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Branched equation modeling of simultaneous accelerometry and

heart rate monitoring improves estimate of directly measured

physical activity energy expenditure

Søren Brage^{1,2*}, Niels Brage^{1*}, Paul W. Franks², Ulf Ekelund^{2,3}, Man-Yu Wong^{2,4}, Lars Bo

Andersen⁵, Karsten Froberg¹, & Nicholas J. Wareham²

*) SB and NB contributed equally to this work

1) Institute of Sport Science & Clinical Biomechanics, University of Southern Denmark, Main

Campus: Odense University, Odense, Denmark

²⁾ Institute of Public Health, University of Cambridge, UK

³⁾ Department of Physical Education & Health, Örebro University, Örebro, Sweden

⁴⁾ Department of Mathematics, Hong Kong University of Science & Technology, Hong Kong

⁵⁾ Institute of Sport Science, University of Copenhagen, Copenhagen, Denmark

Running head: Simultaneous accelerometry and HR to estimate PAEE

Corresponding author and requests for reprints to:

Søren Brage

Institute of Public Health, University of Cambridge, University Forvie Site, Robinson Way,

Cambridge CB2 2SR, UK

Tel.: +44 1223 330316

Fax: +44 1223 330330

E-mail: sb400@medschl.cam.ac.uk

ABSTRACT

2	The combination of heart rate (HR) monitoring and movement registration may improve
3	measurement precision of physical activity energy expenditure (PAEE). Previous attempts
4	have used either regression methods, which do not take full advantage of synchronized data, or
5	have not used movement data quantitatively. The objective of the study was to assess the
6	precision of branched model estimates of PAEE, utilizing either individual calibration (IC) of
7	HR and accelerometry or corresponding mean group calibration (GC) equations. In 12 males
8	(20.6-25.2 kg·m ⁻²), IC and GC equations for physical activity intensity (PAI) were derived
9	during treadmill walking and running for both HR (Polar) and hip-acceleration (CSA). HR and
10	CSA were recorded minute-by-minute during 22hrs of whole-body calorimetry and converted
11	into PAI in four different weightings (P_{1-4}) of the HR vs. the CSA ($1-P_{1-4}$) relationships: If
12	CSA>X, we used the P_1 weighting if HR>Y, otherwise P_2 . Similarly, if CSA \leq X, we used P_3 if
13	HR>Z, otherwise P ₄ . PAEE was calculated for a 12.5hr non-sleeping period as the time-
14	integral of PAI. A priori, we assumed P ₁ =1, P ₂ =P ₃ =0.5, P ₄ =0, X=5counts·min ⁻¹ ,
15	Y=walking/running transition HR, and Z=flex HR. These parameters were also estimated post
16	hoc. Mean±SD estimation errors of a priori models were -4.4±29% and 3.5±20% for IC and
17	GC, respectively. Corresponding <i>post hoc</i> model errors were -1.5±13% and 0.1±9.8%. All
18	branched models had lower errors (p \leq 0.035) than single-measure estimates of CSA (\leq -45%)
19	and HR (\geq +39%), as well as their non-branched combination (\geq +25.7%). In conclusion,
20	combining HR and CSA by branched modeling improves estimates of PAEE. Individual
21	calibration may be less crucial with this modeling technique.

KEY WORDS: Validity, intensity, epidemiology, calorimetry, movement sensor, activity
monitor, energy expenditure, individual calibration
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INTRODUCTION

- 2 Physical activity is a complex behavior and difficult to measure precisely at population level.
- 3 The reasons that precise estimates of physical activity are important include clarification of
- 4 which dimension of activity is most strongly associated with a particular health outcome,
- 5 understanding dose-response relationships, improving the ability to monitor secular trends in
- 6 activity level and the compliance to intervention programs, cross-cultural comparisons, and
- 7 optimization of sample size for the detection of gene-environment interactions (21; 38).
- 8 Accelerometry and heart rate (HR) monitoring are among the available objective methods.
- 9 There are, however, limitations associated with both methods when used alone for the
- assessment of physical activity energy expenditure (PAEE) and its first time derivative,
- physical activity intensity (PAI). The limitations of HR monitoring are largely due to
- biological variance. For example, the HR-PAI relationship is affected by age, sex, training
- state, stroke volume, hemoglobin concentration of the blood and its O₂ saturation, mental
- stress, ambient temperature, hydration, and quantity of muscle mass involved in the activity
- 15 (20; 36; 41). Some of these limitations can be overcome by individual calibration. In contrast,
- the limitations of accelerometry are primarily biomechanical, in that the accelerometry-PAI
- 17 relationship during different activities, such as walking on the level and incline, when running,
- stepping, and cycling, and during load-bearing activities, is highly variable (2; 5; 6; 9; 10; 15;
- 19 18; 24; 31; 33). Thus, it is difficult to accurately translate epidemiological accelerometry data
- directly into units of intensity or expended energy. Since the errors associated with the two
- 21 methods are not positively correlated, the combination of HR and accelerometry should
- theoretically yield a more precise estimate of PAEE and PAI than either used independently (5;
- 23 6; 11; 14; 22; 25; 28; 34; 35; 37). Many of the studies, in which the validity of combined heart
- rate monitoring and accelerometry has been assessed, have used multiple regression
- 25 techniques, which produce weighted averages. These depend upon the protocol used and

