1 <u>Title page</u>

- 2 Title: Comparability of children's sedentary time estimates derived from wrist worn GENEActiv and
- 3 hip worn ActiGraph accelerometer thresholds
- 4 Lynne M. Boddy¹, Robert J. Noonan¹, Youngwon Kim², Alex V. Rowlands^{3,4,5} Greg J. Welk⁶ Zoe R.
- 5 Knowles¹, Stuart J. Fairclough^{7,8}
- 6 ¹Physical Activity Exchange, Research Institute for Sport and Exercise Sciences, Liverpool John Moores
- 7 University, Liverpool, UK
- 8 ² MRC Epidemiology Unit, University of Cambridge School of Clinical Medicine, Cambridge
- 9 Biomedical Campus. Cambridge, UK
- ³Diabetes Research Centre, University of Leicester, Leicester General Hospital, Leicester, UK;
- ⁴NIHR Leicester Biomedical Research Centre, UK;
- ⁵Alliance for Research in Exercise, Nutrition and Activity (ARENA), Sansom Institute for Health
- 13 Research, Division of Health Sciences, University of South Australia, Adelaide, Australia.
- 14 ⁶Department of Kinesiology, College of Human Sciences, Iowa State University, Ames, IA
- 15 ⁷Department of Sport and Physical Activity, Edge Hill University, Ormskirk, UK
- 16 ⁸Department of Physical Education and Sport Sciences, University of Limerick, Limerick, Ireland

17	Corresponding author details:	Dr Lynne M. Boddy			
18		The Physical Activity Exchange,			
19		Research Institute for Sport & Exercise Sciences,			
20		Liverpool John Moores University,			
21		62 Great Crosshall Street,			
22		Liverpool,			
23		L3 2AT			
24		UK			
25		Email: L.M.Boddy@ljmu.ac.uk			
26		<u>Tel: +44 (0)151 231 4275</u>			
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34 hip worn ActiGraph accelerometer thresholds

36 Abstract:

Objectives: to examine the comparability of children's free-living sedentary time (ST) derived from raw
acceleration thresholds for wrist mounted GENEActiv accelerometer data, with ST estimated using the
waist mounted ActiGraph 100 count·min⁻¹ threshold.

40 Design: Secondary data analysis

Method: 108 10-11-year-old children (n=43 boys) from Liverpool, UK wore one ActiGraph GT3X+ and one GENEActiv accelerometer on their right hip and left wrist, respectively for seven days. Signal vector magnitude (SVM; *mg*) was calculated using the ENMO approach for GENEActiv data. ST was estimated from hip-worn ActiGraph data, applying the widely used 100 count·min⁻¹ threshold. ROC analysis using 10-fold hold-out cross-validation was conducted to establish a wrist-worn GENEActiv threshold comparable to the hip ActiGraph 100 count·min⁻¹ threshold. GENEActiv data were also classified using three empirical wrist thresholds and equivalence testing was completed.

Results: Analysis indicated that a GENEActiv SVM value of 51mg demonstrated fair to moderate
agreement (Kappa: 0.32-0.41) with the 100 count·min⁻¹ threshold. However, the generated and empirical
thresholds for GENEActiv devices were not significantly equivalent to ActiGraph 100 count·min⁻¹.
GENEActiv data classified using the 35.6 mg threshold intended for ActiGraph devices generated
significantly equivalent ST estimates as the ActiGraph 100 count·min⁻¹.

53 Conclusions: The newly generated and empirical GENEActiv wrist thresholds do not provide equivalent 54 estimates of ST to the ActiGraph 100 count·min⁻¹ approach. More investigation is required to assess the 55 validity of applying ActiGraph cutpoints to GENEActiv data. Future studies are needed to examine the 56 backward compatibility of ST data and to produce a robust method of classifying SVM-derived ST.

