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1 **Predicting basal metabolic rate in men with motor complete spinal cord injury**

2

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12 **Running Title:** Basal Metabolic Rate in SCI

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29

30

31 **Abstract**

32 **Purpose:** To assess the accuracy of existing basal metabolic rate (BMR) prediction
33 equations in men with chronic (> 1 year) spinal cord injury (SCI). The primary aim is
34 to develop new SCI population-specific BMR prediction models, based on
35 anthropometric, body composition and/or demographic variables that are strongly
36 associated with BMR.

37 **Methods:** Thirty men with chronic SCI (Paraplegic; n = 21, Tetraplegic; n = 9), aged
38 35 ± 11 years (mean \pm SD) participated in this cross-sectional study. Criterion BMR
39 values were measured by indirect calorimetry. Body composition (dual energy X-ray
40 absorptiometry; DXA) and anthropometric measurements (circumferences and
41 diameters) were also taken. Multiple linear regression analysis was performed to
42 develop new SCI-specific BMR prediction models. Criterion BMR values were
43 compared to values estimated from six existing and four developed prediction
44 equations

45 **Results:** Existing equations that use information on stature, weight and/or age,
46 significantly ($P < 0.001$) over-predicted measured BMR by a mean of 14–17% (187–
47 234 kcal/day). Equations that utilised fat-free mass (FFM) accurately predicted BMR.
48 The development of new SCI-specific prediction models demonstrated that the
49 addition of anthropometric variables (weight, height and calf circumference) to FFM
50 (Model 3; $r^2 = 0.77$), explained 8% more of the variance in BMR than FFM alone
51 (Model 1; $r^2 = 0.69$). Using anthropometric variables, without FFM, explained less of
52 the variance in BMR (Model 4; $r^2 = 0.57$). However, all the developed prediction
53 models demonstrated acceptable mean absolute error $\leq 6\%$.

54 **Conclusion:** BMR can be more accurately estimated when DXA derived FFM is
55 incorporated into prediction equations. Utilising anthropometric measurements

56 provides a promising alternative to improve the prediction of BMR, beyond that
57 achieved by existing equations in persons with SCI.

58

59 **Key Words:** Basal Metabolism, Anthropometry, Body Composition, Spinal Cord
60 Injuries, Indirect Calorimetry.

61

62 **Introduction**

63 A critical determinant of body weight fluctuations over time is the imbalance between
64 energy intake and expenditure (kcal). Energy intake reflects the ingestion of
65 macronutrient food groups (carbohydrate, protein, fat and alcohol), whereas energy
66 expenditure can be partitioned into three components; basal metabolic rate (BMR),
67 dietary induced thermogenesis (DIT) and activity energy expenditure (AEE). BMR
68 represents the energy required to maintain homeostasis and the metabolic activities of
69 cells at rest. It is the largest component of total daily energy expenditure (TDEE),
70 approximately 70% for inactive persons with chronic spinal cord injury (SCI) (1). In
71 comparison to non-disabled controls, BMR is significantly reduced by 14 – 27% in
72 persons with SCI, although, values were comparable between groups when adjusted
73 for fat free mass (FFM) (2). Reductions in BMR after SCI are primarily driven by
74 skeletal muscle disuse atrophy below the level of the injury (3, 4). The adoption of a
75 more sedentary lifestyle after SCI reduces AEE (1, 5), further eroding TDEE, which
76 can lead to a sustained positive energy balance and thus the accumulation of excess
77 adiposity. Obesity, and its associated negative metabolic sequelae (i.e. impaired
78 glucose tolerance, insulin resistance and dyslipidaemia), commonly occurs at a
79 heightened frequency in persons with SCI (6-8).