1 consequently may not perform well in other diverse scenarios. Moreover, this approach does 2 not exploit the time synchronization between the accelerometry and heart rate to estimate the 3 model coefficients. The selective use of data derived from either heart rate or accelerometry, 4 depending on the characteristics of the activity assessed, may improve the generalizability of 5 PAEE and PAI estimates to the free-living scenario. Some investigators have adopted this 6 approach (28; 34; 35), although they have not explored the potential use of body movement 7 data in a quantitative manner when combining the two data sources. Combining the methods in 8 this way may provide more robust estimates of intensity and energy expenditure in the 9 intensity region around the flexHR, a HR used to discriminate between activity and non-10 activity. Definition of flexHR is critical, as the majority of time is spend in this intensity region 11 (28). Although Rennie et al demonstrated the utility of a single piece combined monitor (28), 12 this device is not commercially available. Other studies involved equipping subjects with a 13 Polar heart rate monitor and two Computer Science & Applications (CSA) accelerometers 14 model 7164 (now also known as MTI Actigraphs; Manufacturing Technology Inc., Fort 15 Walton Beach, FL, USA), one on the arm and one on the leg (34; 35). This combination 16 performed better for the prediction of energy expenditure than when HR monitors, hip mounted 17 CSAs, or hip mounted pedometers were used separately. However, as with the study by Rennie 18 et al, this combined method did not use the accelerometer data quantitatively and required even 19 more individual calibration than is commonly undertaken in the epidemiological setting. 20 Additional potential problems with using three separate measurement units in epidemiological 21 studies include lower response rates and increased Hawthorne effect. Although the 22 combination of a Polar HR monitor and a hip-mounted CSA accelerometer is not perfect, it is 23 more feasible, and is thus currently being used in larger cohorts. Additionally, the precision of 24 any objective assessment method, especially energy expenditure estimated from HR, is that it 25 is dependent upon some level of individual calibration. However, this procedure places

1 additional demands on both experimenter and participant. Thus, a key question is whether the 2 combination of HR and accelerometry will be sufficiently precise to preclude the need for 3 individual calibration. Therefore, the aim of this study was to compare the time integral of minute-by-minute estimated PAI (in kJ·kg⁻¹·min⁻¹) from the combination of a hip mounted 4 5 CSA accelerometer and a Polar HR monitor against whole-body calorimetry PAEE (in kJ·kg⁻¹). This was done by using both accelerometry and HR data in a quantitative manner, and with and 6 7 without individual calibration. 8 9 RESEARCH DESIGN AND METHODS 10 **Subjects** 11 Twelve male subjects (22.7-30.0 years, 63.9-91.2 kg, 169-199 cm, 20.6-25.2 kg·m⁻²) performed 12 individual calibration on a treadmill, following which they spent 22 hrs in a whole-body heat-13 sink calorimeter, in which they performed various activities of daily living. All subjects were healthy and well trained with a mean peak VO₂ (fitness) of 61.5 ml·min⁻¹·kg⁻¹ (range: 51.0-71.5 14 ml·min⁻¹·kg⁻¹). Informed written consent was obtained from each participant on entry to the 15 16 study, which was approved by the local research ethics committee (Denmark). 17 18 **Calorimetry study** 19 Calorimeter: The calorimeter protocol has been described in detail previously (17). In brief, 20 the method relies on the principle that all expended energy is converted to heat (sensible 21 power) or used to evaporate water (evaporative power). An example of the calorimeter output 22 is shown in **Figure 1**. The sum of the two time integrals of the sensible and the evaporative 23 power represents the overall energy production for the time interval. Due to the large volume 24 of this heat-sink calorimeter, the response time of the two power readings, e.g., after an activity 25 bout, is rather slow and thus data must be analyzed in extended epochs. Nonetheless, the

- 1 energy expenditure measurements from these epochs are precise within $\pm 2\%$, owing to high
- 2 precision of the sensible and evaporative power (+1.4% and +4.0%, respectively) (17).
- 3 The total energy expenditure (TEE) is comprised of three main components. These are resting
- 4 energy expenditure (REE), diet-induced thermogenesis (DIT), and physical activity energy
- 5 expenditure (PAEE). In order to assess the predictive capabilities for PAEE of CSA and HR
- 6 used independently and in combination, PAEE was calculated as TEE (REE+DIT) and
- 7 expressed in kilojoules (kJ) per kg body weight. The data were analyzed in two epochs: a
- 8 night-time period from 0:00-07:00hrs and a day-time period from 07:30-20:00hrs.
- 9 HR and movement measurement: During both sleeping and waking hours, subjects wore a
- 10 Polar Vantage NV heart rate monitor (Polar Electro, Kempele, Finland), set to measure HR (in
- beats per minute, bpm) every 15 seconds and two CSA accelerometers on each hip, sampling at
- 12 15 and 60 second epochs, respectively. All CSA and HR data were compiled to a minute-by-
- minute file for each individual. We used the mean of four CSA monitors to make our estimates
- more generalizable, as differences between CSA monitors and sites of placement on the hip
- have been reported (4; 39). We defined resting HR (RHR) as the tenth lowest HR observed
- during sleeping to obtain a robust estimate of this key parameter.
- 17 Study protocol: Each subject was instructed not to exercise during the two hours immediately
- preceding their arrival at the laboratory at 21:30hrs and to refrain eating and drinking (other
- than water) from 20:30hrs onwards. Height and body weight were assessed by standard
- anthropometric methods. Body composition was assessed by the impedance technique (TBF-
- 21 300, Tanita Europe GmbH, Germany), using an average between the 'Standard' and the
- 22 'Athlete' settings on the impedance scale. Subjects entered the calorimeter at 22:00hrs and left
- 23 the calorimeter 22hrs later. **Figure 1** shows the overall activity protocol for the 22hrs of
- calorimetry. Each subject performed a standardized protocol, which aimed to emulate the types
- of activity the subjects would undertake during a typical day. This involved periods of