57 Keywords: children, physical activity, inactivity, accelerometry, measurement

58 Introduction

Sedentary behaviour is increasingly viewed as an important health risk factor in children ¹, and the 59 60 detrimental effects of reallocating PA time to sedentary behaviours have been established². Sedentary 61 behaviour is defined as 'any waking behaviour characterized by an energy expenditure ≤ 1.5 METS while in a sitting, reclining or lying posture' ³ however for children the recommended upper boundary 62 of energy expenditure is ≤ 2 METs or ≤ 1.5 child-METs⁴. It is common for researchers to assess 63 sedentary time (ST) which is commonly defined as the time spent below the threshold of proprietary 64 65 accelerometer counts representing light physical activity, rather than focussing on sedentary behaviour 66 per se.

67 Accelerometers have been used for several years to quantify children's ST, but heterogeneous data processing and researcher decisions related to for example, device location, wear time criteria, and 68 69 choice of thresholds, often mean that study methods lack consistency and comparability. The advent of 70 newer accelerometer devices capable of raw acceleration data collection removes the reliance on 71 proprietary counts and allows researchers more autonomy when examining data, whilst producing estimates of acceleration that in theory should be comparable between devices ⁵. Therefore, devices that 72 73 produce raw acceleration data for researchers to use, such as the GENEActiv and ActiGraph GT3X+ 74 offer an opportunity to increase comparability between studies aiming to estimate ST using 75 accelerometers.

76 Raw acceleration data from GENEActiv and ActiGraph accelerometers are increasingly being processed 77 in the open source R package GGIR (http:/cran.r-project.org). GGIR auto-calibrates the data using local gravity as a reference ⁶, detects sustained abnormally high values and generates the average magnitude 78 of dynamic acceleration (termed the Euclidean Norm Minus One (ENMO))^{5, 7-9}. Recently, the ENMO 79 80 metric has been used to estimate ST and physical activity in both children and adults⁹⁻¹¹, but significant 81 differences have been reported for ST and PA estimated from counts and from raw acceleration signals. Authors have attributed these differences to the various intensity thresholds used to classify acceleration 82 data across the reduction approaches and differences in wear-site¹¹, but they may also be due to the 83 84 inherent differences between the proprietary counts and raw acceleration data. One recent study,

conducted in children, provided a method of calibrating raw acceleration data from wrist-worn monitors 85 86 to counts based hip-warn physical activity estimates in an effort to harmonise data⁹. The study classified 87 raw accelerations using a range of ENMO thresholds for wrist-worn monitors and aligned these to 88 counts-based thresholds for hip-worn monitors, demonstrating that incremental thresholds enable simple group level comparisons to past estimates of physical activity derived from hip-worn accelerometer 89 counts cutpoints. For traditional accelerometer counts-based protocols using hip-worn ActiGraphs, 90 91 studies have widely adopted 100 vertical axis count min⁻¹ as the upper threshold for ST in children ¹². 92 To date, the comparability of wrist-worn GENEActiv ENMO ST estimates to those generated using the ActiGraph 100 vertical axis count min⁻¹ method is unknown. Studies have utilised the ENMO regression 93 equation published by Hildebrand et al.⁸ which was generated using a laboratory protocol to classify 94 ST, however, these thresholds have not been cross-validated for classifying ST or examined in 95 comparison with other methods. More recent studies ¹³ utilised the Hildebrand et al. ⁸ laboratory 96 protocol to general thresholds then examined the agreement between ST and activPAL (which was 97 considered as a criterion reference standard measure) using free-living data. The thresholds 98 99 demonstrated low specificity, overestimating sedentary time in comparison to the activPAL. The equivalence of wrist worn data classified using these approaches to the 100 count min⁻¹ standard is 100 101 unknown. Therefore, researchers wishing to represent raw accelerations through ENMO cannot compare 102 ST to previous counts-based research, and so a pragmatic solution to classifying ST is required.