80

81 Considering BMR accounts for the greatest proportion of TDEE in inactive
82 populations, its accurate measurement is of utmost importance. Multiples of BMR can
83 be used to derive an individual's daily energy needs and inform energy intake
84 adjustments in a clinical setting. From a public health perspective, the prescription of
85 a calorie-restricted diet is integral for obesity management, through the creation of a
86 sustainable energy deficit. The gold standard method for assessing BMR is indirect
87 calorimetry. However, this approach requires expensive, specialised equipment (i.e.
88 metabolic cart) which typically restricts its use to research settings. Accurate BMR
89 measurements should be performed upon waking in a quiet, darkened, thermal neutral
90 room, following an overnight fast, with participants in a complete resting posture. To
91 achieve these appropriate conditions, BMR is usually measured following an
92 overnight in-patient stay, which may be impractical. Consequently, in clinical
93 practice, BMR is often predicted using equations which feature variables that are
94 easily measured; body weight, stature and/or age (9-11). However, a recent review
95 reported that such equations, derived from able-bodied populations, over-predicted
96 BMR by 4 – 92% in persons with SCI (12). Variations in the prediction error across
97 studies likely reflect both error intrinsic to the equations themselves and variance
98 between study populations. For example, when using the equation from the seminal
99 work of Harris and Benedict (9), Aquilani *et al*, (13) observed only a 4%
100 overestimation compared to criterion BMR. Not only did these participants have sub-
101 acute injuries (~2 months post traumatic SCI) but they were also hypermetabolic due
102 to the presence of urinary tract infections and pressure injuries, which may explain the
103 reduced overestimation. Therefore, the accuracy of commonly used BMR prediction
104 equations remains to be assessed in a cohort representative of men with chronic (>1
105 year) SCI.

106 A major disadvantage of equations that utilise body weight to predict BMR is that this
107 variable is unable to distinguish between FFM and fat mass (FM). FFM has been
108 shown to explain most of the variance in BMR (14-16), with other studies
109 demonstrating an independent, secondary contribution of FM (17). In persons with
110 SCI, recent evidence would suggest incorporating FFM measured via dual energy X-
111 ray absorptiometry (DXA) more accurately predicts BMR than using height and
112 weight measurements (16). However, it is possible that prediction models utilising
113 FFM alone might not be sensitive enough to estimate individual BMR, and perhaps
114 other sources of variation (i.e. age and injury characteristics) should also be
115 considered (18, 19). Moreover, equations incorporating FFM also require the
116 acquisition of body composition data using expensive equipment (i.e. DXA), which
117 might not be available in a clinical setting, or inaccurate techniques (i.e. bioelectrical
118 impedance). Therefore, anthropometric measurements (i.e. circumferences and/or
119 diameters) might improve BMR prediction accuracy, with a trivial increase in
120 clinician/nutritionist workload to attain desirable predictor variables.

121

122 It remains to be seen whether the incorporation of injury characteristics could act as
123 surrogates for FFM or anthropometric measurements in the prediction of BMR. Both
124 level of injury and time since injury (TSI) influence body composition parameters (3,
125 20). Significant differences have been reported in BMR measured via indirect
126 calorimetry between paraplegic and tetraplegic participants (21). Utilising such easily
127 attainable injury characteristics to predict BMR in persons with SCI would further
128 reduce the burden on clinicians/nutritionists. The primary aim was to develop new
129 SCI population-specific BMR prediction models, based on injury characteristics or
130 anthropometric variables that are strongly associated with BMR. The secondary aim

131 of this study was to assess the accuracy of existing BMR predictive equations in men
132 with chronic (> 1 year) SCI.

133

134 **Methods**

135

136 *Participants*

137 Thirty men with chronic (> 1 year) motor complete (American Spinal Injury
138 Association Impairment Scale classification; A or B) SCI participated in this study.
139 All participants had lesion levels below C5 and were aged between 18 – 65 years old
140 with a BMI less than 32 kg/m². Exclusion criteria included; cardiovascular disease,
141 hypertension, type II diabetes, pressure ulcers greater than grade II and urinary tract
142 infection or symptoms. This experimental protocol was approved by the McGuire
143 Veteran Affairs Investigational Research Board and the Virginia Commonwealth
144 University (VCU) Office of Research and Innovation. All participants provided
145 written informed consent and procedures were conducted in accordance with the
146 principles of the Declaration of Helsinki.