- 1 rest/reading and bouts of different forms of exercise. During periods when no activity other
- 2 than reading was scheduled, subjects were allowed to use the telephone in the calorimeter. Of
- 3 the 12.5hrs when subjects were awake, they spent 13.1% on regular physical activity, which
- 4 comprised of 4.7% cycling, 4% walking, 2.4% stepping, and 2% jogging. The subjects went to
- 5 bed at 23:00hrs and were woken at 07:30hrs. Breakfast was served at 08:15hrs, lunch at
- 6 13:30hrs, and snacks (fruit or chocolate) at 10:15hrs, 15:45hrs, and 17:30hrs. Ad libitum
- 7 quantities and compositions of breakfast and lunch were selected by the subject from a limited
- 8 menu.
- 9 Diet-induced thermogenesis: All consumed foods were registered and analyzed by a national
- 10 food database (DanKost 2000, Dansk Catering Service A/S, DK) to yield energy intake (EI)
- and macronutrient composition. DIT was estimated from the absolute energy yield (in kJ) of
- the three macronutrients, according to the equation DIT = $0.025 \cdot \text{fat EI} + 0.07 \cdot \text{carbohydrate EI}$
- 13 + 0.275 · protein EI (19).
- 14 Resting energy expenditure: A heat source yielding exactly 6 kJ·min⁻¹ was introduced from
- 15 02:00-05:00hrs, as a means of internal validation. This procedure enables the examination of
- the precision of the calorimeter's response to an increase in energy expenditure. The sleeping
- metabolic rate (SMR) was calculated for the periods 0:00-02:00hrs (SMR₁) and 05:00-07:00hrs
- (SMR_3) and then averaged (SMR_{1+3}) . This was compared to the calculated SMR from 02:00-
- 19 05:00hrs (SMR₂), which ideally should be 6 kJ·min⁻¹ higher if the calorimeter was 100 %
- 20 accurate, assuming that SMR₁₊₃ did indeed approximate SMR during the heat supplementation.
- 21 The total heat supplement was 1080 kJ, so the SMR for the whole night was calculated as SMR
- = $(TEE_{0:00-07:00hrs} 1080kJ) / 7hr$. The resting metabolic rate (RMR) was assumed to equal 105
- 23 % SMR (13). The RMR value was used as a baseline in derivation of the calibration equations
- 24 (see below). Resting energy expenditure (REE) during time awake was obtained by integrating

- 1 RMR over 12.5 hrs. This was also obtained by prediction equations using the impedance-
- 2 derived body composition data (16).

- 4 Calibration study 5 The calibration procedure was carried out in duplicate on a treadmill approximately four 6 months before the calorimetry study, as described previously (6). The subjects did not change 7 their overall (self reported) physical activity level during this interim period. Briefly, the 8 calibration protocol consisted of 5min intervals (continuous) at the following treadmill velocities: 3 and 6 km·h⁻¹ of walking and 8, 9, 10, 12, 14, 16, 18, and 20 km·h⁻¹ of running until 9 10 volitional exhaustion. On both these treadmill tests, oxygen consumption was measured by an 11 automated system (EO Sprint, Erich Jaeger GmbH, Germany). Aerobic fitness (peak VO₂) was 12 determined as the maximal observed value in either of the two treadmill tests. For each 13 velocity, steady state VO₂ and HR were calculated as the mean of minutes 3.5–5 following 14 change of speed and CSA output was expressed as the mean of 4min, i.e., four epochs not 15 overlapping different speeds. Body mass specific PAI was calculated as VO₂ minus measured RMR from the calorimeter and expressed in kJ·kg⁻¹·min⁻¹ by assuming an energetic value of 1 16 17 L oxygen ~ 20.35 kJ (7). This value assumes that energy is derived equally from fat and 18 carbohydrate and has been used elsewhere (28). Parallel estimates of PAI were also obtained 19 using predicted RMR. All HR values are expressed as absolute values minus resting heart rate 20 (**R**R). 21 The calibration equations for PAI (calculated both using measured and predicted RMR) were 22 derived at group level (N=12) and individual level for CSA, HR, and their combination as 23 follows:
- 24 CSA to PAI conversion: One-dimensional accelerometers, such as the CSA record virtually the
- 25 same value across running speeds but increases linearly across walking through to jogging

