).

* p<0.05.
† Individuals with diagnosed diabetes were excluded.
‡ Individuals with diagnosed hypertension were excluded.
§ Not adjusted for age-difference.
¶ In 15% of the dyads, one member did not complete the step test.
BMI, body mass index; HbA1c, haemoglobin A1c.
partly explained by the high heritability of VO\textsubscript{2}\text{-max}.\cite{21}

siblings. The high resemblance in fitness status is highest ICC at household level and standardised regression on a single type of dyad\cite{19,20} or a single type of risk factor.\cite{35} We are not aware of studies from low-income countries investigating household or dyad resemblance in risk factors for T2D.\cite{12,39} However, studies of assortative mating and/or convergence over time are often used to explain spousal resemblance in risk factors for T2D.\cite{12,39} However, studies of assortative mating and risk factors for T2D are almost exclusively from high-income settings, and preferences for choice of spouse may differ across geographical, social and ethnic settings. For instance, overweight has traditionally been viewed as a desirable feature in SSA settings,\cite{40} whereas it is more stigmatising in high-income settings.\cite{41} Further, until recently the prevalence of obesity in SSA was low, and results from a Danish study showed a tendency to an increase in assortive marriages between obese spouses along with the obesity epidemic.\cite{39}

In contrast to other studies of genetically related individuals, we did not find a relationship in BMI for parent–offspring\cite{18,42} or sibling dyads.\cite{19,36,43} Concerning parent–offspring, a study from the USA including children from 2 to 16 years of age suggested that pubertal children are less likely to resemble their parents in BMI than prepuberty children, as they grow more independent of parents’ eating and exercise behaviours.\cite{42} This could explain the lack of parent–offspring relationship in our study where some of the parent–offspring dyads included adult offspring. However, stratifying parent–offspring dyads into adolescents and adult offspring or above/below median age difference did not change the lack of statistical associations. In terms of siblings, other studies found that sibling dyads resembled in BMI,\cite{19} but that the sibling BMI correlations were less pronounced during adolescence,\cite{36} decreased with increasing age difference\cite{19} and were higher among home living adolescents than adult siblings living apart.\cite{43} The mean sibling age difference (7 years) in our study was not markedly different from the mentioned studies, and the siblings lived together. Thus, these factors cannot entirely explain the lack of relationship.

The last relationship with a genetic component examined in the present study was grandparent–grandchild dyads. Again no relationship was seen in BMI, which is supported by data from a Korean population,\cite{44} but in

Table 4 OR of having a condition as a function of the disease or risk factor status in other members of the same household (adjusted for age, sex and household size)

<table>
<thead>
<tr>
<th>Exposure status</th>
<th>Outcome</th>
<th>Household level</th>
</tr>
</thead>
<tbody>
<tr>
<td>Diagnosed diabetes</td>
<td>Dysglycaemia</td>
<td>0.8 (0.4 to 2.0)*</td>
</tr>
<tr>
<td>Dysglycaemia</td>
<td></td>
<td>19.8 (11.0 to 35.5)*</td>
</tr>
<tr>
<td>Diagnosed hypertension</td>
<td>Diagnosed or undiagnosed hypertension</td>
<td>2.6 (1.5 to 4.5)*</td>
</tr>
<tr>
<td>Undiagnosed hypertension</td>
<td>Diagnosed or undiagnosed hypertension</td>
<td>4.8 (2.9 to 8.0)*</td>
</tr>
<tr>
<td>Short stature</td>
<td>Short stature</td>
<td>10.9 (6.9 to 17.0)*</td>
</tr>
<tr>
<td>Overweight or obesity</td>
<td>Overweight or obesity</td>
<td>9.0 (6.1 to 13.2)*</td>
</tr>
<tr>
<td>Underweight</td>
<td>Underweight</td>
<td>13.7 (7.1 to 26.3)*</td>
</tr>
<tr>
<td>Unfit</td>
<td>Unfit†</td>
<td>11.2 (7.4 to 17.1)*</td>
</tr>
<tr>
<td>Smoker</td>
<td>Smoker</td>
<td>33.7 (15.8 to 71.8)*</td>
</tr>
<tr>
<td>Former smoker</td>
<td>Former smoker</td>
<td>18.9 (9.4 to 38.0)*</td>
</tr>
</tbody>
</table>

Values are presented as ORs (95%CI). *p<0.05. †Unfit is defined as a fitness level below middle derived from VO\textsubscript{2}\text{-max and grouped according to Astrand.\cite{34}}

indicating an effect of shared lifestyle behaviours. For other cardiometabolic risk factors, the resemblance was more prominent between siblings and parent–offspring dyads, whereas grandparent–grandchild dyads were less alike.

To our knowledge, this is the first study to investigate the resemblance of multiple cardiometabolic risk factors in household clusters including several generations living and eating together on a daily basis. A German study of aerobic fitness found an ICC of 0.22 in fitness status in nuclear families but no association when restricting the analyses to spouses.\cite{35} Our findings of dyad resemblance in HbA1c, blood pressure, height and fitness status are in agreement with other epidemiological studies focusing on a single type of dyad\cite{19,20} or a single type of risk factor.\cite{35,36} We are not aware of studies from low-income countries investigating household or dyad resemblance in risk factors for T2D.