The aims of this secondary data analysis were to examine the comparability of children's free-living ST derived using the ENMO metric for wrist mounted GENEActiv accelerometer data, with ST estimated using the waist mounted ActiGraph 100 count·min⁻¹ threshold. This aim was addressed by examining, [1] if comparable ST estimates could be attained from wrist-mounted GENEActiv raw acceleration data anchored to the widely adopted 100 count·min⁻¹ uniaxial hip-mounted ActiGraph ST threshold, and [2] the equivalence of ST estimates between the newly generated threshold, those published by Hildebrand et al. ^{8, 13} and the 100 count·min⁻¹ uniaxial hip-mounted ActiGraph ST threshold.

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111 <u>METHODS</u>

This is a secondary data analysis of data generated by a previous study ⁷. After gaining University ethics approval, informed parental consent, and participant assent 108 10-11-year-old children (n=43 boys) were involved in this study. Data collection took place on school sites from January to May 2014. Stature and body mass were assessed to the nearest 0.1cm using a portable stadiometer (Leicester Height Measure, Seca, Birmingham, UK) and nearest 0.1kg (Seca, Birmingham, UK) respectively using standard techniques ¹⁴. Body mass index (BMI), was calculated for each participant.

Sedentary time was assessed using two tri-axial accelerometers, one worn on the non-dominant wrist (GENEActiv; Activinsights, Cambs, UK) and one worn on the right hip (ActiGraph GT3X+; ActiGraph, Pensacola, FL). Both monitors were initialised using the same computer to record at a frequency of 100 Hz, and participants were asked to wear the monitors at all times for 7 consecutive days except when sleeping and engaging in water based activities (e.g., bathing, swimming).

123 ActiGraph monitors were analysed using ActiLife v 6.11.4 software (ActiGraph, Pensacola, FL). 124 Twenty minutes of consecutive zero counts (1 minute spike tolerance) defined non-wear time, and these periods were subtracted from daily wear time ¹⁵. Sedentary time was coded as ≤ 100 count min⁻¹¹². Valid 125 days were defined as \geq 540 min for a weekday ¹⁶ and \geq 480 min for weekend days ¹⁷. For each participant 126 the valid weekday and weekend day with the longest wear time were selected and retained for analysis. 127 For participants with no valid weekend data, the valid weekday only with the longest wear time was 128 included within analysis. After establishing daily wear time, data for the included days were converted 129 to 1-s epoch csv output files for further analysis. 130

GENEActiv data were downloaded using GENEActiv v 2.2 software (Activinsights, Cambs, UK) and saved as binary files. These were then processed in R (http://cran.r-project.org) using the GGIR package (version 1.1-4). To correct for sensor calibration error autocalibration was completed ⁶. GGIR processing produced files in csv format. Each csv file contained the ENMO-derived average magnitude of dynamic acceleration values expressed in average mg ¹⁸. GENEActiv csv files corresponding to the selected ActiGraph weekday and/or weekend days were taken forward to the next stage of analysis. ActiGraph and GENEActiv time stamped data were synched, resulting in one csv file for each participant containing date- and time-stamped ActiGraph and GENEActiv data in 1 s epochs. Non-wear times were removed from each merged file according to the ActiLife wear time details generated for each participant's ActiGraph data. For the ROC analysis each participant's ActiGraph and GENEActiv data were then summed into 1 min epochs to allow data scoring using the ActiGraph vertical axis 100 count·min⁻¹ as the reference value for sedentary time ¹². These data were then stacked into one csv file to create a dataset including all participants (n = 108, 43 boys).

To establish GENEActiv classification criteria anchored to the ActiGraph 100 count min⁻¹ ST threshold, 144 ROC analysis was performed on the whole sample, which represented 126,999 minutes of monitor wear 145 time. Threshold values were cross-validated using 10-fold hold-out groups stratified by sex ¹⁹, whereby 146 147 separate cross-validation analyses were conducted with a randomly selected hold-out group for each iteration (11 participants [6 girls and 5 boys] per analysis cycle)²⁰. Therefore, each ROC analysis was 148 completed with 97 participants with 11 excluded to enable cross-validation. For each hold-out group 149 2x2 contingency tables were used to check classification agreement based on the GENEActiv 150 classifications generated from each cross-validation ROC analysis. Computed sensitivity and specificity, 151 152 Cohen's kappa coefficients, and percentage agreement between classifications were assessed.