- speeds (6; 12; 26). Therefore, we used linear regression to produce prediction equations for the
- 2 CSA-PAI relationship in the 3-9 km·h⁻¹ range of CSA output (two walking and two running
- 3 speeds). This relationship was extrapolated to a CSA flex point, defined as 50% of the mean
- 4 CSA output at 3 km·h⁻¹. Between this flex point and the origin (0 counts·min⁻¹, 0 kJ·kg⁻¹·min⁻¹),
- 5 we assumed a straight line.
- 6 HR to PAI conversion: All subjects completed the calibration protocol including the 16 km·h⁻¹
- 7 interval. Therefore, prediction equations for the HR-PAI relationship were produced using only
- 8 data in the 3-16 km·h⁻¹ intervals by quadratic regression (8). This regression was forced
- 9 through the origin (0 bpm, 0 kJ·kg⁻¹·min⁻¹), thus effectively assuming that the energy
- expenditure is equal to REE when the absolute HR is equal to RHR. In the flex HR method,
- 11 this relationship was used for all HR values above the flex HR (defined as the 10 bpm + the
- 12 average of RHR and the mean HR at 3 km·h⁻¹). For HR values below the flex HR, we assumed
- 13 PAI to be 0 kJ·kg⁻¹·min⁻¹. This approach is similar to the one used by Spurr and colleagues
- 14 (32).
- 15 Non-branched CSA+HR to PAI conversion (MLR): Multiple linear regression on the treadmill
- data in the 3-9 km·h⁻¹ range was used to produce a non-branched equation, containing both
- 17 CSA and HR.
- 18 Branched CSA+HR to PAI conversion (a priori): We constructed a branched model, the
- structure of which is shown in **Figure 2**. A priori, we assumed values of Y = the
- walking/running transition HR (mean HR between the fastest walking and the slowest running
- on the treadmill), Z = flex HR, $P_1 = 100\%$, $P_2 = P_3 = 50\%$, and $P_4 = 0\%$. Pilot testing indicated that 5
- 22 counts·min⁻¹ is moderately exceeded in cycling activity, so to ensure that cycling was not
- quantified by Box 4 in **Figure 2**, we set X = 5 counts·min⁻¹.

- 1 The day-time period in the calorimeter lasted 750min, and for each minute we converted the
- 2 CSA and HR into PAI (kJ·kg⁻¹·min⁻¹), using the derived calibration equations. Estimates of
- 3 PAEE were obtained as the sum (time integral) of the 750 estimated values of PAI. This
- 4 produced estimates of PAEE (in kJ·kg⁻¹) for each individual, which were then compared with
- 5 the measured value. A priori, a total of eight models using measured RMR in the calibration
- 6 and eight models using predicted RMR in the calibration were tested against measured PAEE.

8

Post hoc branched CSA+HR model estimation

- According to the branched model (**Figure 2**), we also estimated the parameters X, Y_{1-2}, Z_{1-2} ,
- and P₁₋₄, using both the individually calibrated conversion equations and the group mean
- calibration equations. This estimation was done by minimizing the standard error of the
- estimates (SEE), calculated as the square root of the mean squared error between the estimated
- and the measured PAEE for all potential models. This is essentially normal linear regression,
- i.e., all possible combinations of the parameters X, Y_{1-2} , Z_{1-2} , and P_{1-4} , are considered and their
- 15 time integral (PAEE) of the resulting minute-by minute PAI compared to the measured PAEE.
- Indeed, this procedure would be preferred if the criterion measure was minute-by-minute PAI,
- but since this is not the case, it is necessary to restrict the flexibility of the model to only move
- within reasonable boundaries around the parameters set in the CSA+HR (a priori) model.
- 19 Thus, constraints of parameters were specified as follows: X range 0 60 counts·min⁻¹, Y
- range 24 105 bpm, Z range 5.5 34 bpm (both obtained with the Y_1 and Z_1 range ± 5.0 , and
- 21 the Y₂ and Z₂ range ± 250 bpm), and P₁ \geq P₂ \geq P₃ \geq P₄ (all range 0 100%). The Y and Z ranges
- were determined by a 50 % expansion of the ranges of transition HR and flex HR, respectively.
- 23 To assess the robustness of the estimated model parameters, the SEE minimization procedure
- 24 was re-run disregarding the maximum and the minimum error, resulting in a different set of
- 25 parameters with a 'trimmed' SEE. The relative contribution from HR and CSA was calculated

1 as the fraction of observations being quantified by boxes 1-4 in Figure 2 times their weighting 2 $(P_{1-4} \text{ and } 1-P_{1-4} \text{ for HR and CSA, respectively}).$ 3 4 **Statistics** 5 Mean CSA output and mean HR during day-time were calculated and denoted CSA_{day} and 6 HR_{day}, respectively. CSA_{night} and HR_{night} were calculated in a similar manner. For comparison 7 purposes, the associations between PAEE and either CSA_{day}, HR_{day}, or their combination were 8 also modeled by linear regression. 9 Agreement between the estimates of PAEE and the measured PAEE values was assessed in 10 multiple ways. Firstly, differences were calculated and tested with Student's paired t-tests. 11 Differences are expressed as percentages of the measured PAEE and TEE values. Secondly, 12 heteroscedasticity was explored by inspection of modified Bland-Altma plots (errors plotted 13 against measured PAEE) and quantified with Pearson correlation (1; 3). Thirdly, precision of 14 the models was assessed by the standard error of the estimates (SEE). Difference in model 15 precision was tested with Student's paired t-tests on the squared estimation errors. To test for 16 bias and confounding, Pearson correlation was used to test whether estimation errors could be 17 explained by weight changes, aerobic fitness, body composition, RMR, or energy intake. The 18 agreement between SMR₁₊₃ and SMR₂ was tested with Student's paired t-test. Statistical 19 significance was set at the 0.05 level. All analyses were performed with STATA version 7.0 20 (Stata Corp. TX, USA), except the parameter estimation for the post hoc models, which using a 21 macro (available on request), programmed in JAVA version 1.4.1 (Sun Microsystems Inc, 22 USA). 23 24 **RESULTS**

25 Internal validation of the calorimeter

- 1 During the night, SMR decreased by 14.4% (p<0.001) when comparing SMR₁ and SMR₃.
- 2 Mean (SD) difference between SMR₁₊₃ and SMR₂ was 6.2 (0.6) kJ·min⁻¹, which was not
- 3 significantly different (p=0.31) from the 6 kJ·min⁻¹ supplied by the heat source.