147

148 *Basal metabolic rate*

149 Participants were woken up ~6.30 am, following a 12 hour overnight fast. All BMR
150 measurements were completed in a darkened, thermoneutral environment (ambient
151 temperature between 20-25°C). Participants abstained from caffeine, nicotine and
152 alcohol \geq 12 hours, in accordance with minimal criteria for best practice BMR
153 guidelines (22). A portable metabolic system (COSMED K4b², Rome, Italy) was used
154 to measure BMR. The unit was calibrated prior to use according to manufacturer's
155 instructions and has been demonstrated to be valid (23). Following calibration, a

156 canopy was placed over the participant's head as they lay in a supine position, with
157 continuous breath-by-breath measurements made over a 20-minute period. Gas
158 exchange values for the first 5 minutes were discarded, with BMR (kcal/day)
159 averaged over the last 15 minutes. Energy expenditure was determined using the Weir
160 equation (24). If respiratory exchange ratio (carbon dioxide production / oxygen used)
161 values were < 0.70 or > 1.00 participants were excluded from the analysis, as these
162 values are deemed indicative of protocol violations or inaccurate gas measurements
163 (22).

164

165 *Anthropometric measurements*

166 Prior to performing anthropometric measurements, participants were instructed to
167 void their bladder. Body mass (kg) was obtained using a digital wheelchair scale
168 (Tanita PW-630U, IL, USA), with the weight of the wheelchair subtracted from the
169 combined weight of participant and wheelchair to derive the participants mass.
170 Participants' height was measured in a supine position following transfer onto a mat.
171 The distance between two wooden boards, one at the apex of the head and the other
172 positioned at the sole of the foot, was measured using a Holtain height caliper to the
173 nearest 0.1 cm. For participants with knee flexion contracture, segmental measures
174 were taken from the greater trochanter to the lateral knee joint and from the lateral
175 knee to the lateral aspect of the sole of the foot.

176

177 Circumference measurements were taken using a standard inflexible measuring tape
178 (MFG, Lufkin, Executive Diameter Pocket Tape measure). The mean of three values
179 (within 0.5 cm of each other) was recorded to the nearest 0.1 cm. Abdominal
180 circumference was measured at the level of the umbilicus. Waist circumference was

181 measured at the midpoint between the crest of the ilium and the inferior margin of the
182 last rib. Hip circumference was measured around the widest part of the trochanters.
183 These measurements were taken after exhalation of a preceding deep breath. Thigh
184 and calf circumferences were also measured on the right leg. Thigh circumference
185 was measured at the midpoint between the anterior superior iliac spine and the
186 superior border of the patellar. Calf circumference was taken at the widest point. All
187 circumference measurements were taken in a supine position, except for the calf,
188 which was taken with participants sitting in their wheelchair. Sagittal and transverse
189 abdominal diameters (SAD and TAD) were also measured at the level of the
190 umbilicus in a supine position, using a Holtain-Kahn abdominal caliper.

191

192 ***Dual energy X-ray absorptiometry***

193 A trained operator measured body composition using a dual energy X-ray
194 absorptiometry (DXA) scanner (Lunar Prodigy Advance DXA scanner, WI, USA).
195 Whole-body lean mass, FM and bone mineral content (BMC) were extracted from
196 DXA computer software. FFM was calculated by adding BMC and lean mass. Whole-
197 body FFM was also predicted from body weight using the following equation, Gorgey
198 *et al.*, (25): $0.288 \times \text{body weight (kg)} + 26.3$. This was to assess whether, in the
199 absence of a direct DXA FFM measurement, predicted FFM could be used to
200 accurately predict BMR in persons with chronic SCI.

201

202 ***Basal metabolic rate prediction equations***

203 BMR (kcal/day) was estimated using three established equations, which incorporated
204 weight, height and age (9-11). For male adults, the Schofield equation utilised three
205 separate equations to predict BMR from weight, depending on the participants' age

206 group (age 18-30, 30-60, >60 years). This equation was previously used by the Food
207 and Agricultural Organization, World Health Organisation and United Nations
208 University (FAO/WHO/UNU) technical report series (26). BMR was also estimated
209 using body composition parameters (FFM and FM) (14, 16, 17). These equations are
210 described in full in Table 1.