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After generating the classification threshold, ST data were scored using 1 minute epochs. Data were 154 classified for each participant using the newly generated GENEActiv threshold, ActiGraph 100 155 count min⁻¹. Additionally GENEActiv ST was scored using the solved regression equation published by 156 Hildebrand et al ⁸, where ST was defined as ≤ 1.5 child-METS ⁴, resulting in a threshold of 22.6 mg. 157 158 GENEActiv ST was also scored using the 56.3 mg GENEActiv and 35.6 mg ActiGraph thresholds from the Hildebrand et al. 2016 study ¹³. The ActiGraph threshold was included as theoretically using the raw 159 data methods should allow the application of the threshold to the GENEActiv device. Pairwise 160 equivalence testing was completed between all combinations of the thresholds. For this study a 95% 161 equivalence test was performed to examine whether the 90% confidence intervals for mean ST for each 162 163 classification method completely fell within the proposed equivalence zone ($\pm 10\%$ of the mean of ST)

defined by the other classification method, representing statistically significant equivalence.
Equivalence testing has been increasingly used in recent PA research where differences testing is not
appropriate ^{11, 21-24}. Difference testing provides information on whether two methods are statistically
different, where in this context it is more useful to know whether two methods are statistically equivalent
at the group level, thus providing similar estimates. Analyses were conducted using IBM SPSS Statistics
v.22 (IBM, Armonk, NY) and Microsoft Excel 2010 (Microsoft, Redmond, WA) and R for Windows
(http://cran.r-project.org).

171

- 172
- 173 <u>Results</u>

Mean anthropometric data, weekend and weekday accelerometer wear times and the number of daysincluded within analysis for boys and girls are displayed in Table 1.

176

177 [TABLE 1 ABOUT HERE]

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The ROC curve for the whole cohort (N = 108) indicated that a GENEActiv threshold of 51 mg179 180 (sensitivity = 81.2%, specificity = 57.4%, AUC 0.760, 95% CI = 0.758, 0.763) provided the most accurate classification of ST. The ROC generated cutpoints, sensitivity and specificity, agreement, and 181 Kappa values for each hold-out analysis for ST can be viewed in supplementary material A. The hold-182 out analysis found that the ST ENMO threshold performed significantly better than random 183 classification, with agreement ranging from 64.7-69.7% and Kappa values ranging from 0.32-0.41 (fair 184 to moderate agreement²⁵). The mean GENEActiv ST cutpoint generated was 51 mg, corresponding with 185 the whole group threshold, therefore 51 mg was used for subsequent equivalence analysis. 186

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Figure 1 displays the results of the equivalence testing using ActiGraph count·min⁻¹ as the reference threshold. Mean time spent in ST for each classification is displayed in supplementary file B. None of the 90% CIs for the newly generated GENEActiv 51mg (630.6-666.7 min), Hildebrand 2014 22.6 mg

(323.2-362 min) or Hildebrand 2016 GENEActiv 56.3mg (673.5-711.1 min) were completely included 191 within the zone of equivalence for the ActiGraph 100 count min⁻¹ (443.2-541.6 min), suggesting no 192 193 statistically significant equivalence between the cut-points compared and the ActiGraph 100 count min-¹, on average. The Hildebrand ActiGraph 2016 35.6mg threshold, applied to GENEActiv data yielded 194 90% CIs (492.9-527.5) that fell within the zone of equivalence, so is considered statistically equivalent 195 196 to the GENEActiv, on average. The newly generated GENEActiv threshold ST estimates were, on 197 average, significantly equivalent to the 2016 Hildebrand GENEActiv threshold, with the 90% CIs for 198 the Hildebrand 2016 GENEActiv threshold of 56.3mg falling within the zone of equivalence for the threshold generated by our study (689.4-695.1 min, zone of equivalence 583.8-713.5 min). No other 199 200 combinations exhibited statistically significant equivalence.