- 5 *Calibration equations*
- 6 CSA flex was 497 counts·min⁻¹ at group level (range: 432-563 counts·min⁻¹). The group
- 7 calibration equations for the prediction of PAI were PAI = $0.053 \cdot \text{CSA} + 47.88$ for CSA values
- 8 above CSA flex and PAI = 0.15·CSA for CSA values below. The PAI-HR relationship was:
- 9 PAI = $0.011 \cdot HR^2 + 5.82 \cdot HR$, and the non-branched CSA+HR model was PAI = $0.028 \cdot CSA +$
- 10 4.04·HR 38.3 with all HR values expressed as bpm above RHR. There was no difference
- between the two treadmill tests for any of these relationships (p>.80).

- 13 Summary of collected variables during awake
- Mean (SD) percentage body fat was 9.8 (2.1) %. Mean (SD) values of REE, PAEE, and EI
- were 52.4 (7.4) $kJ \cdot kg^{-1}$, 33.6 (7.0) $kJ \cdot kg^{-1}$, and 153.5 (17.8) $kJ \cdot kg^{-1}$, respectively for the 12.5hrs
- when subjects were awake. Mean predicted REE was not significantly different from mean
- measured REE (p=.538) but there was a negative trend (r=-.73, p=.007) in the Bland-Altman
- plot (not shown). Energy expenditure from physical activity accounted for an average (SD) of
- 19 33 (5.8) % of TEE. DIT accounted for 12 (1.3) %, and REE for the remaining 55 (5.2) %.
- 20 Mean (SD) RHR was 43.6 (6.3) bpm. Average day-time HR ranged from 14.9 to 27.9 bpm
- above RHR, with a mean (SD) of 21.5 (3.8) bpm above RHR. All observed CSA values were
- in the range 0 14,636 counts·min⁻¹, with a third of the daytime observations being 0 or 1
- 23 count·min⁻¹ and 1% above 7,500 counts·min⁻¹. Average daytime CSA output had a group mean
- 24 (SD) of 290.2 (64.3) counts·min⁻¹, whereas means for each subject ranged from 173.6 to 397.3
- 25 counts·min⁻¹.

- 2 PAEE estimated from CSA, flexHR, CSA+HR (MLR), and CSA+HR (a priori)
- 3 Measured PAEE was significantly correlated with CSA_{day} (R²=0.55, p=0.006, SEE=4.96 kJ·kg⁻¹
- 4 ¹), HR_{day} (R^2 =0.35, p=0.044, SEE=5.95 $kJ\cdot kg^{-1}$), and their combination (R^2 =0.78, p=0.001,
- 5 SEE= $3.67 \text{ kJ} \cdot \text{kg}^{-1}$).
- 6 Using the treadmill calibration on both individual and group levels produced the estimates of
- 7 PAEE from CSA, flex HR, and their combinations that are shown in **Table 1**. The mean (SD, p
- 8 for difference from measured value) percentage errors of the CSA estimates of PAEE were -
- 9 50.8 % (10.0 %, p<0.001) and -45.1 % (7.3 %, p<0.001) for the individually calibrated and the
- group calibrated estimates, respectively. Corresponding values were 39.1 % (58.0 %, p=0.047)
- and 48.8 % (37.7 %, p=0.001) for flexHR, 29.9 % (71.8 %, p=0.176) and 25.7 % (25.6 %,
- 12 p=0.004) for the non-branched CSA+HR model, and -4.4 % (29.0 %, p=0.612) and 3.5 % (20.1
- 13 %, p=0.477) for the branched CSA+HR (*a priori*) model.
- 14 The modified Bland-Altman plots (**Figure 3.A+B**) illustrated that differences between CSA
- estimates and measured values of PAEE were negatively correlated with PAEE (r=-0.88,
- p<0.001 for both individual and group calibration estimates). Estimation errors from flex HR
- demonstrated a different relationship (**Figure 3.C+D**) with r=0.15 (p=0.631) and r=0.53
- 18 (p=0.079) for individual and group calibration estimates, respectively. For the two non-
- branched CSA+HR (*MLR*) models (**Figure 3.E+F**), corresponding values were r=0.25
- 20 (p=0.426) and r=0.28 (p=0.387), and for the two branched CSA+HR (a priori) models (**Figure**
- 21 **3.G+H** values were r= -0.08 (p=0.803) and r=0.19 (p=0.561). For all eighta *priori* models,
- 22 estimation errors were not significantly related to weight change between the calibration and
- 23 the calorimeter test ($p \ge 0.40$), weight change during the calorimeter test ($p \ge 0.07$), fitness
- 24 (p \ge 0.17), body composition (p \ge 0.10), RMR (p \ge 0.10), or energy intake (p \ge 0.21).