211

212 *[PLEASE INSERT TABLE 1 ABOUT HERE]*

213

214 ***Statistical Analysis***

215

216 *Data modelling*

217 To explore the associations between criterion BMR and potential predictive traits,
218 simple univariate linear regressions were performed to derive Pearson correlation
219 values (r). A multivariate regression analysis, with both forward inclusion and
220 backward deletion, was then performed to develop SCI-specific BMR prediction
221 Models, incorporating the best combination of predictor variables (demographic
222 characteristics, anthropometric measurements and body composition parameters) that
223 explain the greatest variance in criterion BMR. Standard error of the estimate (SEE)
224 was also calculated to determine the accuracy of these prediction models. A 95%
225 Limits of Agreement (LoA) analysis was performed (mean difference \pm 1.96 SD)
226 comparing criterion and predicted BMR, with data displayed using Bland-Altman
227 plots.

228

229 *Error statistics*

230 Predicted BMR from each of the six established equations and generated prediction
231 models was compared to corresponding criterion BMR for each participant.
232 Comparison statistics included mean signed error (MSE) and mean absolute error
233 (MAE). Error of estimate data is presented as a percentage [Eq. Percentage error =
234 (Estimated BMR – criterion BMR) / criterion BMR × 100]. Differences between
235 predicted and criterion BMR were also compared by paired *t*-tests, with a Bonferroni
236 stepwise correction applied to correct for multiple comparisons. Statistical
237 significance was set at a priori of $\alpha < 0.05$ and all analyses were performed using
238 SPSS Statistics 25 for Windows (IBM, NY, USA).

239

240 **Results**

241 Participant demographics are presented in Table 2. Mean \pm SD measured BMR and
242 respiratory exchange ratio (RER) was 1499 ± 162 kcal/day and 0.83 ± 0.04 ,
243 respectively.

244

245 *[PLEASE INSERT TABLE 2 ABOUT HERE]*

246

247

248 *[PLEASE INSERT FIGURE 1 ABOUT HERE]*

249

250 *Associations between predictive traits and basal metabolic rate*

251 FFM measured by DXA explained most of the variance (69%) in BMR ($r = 0.83$; $P <$
252 0.01). Predicted FFM using Gorgey *et al.* (17) did not explain anymore of the
253 variance in BMR than weight, however, both were strongly associated with criterion
254 BMR ($r = 0.56$, $P < 0.01$). The predicted FFM equation significantly under-estimated

255 FFM by 3.6 kg ($P < 0.001$). Height and other anthropometric measurements (supine
256 waist and abdominal circumference, sitting calf circumference) were moderately
257 associated with BMR (Table 3). None of the demographic or injury characteristics
258 were associated with BMR.

259

260 *[PLEASE INSERT TABLE 3 ABOUT HERE]*

261

262 *Accuracy of developed prediction models*

263 The addition of circumferences and diameters to FFM (Model 2) slightly improved
264 the prediction of BMR in comparison to just FFM alone (Model 1) (Table 4).

265 However, the best prediction algorithm generated was Model 3 (incorporating FFM,
266 weight, height and calf circumference as predictor variables), which explained 77% of
267 the variance in BMR. For researchers/clinicians without access to expensive scanning
268 equipment (DXA), a final prediction algorithm was generated (Model 4), with the
269 FFM predictor variable removed. This explained the least variance in criterion BMR
270 ($r^2 = 0.57$). Relative to criterion BMR, mean bias for all the generated prediction
271 models was zero. The 95% limits of agreement (indicative of random error) were
272 greatest for Model 4 (anthropometrics alone: ± 207 kcal/day) and the smallest for
273 Model 3 (FFM plus anthropometrics: ± 152 kcal/day) (Figure 1). Entering predicted
274 FFM into Model 1 resulted in a mean bias $\pm 95\%$ LoA of -84 ± 262 kcal/day.