201

202 [FIGURE 1 ABOUT HERE]

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205 Discussion

206 The aims of this secondary data analysis were to examine the comparability of children's free-living 207 sedentary time (ST) derived from raw acceleration thresholds for wrist mounted GENEActiv 208 accelerometer data, with ST estimated using the waist mounted ActiGraph 100 count min⁻¹ threshold. A 209 GENEActiv wrist ST threshold of 51 mg was generated which demonstrated fair to moderate agreement 210 between the cross-validation and whole samples. The fact that the free-living data reflected a typical 211 range of sedentary activities undertaken by children gave it a high degree of ecological validity. Irrespective of this, ST estimated using the 51 mg was not equivalent to the ActiGraph 100 count min^{-1} 212 213 threshold and therefore is not an acceptable value to use to generate ST estimates from GENEActiv 214 wrist accelerations that are compatible with estimates from waist-worn ActiGraphs. However, when applied to the GENEActiv data, the Hildebrand 35.6 mg ActiGraph wrist acceleration threshold 215 produced significantly equivalent estimates of ST as the waist ActiGraph 100 count min⁻¹ suggesting 216 that this threshold could potentially be applied to GENEActiv data to provide comparable estimates of 217 218 ST. Whether this provides an accurate estimate of ST when compared to criterion reference methods

such as activPAL warrants further investigation, however this was not the purpose of the analysisconducted.

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222 Field-based approaches to generating acceptable ST thresholds may be desirable because of their greater ecological validity, and because they may reduce the risk of misclassification associated with 223 laboratory-derived thresholds being used in the field ²⁶. However, our findings suggest that the current 224 225 thresholds used to classify ST using ENMO do not produce comparable estimates to those reported when using the standard 100 count min⁻¹ approach. The challenges of estimating ST from wrist 226 accelerometry are becoming more established ²⁷. Accelerometers are predominantly designed to 227 measure movement rather than postural allocations. Accelerations from hip- and wrist-worn 228 229 accelerometers are highly correlated in children during ST and physical activities of moderate through to vigorous intensities ²⁸. However, correlations are weaker during stationary light intensity physical 230 activity which can involve a combination of sitting and standing activities, as well as transitions 231 between the two ²⁸. Sitting and standing often encompass a combination of sedentary time and time in 232 233 light intensity physical activity, whereby a high degree of hip and wrist acceleration decoupling occurs. 234 For example, an individual may be sitting but gesturing with their hands, or standing and throwing a ball, both of which involve movements that a hip monitor may not detect but that could be detected by 235 236 a wrist mounted device. This lack of consistency between hip and wrist accelerations during some 237 sedentary and light intensity activities provides some explanation of the moderate levels of agreement observed in the cross-validation analyses, and the lack of equivalence with the hip 100 count min⁻¹ 238 239 threshold in particular.

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The accuracy of classifying ST is not explored in this study, we simply looked at the comparability of the GENEActiv thresholds to the standard ActiGraph vertical axis 100 count·min⁻¹ threshold. Whether the standard approach provides a more or less accurate estimate of ST is not examined and warrants further evaluation. To examine the accuracy of ST thresholds within a field-based protocol, a criterion measure, such as an inclinometer is needed. Theoretically this would increase participant burden through the need to wear two devices, increase the cost of undertaking the research and data would still not allow for cross-comparisons between previous counts based studies. An alternative approach, that negates the need for additional devices, is to use accelerometers to examine assumed postural changes relative to arm elevation and wrist orientation (i.e., the Sedentary Sphere approach ²⁷). Recent evidence suggests that the Sedentary Sphere method provides comparable estimates of ST in adults when compared to the activPAL ²⁹, however, this method has not been validated in children, and so further work is required to examine its utility of this method in this population.