Among the measured risk factors, fitness status had the highest ICC at household level and standardised regression coefficient among spouse, parent–offspring and sibling dyads. The high resemblance in fitness status is partly explained by the high heritability of VO\textsubscript{2}\text{-max}.\cite{21} However, in contrast to the German study,\cite{35} we also found a high association in spousal fitness status suggesting that also shared physical activity patterns may contribute to the high fitness status resemblance in our study population. In the Ugandan situation, a peasant’s wife is most often also a peasant, and offspring help cultivating the family land. Shared daily activities as the explanation for spousal resemblance in fitness status is supported by a French study finding that spouses’ physical activity patterns were only similar during weekend days.\cite{37} In addition, walking was the primary means of transportation for most of the study participants, giving all individuals in the same household the same walking distance and elevation differential when for example, going to the nearest trading centre. However, adjusting for elevation gave only a modest attenuation of the household ICC or the dyad resemblance in fitness status.

In line with the results of a meta-analysis,\cite{38} spouses resembled each other with regard to systolic blood pressure. Contradicting other studies,\cite{10,39} we did not find a statistically significant spousal association for BMI, diastolic blood pressure or HbA1c. Discordance in ethnicity of spouses, low numbers of people living in the household and higher SES have previously been shown to attenuate the spousal association in BMI.\cite{15} However, none of these factors were present or affected the absence of a spousal BMI association on our study. Assortative mating and/or convergence over time are often used to explain spousal resemblance in risk factors for T2D.\cite{12,39} However, studies of assortative mating and risk factors for T2D are almost exclusively from high-income settings, and preferences for choice of spouse may differ across geographical, social and ethnic settings. For instance, overweight has traditionally been viewed as a desirable feature in SSA settings,\cite{40} whereas it is more stigmatising in high-income settings.\cite{41} Further, until recently the prevalence of obesity in SSA was low, and results from a Danish study showed a tendency to an increase in assortive marriages between obese spouses along with the obesity epidemic.\cite{39}
contrast to a study from Belgium finding a direct association in obesity measures through three generations. Neither the Korean nor the Belgian study reported that grandparents and grandchildren lived together, which they did in our study and could have increased the chance of resemblance in BMI. However, Uganda is a country in transition in terms of both disease burden and nutrition. In addition, the Rwenzori Mountain region in Kasese district was the centre of civil strife with a civil war in 1962–1982 and again from 1996 to 2002, making it likely that grandparents and grandchildren were exposed to very different intrauterine environments and growth conditions. This hypothesis is supported by the findings of a statistically significant height increment between each of the three generations in our cohort (data not shown), which was not reported in the study from Korea including three generations. Potential unmeasured confounders for BMI may have been unreported/undiagnosed infectious disease such as tuberculosis or HIV/AIDS: both have a fairly high prevalence in the study setting and both affect body weight.

The high ORs in smoking status may partly be explained by a low overall smoking prevalence (7.6%). Further, 63% of the smokers lived together with at least one other smoker. The high resemblance in smoking status is supported by results of studies finding a high spousal resemblance in smoking status and that both smoking and quitting smoking spread in social ties in social networks.

Strengths and limitations
One of the main strengths of this study is the household-based approach. Visiting the families in the home setting resulted in a high individual response rate (97.5%) and thus only minimal risk of selection bias in dyad representativeness. The cross-sectional design prevents us from concluding on whether the spousal resemblance was due to shared risk behaviours or assortative mating, and for the genetic relationships, we cannot distinguish between shared genes and shared environment/behaviours. The ICCs reflect the proportion of variances, whereby the sizes of the ICCs cannot be compared with other cohorts or settings. Thus, the size of ICCs should only be interpreted as a tool to investigate which risk factors resemble most strongly at the household level in the present cohort. The application of HbA1c as a diagnostic tool in African populations is debated. However, in the present study, HbA1c was used to investigate resemblance in dyad members and not to diagnose diabetes. Due to the initial sampling of this study population, 50% of the households had a member with diagnosed T2D. We have previously shown that having diagnosed T2D in the household may have positive spillover effects on the other members potentially due to changes in diet and physical activity due to the diabetes status. This could explain the difference between diagnosed T2D and dysglycaemia in the household as a risk factor for dysglycaemia in other members of the household.

CONCLUSION
The moderate to strong correlations in T2D risk factors at household level and between spouses, parent–offspring and sibling dyads suggest that shared behavioural and environmental factors such as physical activity may influence the risk factor level among cohabiting individuals. The marked degree of household resemblance for certain T2D risk factors highlights the potential of the household setting for screening and prevention of T2D. Thus, when one household member presents with elevated glucose, blood pressure or physical inactivity, the entire household could benefit from lifestyle interventions.

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Contributors JN developed the study protocol, collected the data, performed the statistical analyses and the interpretation of data, and drafted, revised and finalised the article. SKB contributed to the protocol with substantial knowledge concerning diabetes in Uganda and specifically in Kasese district, took part in the later stage and final interpretation of data, and participated in developing, revising and finalising the manuscript. SRW, DWM and ICB contributed to the development of the study protocol, took part in the later stage and final interpretation of data, and participated in developing, revising and finalising the manuscript. DRW performed the statistical analysis and the interpretation of data, and participated in developing, revising and finalising the manuscript.

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Household and familial resemblance in risk factors for type 2 diabetes and related cardiometabolic diseases in rural Uganda: a cross-sectional community sample

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