275

276 *[PLEASE INSERT TABLE 4 ABOUT HERE]*

277

278 *[PLEASE INSERT FIGURE 1 ABOUT HERE]*

279

280 *Accuracy of established and developed prediction models of basal metabolic rate*

281 The variability in error of established and newly developed BMR prediction equations
282 are displayed in Figure 2. Established equations, which feature variables that are
283 easily measured (body weight, stature and/or age), significantly ($P < 0.001$) over-
284 predicted measured BMR by a mean of 14 – 17% (187 – 234 kcal/day). Established
285 equations that utilised FFM (highlighted in grey) more accurately predicted measured
286 BMR in persons with SCI. The Nelson *et al*, (17) equation, which also incorporated
287 FM, significantly ($P < 0.001$) under-predicted BMR by $5 \pm 6\%$ (82 ± 95 kcal/day).
288 The remaining two established equations were not significantly different from the
289 criterion BMR and displayed negligible mean bias \pm SD; $-1 \pm 6\%$ (-20 ± 92 kcal/day)
290 and $1 \pm 6\%$ (3 ± 91 kcal/day) using the Cunningham, (14) and SCI-specific (16)
291 equations, respectively. Mean absolute percentage error for the generated Models
292 were small ($\leq 6\%$) and comparable to the Cunningham (14) and Chun *et al*, (16)
293 prediction equations. There was a trend ($P = 0.065$) for significantly elevated absolute
294 percentage error using predicted FFM in Model 1 ($8 \pm 6\%$) (not shown on Figure), as
295 opposed to DXA measured FFM ($5 \pm 4\%$).

296

297 *[PLEASE INSERT FIGURE 2 ABOUT HERE]*

298

299 **Discussion**

300 Existing equations developed for non-disabled individuals, which incorporate stature,
301 weight and/or age, significantly over-predicted BMR and are not fit for purpose in
302 person with SCI. Equations that utilise FFM, the Cunningham (14) and newly-
303 developed SCI-specific model (16), were not significantly different to criterion BMR.
304 In this sample of participants with chronic SCI, FFM as a single predictor variable

305 explained the greatest variance in BMR ($r^2 = 0.69$), which is in accordance with
306 previous studies ($r^2 = 0.63 - 0.79$) (2, 15, 27). However, the addition of volumetric
307 (circumferences and diameters) and anthropometric (height and weight)
308 measurements to FFM explained an additional 8% of the variance in BMR. Removal
309 of FFM from generated prediction models increased the prediction error, but offered a
310 useful alternative methodology in the absence of FFM measurement and improved the
311 prediction of BMR relative to existing equations validated for use in non-disabled
312 individuals.

313

314 We hypothesised that it might be possible to use certain demographic and injury
315 characteristics, such as age, level of injury and TSI, which are easily attainable and
316 thus reduce the burden on clinicians/nutritionists to predict BMR. We found no
317 significant differences in BMR between paraplegic (1497 ± 148 kcal/day) and
318 tetraplegic (1467 ± 178 kcal/day) participants. Previous studies have demonstrated
319 increased BMR in paraplegic compared to tetraplegic participants of 224 and 370
320 kcal/day (21, 28), whereas other researchers have shown there to be no difference (16,
321 29). One possible reason for similar BMR's between the subgroups in this current
322 study could be due to race. BMR has been shown to be higher in White than in
323 African-American individuals (30) and in this study, there was a greater percentage of
324 White participants with tetraplegia than paraplegia, 82% and 57%, respectively. Due
325 to the relatively small sample size and the requirement to develop models with
326 external validity to the wider male SCI population, it was not possible to develop
327 race-specific equations. As FFM is strongly associated with BMR, it is surprising that
328 age or TSI are not also associated with BMR, given the loss of skeletal muscle mass