1 As indicated by the differences in SEE (**Table 1**), the branched CSA+HR (*a priori*) models 2 were more precise than both the corresponding single-measure models and the non-branched 3 CSA+HR models (p=.035 and p=.007 for the models using individual and group calibration, 4 respectively). Of all four single-measure models, only the flex HR model using group 5 calibration was significantly less precise than the non-branched CSA+HR model at the same 6 level of calibration (p=.048). Only the non-branched model of CSA+HR using group 7 calibration lost a statistically significant amount of precision when utilizing predicted RMR 8 instead of measured RMR in the calibration. Other models were either unaffected or showed 9 improvement. 10 11 PAEE estimated from branched CSA+HR (post hoc) 12 The PAEE estimates of the two branched post hoc models are shown in **Table 1**, together with 13 the *a priori* model estimates. The branched model parameters underlying these results are 14 displayed in **Table 2**. The mean (SD, p for difference from measured PAEE) percentage errors 15 of the estimates were -1.5 % (13.0 %, p=0.452) and 0.1 % (9.8 %, p=0.843) for the individually 16 calibrated and the group calibrated estimates, respectively. Mean error in percent was -2.36% 17 for the individual calibration model (individual estimates from -24 to 16%) and +0.54% 18 (individual estimates ranging within +14%) for the group calibration model, corresponding to 19 about 0.18+4.6% of TEE. The branched post hoc models were also more precise than their 20 non-branched counterparts (p=.026 and p=.007 for the models using individual and group 21 calibration, respectively). The Bland-Altman plots (Figure 3.I+J) illustrated that estimation 22 errors were not significantly correlated with measured PAEE (r=-0.40, p=0.202 and r=-0.36, 23 p=0.244 for individual and group calibration, respectively). Estimation errors of the two post

hoc models were not significantly related to weight change between the calibration and the

- calorimeter test ($p \ge 0.51$), weight change during the calorimeter test ($p \ge 0.08$), fitness ($p \ge 0.16$),
- body composition (p \geq 0.24), RMR (p \geq 0.27), or energy intake (p \geq 0.24).

25

4 **DISCUSSION** 5 The calorimeter used in this study has been previously shown to measure energy expenditure 6 with a precision of +2% (17). This level of precision is similar to that estimated through our 7 internal validation using the fixed heat source. For the calculation of the criterion measure, 8 PAEE, it was necessary to make assumptions on the magnitude to which diet increases TEE. 9 There is some controversy about the magnitude of DIT for a given individual, but it is 10 generally agreed that DIT is modified by age, gender, obesity, and the macro-nutrient intake 11 (19; 23; 27; 29; 40). With the exception of macro-nutrient intake, our sample was relatively 12 homogeneous, with regard to these modifying parameters, which helps justify our choice of 13 method to calculate DIT in the present study. Furthermore, DIT estimated in this way was 14 comparable to the more simplistically calculated and widely used value of 0.1 TEE (30). 15 Although the results of the present study change with different assumptions of DIT, the relative 16 differences between the models persist, therefore demonstrating the utility of combining 17 accelerometry with HR. 18 The CSA accelerometer and the Polar HR monitor are among the most widely used physical 19 activity monitors. Although, this combination has been studied previously, the specific 20 modeling techniques described in the present study have not been reported (5; 6; 34; 35). The 21 combination method used here is based on three assumptions. The first is that resting HR is 22 measured, preferably overnight. The second is that some level of calibration has been 23 undertaken. And the third is that a valid measure of REE is obtained to provide a base for the 24 PAI calculations. Our measurement of REE was obtained by the gold standard technique of

whole-body calorimetry, including overnight assessment. This procedure is infeasible for large-

1 scale population studies, but was appropriate in this study for the purpose of validating the 2 method we describe. An alternative to our procedure would have been measurement of REE 3 with indirect calorimetry as a part of the calibration, or by calculating REE from prediction 4 equations. Although the latter showed regression to the mean in our study, this did not 5 significantly affect the precision of the branched models. Additionally, we used four CSA 6 accelerometers in this study to correct for unit differences (4) and effect of placement on the 7 hip (39) but this not necessary in the epidemiological setting if CSA units are thoroughly 8 calibrated before use and consistently placed on the same site. 9 Our main finding was that the combination of HR and accelerometry improves the estimates of 10 PAEE when using treadmill derived calibration equations in a branched model. This is mainly 11 because estimates of PAEE derived using the CSA are usually underestimates of the true value, 12 whereas the flex HR method usually overestimates PAEE. The group level calibration 13 equations used individually in this study are not independent of the subjects, as each of them is 14 represented by a weight of 0.083 in the equations. Therefore, the comparisons between the 15 estimates with and without individual calibration should be interpreted with some caution and 16 merely taken as an indicator of how the error structure and optimal weighting between 17 accelerometry and HR data changes as one moves away from individual calibration. 18 Nonetheless, moving towards a higher degree of individual calibration resulted in a slightly 19 more precise group mean and less heteroscedasticity (correlation between estimation error and 20 PAEE) only for the flexHR method, although the SEE tended to increase (non-significant). For 21 all other models, the group mean estimate and the SEE tended to decrease (only significant for 22 the CSA and CSA+HR (MLR models). Although HR is expressed in beats above resting, 23 which reduces a considerable amount of the inter-individual variance in the HR-PAI 24 relationships, this was not anticipated. Even though genetic and other non-variable components 25 of the inter-individual variance in the PAI relationships would be removed by individual