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254 The Hildebrand 22.6 mg ST threshold is based on GENEActiv wrist ENMO values, but was generated using VO₂ data rather than ActiGraph counts as in the current study. This may explain why the 255 256 thresholds were not equivalent. In addition, the laboratory protocol used by Hildebrand et al.^{8, 13} only included lying watching TV and sitting using a computer as sedentary activities⁸. Whilst such activities 257 are common among children they do not reflect the wide range of free-living sedentary behaviours that 258 the children involved in this study were likely to have engaged in. Further, the Hildebrand et al. (2014) 259 22.6 mg estimated ST threshold was calculated from a regression equation anchored to energy 260 expenditure. As sedentary behaviours are characterised by posture and low energy expenditure, 261 262 determining sedentary time using energy expenditure alone without posture classification may be a less accurate approach than using criterion measures such as inclinometers or direct observation ²⁷. 263 264 Hildebrand et al.'s 2016 GENEActiv 56.5mg wrist threshold was similar to our 51mg threshold, though 265 the former demonstrated low specificity, overestimating sedentary time in comparison to the activPAL when examining free-living data, which may be due to the limited number of sedentary stations 266 267 included in the original laboratory protocol.

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There are a number of limitations to this study. Our study was conducted in one geographical area of the UK and as such the results may not be representative of other populations. To classify GENEActiv data against the 100 count·min⁻¹ criterion, we used a 1-minute epoch setting. Though this would likely result in the inability to detect movement at higher intensities, as sedentary behaviour is characterised by a lack of movement the 1-minute epoch setting would have less impact upon the ST estimates generated. We did not use a criterion reference standard device such as activPAL within this study. This was by design, as the primary aim was to examine the comparability of simple accelerometer estimates rather than investigate the accuracy of the measurement of ST. Future studies should aim to utilise the activPAL and other reference methods to develop and validate ST thresholds for use in children.

278

279 <u>Conclusions</u>

Despite displaying fair to moderate agreement, the generated GENEActiv ST threshold does not provide an equivalent estimate of ST to the hip mounted ActiGraph 100 count·min⁻¹ approach. Furthermore, ST data generated using Hildebrand thresholds were not equivalent to the 100 count·min⁻¹ method. Future studies are needed to examine the backwards compatibility of ST data and to produce a robust method of classifying ENMO-derived ST.

285

286 **Practical implications**

- Estimates of children's sedentary time generated from GENEActiv wrist ENMO and ActiGraph
 100 count·min⁻¹ are not comparable.
- Researchers should not compare data generated using the two different methods.
- Future studies are required to provide methods of data harmonization and to establish valid and
 reliable sedentary time thresholds for children.

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366 <u>Table legend</u>

367 <u>Table 1. Mean (SD) anthropometric, wear time and number of days included within analysis for boys</u>

- 368 <u>and girls</u>
- 369

	Boys $N = 43$		Girls N = 65	
	Mean or	SD	Mean or	SD
	Frequency		Frequency	
Age (y)	10.03	0.35	10.04	0.31
Height (cm)	139.49	7.89	137.97	7.37
Body mass (kg)	35.64	8.24	34.23	8.60
BMI (kg·m ²)	18.15	3.00	17.78	3.18
ActiGraph weekday wear (min·day ⁻¹)	739.88	115.55	738.75	100.35
ActiGraph weekend day wear (min·day-1)	631.83	110.82	661.50	108.28
ActiGraph valid weekdays included	41	N/A	64	N/A
ActiGraph valid weekend days included	30	N/A	46	N/A
Total valid included days	71	N/A	110	N/A

370

372 Figure legend

Figure 1. ActiGraph 100 count·min⁻¹ zone of equivalence (dotted lines) and 90% confidence intervals for the GENEActiv sedentary time data