329 with aging (31) and post SCI (3). It appears that these variables cannot be used as
330 surrogates for FFM in BMR prediction models for persons with SCI.
331
332 Besides skeletal muscle, bone mineral content (which contributes to FFM) is
333 significantly correlated to BMR ($r = 0.48$). Yilmaz *et al*, (28) demonstrated that hip
334 bone mineral density was significantly associated with BMR ($R_s = 0.41$) in persons
335 with SCI. These results indicate that bone metabolism is a major component of BMR
336 and might explain why height as an anthropometric variable explains 18% of the
337 variance in BMR. To date, no studies in persons with SCI have sought to assess the
338 improvement in the prediction of BMR with the addition of simple anthropometric
339 measurements that can be easily obtained. In non-disabled individuals, the addition of
340 FFM to a regression equation using the predictors of mass, height and age increased
341 the associations between predicted and criterion BMR from $r^2 = 0.71$, (SEE = 125
342 kcal/day) to $r^2 = 0.80$ (SEE = 103 kcal/day) (32). Similarly, the results of this current
343 study demonstrate the addition of anthropometric measurements to FFM (Model 3)
344 explains an additional 8% of the variance in BMR.
345
346 Whilst our generated multiple linear regression models demonstrate a negligible mean
347 bias (Figures 1 & 2), this can be somewhat misleading as under and over-estimations
348 for each participant likely cancel each other out. Using a limits of agreement analysis
349 (exploring the distribution of individual differences) and mean absolute percentage
350 error (ignoring the sign/direction of difference) are alternative approaches that offer
351 greater insight into the accuracy of developed models. The 95% LoA for all the
352 generated models ranged between ± 152 kcal (Model 3) to ± 207 (Model 4), which
353 are less than the values reported previously for the Cunningham (14) and SCI-specific
354 (16) equations, 236 and 231 kcal, respectively. Moreover, the mean absolute

355 percentage error was small, even for Model 4, which utilised only anthropometric
356 measurements (MAE = $6 \pm 4\%$), and were comparable to existing equations that
357 incorporate FFM. Therefore, in the absence of direct analyses of body composition,
358 we posit that the use of anthropometric measurements in models derived specifically
359 for males with chronic SCI can be used to improve the prediction of BMR. This is in
360 accordance with data from non-disabled individuals, which suggests utilizing
361 anthropometric data (height, weight, mid-upper arm and waist and hip
362 circumferences) provides a useful alternative methodology to better predict BMR
363 when detailed information on body composition is not available (33).

364

365 A recent systematic review highlighted the problems in predicting BMR in persons
366 with SCI from existing equations developed for non-disabled individuals (12). The
367 Harris Benedict (9) and Schofield *et al*, (11) equations have previously been shown to
368 over-predict BMR by 15-32% and 6% respectively (2, 34, 35). In conjunction with
369 findings herein, it is therefore not advisable to utilise equations developed for non-
370 disabled individuals that incorporate stature, weight and/or age to predict BMR in
371 persons with SCI. This study cross-validated, for the first time, the SCI-specific BMR
372 prediction equation developed by Chun *et al*, (16). This SCI-specific equation was
373 generated with criterion indirect calorimetry measurements taken between 8:00 and
374 10:00 am, rather than upon waking (~ 6:30am) in a darkened room following an
375 overnight stay. Occasionally in the wider literature, resting metabolic rate (RMR;
376 often measured under less restricted conditions) and BMR (as measured in this
377 current study) are often used interchangeably, but it is important to distinguish the
378 differences in terminology as this can help to reflect differences in prediction error
379 between studies. Moreover, the Chun *et al*, (16) equation was developed in East Asian

380 participants, with a considerably lower mean FFM than participants in this current
381 study (42.1 vs. 51.3 kg). Nevertheless, this equation showed the lowest mean \pm SD
382 bias of the pre-existing equations tested, $1 \pm 6\%$ (3 ± 91 kcal/day) and further
383 highlights the importance of incorporating a measurement of FFM into BMR
384 prediction models.