- 1 calibration, it is possible that drift between the calibration and validation parts of the study, due
- 2 to changes in fitness and/or weight, may explain this observation. However, even though small
- 3 weight changes occurred, this is largely accounted for by expressing all values relative to body
- 4 weight. Furthermore, the estimation errors were not significantly related to weight changes,
- 5 fitness (at the time of calibration), body composition, RMR, or EI, making (residual) bias less
- 6 likely. The greater variance of the PAEE estimation errors for the models incorporating
- 7 individual calibration suggests that errors in the calibration procedure are greater than errors
- 8 resulting from inter-individual variance, with the possible exception of the HR-PAI
- 9 relationships. If this is true, it highlights the importance of choosing an appropriate calibration
- procedure. This is probably more of an issue for the interpolations that were employed to infer
- energy expenditure at the lowest levels of activity. In this study, the interpolated part of the
- 12 CSA-PAI relationship is used more than 90 % of the time, reflecting the relatively long periods
- of low or no activity in the calorimeter. Indeed, a large proportion of time was spent in the
- intensity region around the flex HR, as was also observed by Rennie and colleagues (28).
- Ultimately, any calibration procedure should reflect the activity most commonly engaged in by
- the population in which it is being employed. But this is often hard to define in the free-living
- scenario, especially when one is mindful of minimizing the burden placed upon the
- experimenter and participants. Although in the interests of precision, a representative
- calibration procedure should perhaps involve 24hr whole body calorimetry for each individual
- 20 (25; 37), this is unfeasible in large-scale epidemiological studies, in which access to the
- 21 participant and to the laboratory is often limited.
- 22 The precision of the CSA+HR (a priori) model using individual calibration is comparable to
- 23 that level reported by Rennie *et al*, when expressed in the same way (i.e., relative to TEE) (28).
- The levels of energy expenditure were also similar in our study and theirs (7.6 kJ·kg⁻¹·hr⁻¹ vs.
- 8.0 kJ·kg⁻¹·hr⁻¹, p=0.25), although the protocols differed. However, as already highlighted, the

1 CSA+HR (a priori) model using the group calibration, which we report in this paper, was more 2 precise than the method reported by Rennie, although this may be attributable to our more 3 homogeneous sample. Irrespective of this, the high precision of the CSA+HR (post hoc) 4 models is encouraging; in theory, the model using group calibration could predict PAEE within 5 0.54% on group level, with the individual estimates within +14%, corresponding to about 6 0.18±4.6% of TEE. Interestingly, the partial contributions of HR and CSA were virtually 7 reversed in the branched models, as compared to multiple linear regression derived equations 8 on walking and running alone (5; 6; 22). Although, this is partially due to the absence of high 9 intensity exercise in the protocol, it supports the potential utility of the branched modeling 10 technique, particularly in populations where high intensity exercise is uncommon. However, 11 there was some variation in the estimated parameters as compared to their 'trimmed' 12 counterparts. This was especially the case for X and Y in the model using group calibration, 13 suggesting that these parameters may lack robustness. In contrast, the Z and P parameters were 14 comparable between the individual- and group-calibrated models. Nonetheless, these models 15 should only be used in other populations bearing in mind that the data for these models is 16 derived from a relatively small and homogeneous sample of young men, who undertook a fixed 17 activity protocol in a calorimeter. For example, these branched models are likely to 18 underestimate activities that are characterized by more static types of activity or arm-only 19 work, as opposed to dynamic leg-exercise. Moreover, because the ability to obtain a precise 20 estimate of PAEE is an important factor in accurately establishing dose-responses relationships 21 between physical activity and disease, and secondly because heat-sink calorimeters provide an 22 accurate measure of PAEE but not minute-by-minute PAI, we used PAEE as the criterion 23 measure, as opposed to PAI. The estimated values for PAEE correlated with the measured PAEE values on the same level (R^2 =0.78) as the multiple regression model that used average 24 25 daytime HR and CSA. However, further validation in a more heterogeneous sample, and

1	preferably against doubly labeled water derived estimates of TEE, is needed before any of
2	these methods can confidently be applied to free-living populations. The branched model was
3	designed as a framework to interpret simultaneous HR and accelerometry data into minute by-
4	minute PAI. This is in contrast to the multiple regression model of PAEE (with CSA_{day} and
5	HR_{day}), which can only be used to estimate PAEE. Thus, the logical progression would be to
6	validate the branched model for combining accelerometry with HR data as a measure of PAI in
7	a range of activities, using a similar experimental design as Strath et al (35). It would also be
8	valuable to know how branched models perform in common occupational settings and
9	activities that predominantly involve arm work or static work. The method proposed in this
10	paper to estimate parameters in branched models is based on the same mathematical principle
11	as normal regression, i.e., minimizing the standard errors of the estimates. This approach
12	would, however, improve substantially by increasing the volume of data. Preferably the data
13	would also be derived from a range of different activity modes and intensities. Frequency,
14	duration, and total energy expenditure of physical activity can be derived from such an
15	intensity measurement, provided the time resolution is sufficiently high to capture the changes
16	in intensity.
17	In conclusion, the combination of HR and CSA data in a branched equation model improves
18	the estimate of PAEE in a population of trained young men, compared to either method used
19	alone or when the traditional non-branched combination is used. Our results also suggest that
20	individual calibration may not be as necessary when branched modeling is employed. We
21	hypothesize from these observations that in larger heterogeneous populations, more
22	parsimonious calibration procedures may be sufficiently precise when utilized in conjunction
23	with equations derived in smaller samples.

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TABLE 1Estimates of PAEE from CSA, HR, and their combination in non-branched (*ML*R and branched models (*a priori* and *post hoc*).

		Estimated PAEE, using individual calibration					Estimated PAEE, using group calibration				
		No	on-bran	ched	Bran	ched	No	on-brancl	ned	Bran	iched
Id	PAEE	CSA	HR	MLR CSA+HR	A priori CSA+HR	Post hoc CSA+HR	CSA	HR	MLR CSA+HR	A priori CSA+HR	Post hoc CSA+HR
1	26.9	15.7	57.4	61.5	37.6	31.4	15.9	35.2	39.9	25.5	30.4
2	41.5	17.6	77.6	84.8	47.3	42.7	18.6	66.9	51.8	42.1	37.4
3	28.2	8.6	62.2	57.0	36.2	33.6	12.9	56.8	44.3	36.0	32.3
4	41.2	19.4	22.2	9.7	20.8	34.2	19.7	63.0	52.7	41.5	39.4
5	42.6	18.6	41.1	54.5	29.3	32.3	21.8	49.2	41.3	35.0	36.8
6	33.4	14.3	47.0	27.9	33.1	33.3	18.0	41.2	33.9	31.0	30.3
7	26.9	14.8	19.7	26.8	17.2	21.9	16.5	24.5	24.5	20.5	24.6
8	36.9	16.0	55.0	53.6	37.0	37.8	19.1	58.8	52.3	40.7	38.7
9	42.6	19.9	91.3	97.8	57.4	44.2	25.5	90.6	64.0	57.6	45.8
10	28.0	15.8	33.3	11.4	24.6	28.4	16.3	46.3	39.6	32.4	29.6
11	30.7	21.0	29.2	25.1	24.3	30.4	21.5	52.6	43.9	39.2	33.1
12	24.1	13.4	25.7	22.2	19.6	23.4	13.2	24.4	20.3	18.8	22.6
Mean (SD)	33.6 (7.0)	16.3 (3.4)	46.8 (22.7)	44.4 (28.4)	32.0 (12.0)	32.8 (6.7)	18.3 (3.6)	50.8 (18.7)	42.4 (12.3)	35.0 (10.6)	33.4 (6.6)
SEE	0	18.2 ^{♣‡}	23.7‡	26.9 ^{♣‡}	10.0^{\dagger}	$4.4^{\dagger\ddagger}$	16.0 ^{‡*}	21.8 ^{†‡*}	11.9 ^{‡*}	6.6^{\dagger}	3.2^{\dagger}
SEE_{pred}	0	18.2 ^{♣‡}	20.7	25.0 [‡]	9.7^{\dagger}	5.8^{\dagger}	15.8 [‡]	18.2 [‡]	12.7 [‡]	6.0^{\dagger}	3.5^{\dagger}
R ² (p)	1 -	.37 (.037)	.20 (.143)	.23 (.116)	.27 (.086)	.61 (.003)	.61 (.003)	.59 (.003)	.53 (.004)	.61 (.003)	.78 (.000)

Data are in kJ·kg⁻¹, except R² (p); strength of association between measured and estimated PAEE. SEE is the standard error of the estimate using measured RMR in the calibration, SEE_{pred} is the corresponding SEE when using predicted RMR in the calibration. Significantly different from corresponding model using group calibration (p<.05). Significantly different from non-branched CSA+HR (*MLR*model on the same calibration level (p<.05). Significantly different from branched CSA+HR (*a priori*) model on the same calibration level (p<.05). Significantly different from corresponding model using predicted instead of measured RMR values in the calibration (p<.05).

TABLE 2Estimated parameters in the branched CSA+HR *post hoc* models, with the utilization of individual and group calibration, respectively. The equation structure of the branched models is shown in Figure 2.

	Individual calibration	Group calibration
X	4 (4)	35 (6)
Y_1	2.6 (2.5)	0.4 (-3.5)
Y_2	-63 (-62)	50 (224)
Z_1	-1.1 (-1.0)	-1.0 (-1.0)
Z_2	70 (68)	60 (64)
P_1	.61 (.64)	1.00 (1.00)
P_2	.32 (.27)	.21 (.21)
P_3	.18 (.26)	.21 (.21)
P_4	0.0 (0.0)	.10 (.10)
Utilization (1 / 2 / 3 / 4)	282 / 4260 / 768 / 3690	80 / 2869 / 2922 / 3129
	(408 / 4134 / 389 / 4069)	(230 / 4022 / 1468 / 3280)
HR / CSA contribution	22% / 78%	18% / 82%
	(16% / 84%)	(19% / 81%)

Parameters are obtained by minimizing the SEE or the 'trimmed' SEE (in parentheses).

Utilization is the number of observations that end up being quantified by boxes 1, 2, 3, and 4 in the equation structure. HR/CSA contribution is the sum of the fraction of observations being quantified by boxes 1-4 (Figure 2) times their weighting (P₁₋₄ and 1-P₁₋₄ for HR and CSA, respectively).

Figure legends:

FIGURE 1. Protocol for the calorimeter study with example of calorimeter data output.

SP=Sensible Power, EP=Evaporative Power, HS=Heat Source.

values are absolute HR minus RHR. All "PAI relationships" is determined by calibration. Therefore, this study has two equation complexes, depending on whether individual or group calibration is used. The equation complexes translate minute-by minute data into PAI as follows: If the CSA value is above X, we use Box 1 (with P₁) if the HR value is above Y, otherwise we use Box 2 (with P₂). Similarly, if the CSA value is below or equal to X, we use Box 3 (with P₃) if the HR value is above Z, otherwise we use Box 4 (with P₄). PAEE is obtained by integrating PAI with respect to time. The parameters **X**, **Y**₁₋₂, **Z**₁₋₂, and **P**₁₋₄ are either assumed *a priori* or can be estimated *post hoc* by simulation of all possible models, whilst minimizing the SEE between predicted and measured PAEE.

FIGURE 3. Bland-Altman plots of differences between measured and estimated PAEE.

Panels A+B are CSA estimates, C+D are flexHR estimates, E+F are CSA+HR (*MLR*)

estimates, G+H are branched CSA+HR (*a priori*) estimates, and I+J are branched CSA+HR

(*post hoc*) estimates. All left panels are results of models using individual calibration, right panels are models using group calibration. Lines are regression of the errors against measured PAEE with 95% error bands (broken lines).