385

386 An alternative approach could be to utilise estimates of FFM, although whole-body
387 FFM was significantly under-predicted (3.6 kg) using the Gorgey *et al.*, (25) equation
388 in this study. Consequently, using estimates of FFM in Model 1 significantly ($P <$
389 0.001) under-predicted BMR (mean bias \pm 95% LoA; -84 ± 262 kcal/day), with
390 increased mean absolute percentage error ($8 \pm 6\%$). This equation estimates FFM
391 from weight, and weight itself explains the same amount of variance in criterion
392 BMR. Therefore, in the absence of expensive scanning equipment it is perhaps
393 advisable to use Model 4 (including height, weight and transverse abdominal
394 diameter) to predict BMR in persons with SCI. It is worth noting, that any error in the
395 estimation of BMR will be amplified if these data are used to derive an individual's
396 total daily energy expenditure (TDEE). For context, multiplying BMR by an activity
397 factor of 1.2 [as has been used previously in inactive persons with SCI (36)] would
398 equate to a TDEE of 1799 kcal/day in our sample. Extrapolating the mean absolute
399 error percentage for Model 3 & 4 indicates there is the potential to under or over-
400 predict TDEE by 72 and 108 kcal/day, respectively. Despite our generated equations
401 showing acceptable error ($< 5\%$), it is important for practitioners to be aware of the
402 implications of using predicted BMR to estimate TDEE, when looking to prescribe a
403 suitable energy intake in persons with SCI.

404

405 *Limitations*

406 The accuracy of the generated prediction models was assessed using the same sample
407 of participants that developed the model. In these circumstances evaluation statistics
408 (i.e. mean bias) can be somewhat biased (37). These equations were only tested in
409 men with motor-complete SCI to ensure a more homogenous sample. The
410 performance of these generated Models therefore remains to be assessed in women
411 with SCI, who represent 25% of the entire SCI population. It is possible the
412 development of future sex-specific Models are necessary to accurately predict BMR
413 in women with SCI. Spasticity, whereby motor control of skeletal muscles is
414 disturbed, occurs in more than 80% of persons with SCI (38). If episodes of spastic
415 hypertonia were to occur during the assessment of criterion BMR, this can lead to
416 increased energy expenditure due to excessive co-contraction (39). Therefore, future
417 studies should consider multiple measurements of BMR by indirect calorimetry to
418 accurately evaluate BMR in persons with severe spasticity (15). Although the use of
419 anthropometric measurements can improve the accuracy of BMR prediction and
420 potentially negate the requirement to use expensive scanning equipment (i.e. DXA), it
421 should be noted that transferring participants into the supine position could be
422 difficult. This is especially relevant when assessing persons with higher-level injuries
423 where access to lifting apparatus is not available.

424

425 **Conclusion**

426 Existing equations incorporating age, stature and weight that have been validated in
427 non-disabled individuals show considerable prediction error when used in persons
428 with SCI and are not fit for purpose. When direct measurements of FFM are available,
429 utilising FFM-based prediction equations offers a more accurate estimation of BMR,

430 which can be further improved with the incorporation of anthropometric
431 measurements. Moreover, in the absence of detailed body composition information,
432 utilising anthropometric measurements (height, weight and transverse abdominal
433 diameter) offers a useful alternative methodology to predict BMR in persons with
434 chronic SCI. However, these generated Models should be cross-validated with an
435 independent, larger sample of male and female participants, with a range of body
436 composition characteristics to demonstrate external validity to the wider SCI
437 population.

438

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450

451 **Conflict of Interest**

452 The authors have no conflict of interest to declare. The results of the study are
453 presented clearly, honestly, and without fabrication, falsification, or inappropriate

454 data manipulation. The results of the present study do not constitute an endorsement
455 by the American College of Sports Medicine.

456

457 **References**

- 458 1. Nightingale TE, Williams, S., Thompson, D., Bilzon, J.L.J. Energy balance
459 components in persons with paraplegia: daily variation and appropriate
460 measurement duration. *Int J Behav Nutr Phys Act.* 2017 [cited 2017 Oct
461 5];14(132). Available from:
462 <https://ijbnpa.biomedcentral.com/articles/10.1186/s12966-017-0590-z>. doi:
463 <https://doi.org/10.1186/s12966-017-0590-z>.
- 464 2. Buchholz AC, McGillivray CF, Pencharz PB. Differences in resting metabolic
465 rate between paraplegic and able-bodied subjects are explained by differences in
466 body composition. *Am J Clin Nutr.* 2003;77(2):371-8.
- 467 3. Spungen AM, Wang J, Pierson RN, Bauman WA. Soft tissue body composition
468 differences in monozygotic twins discordant for spinal cord injury. *J Appl*
469 *Physiol.* 2000;88(4):1310-5.
- 470 4. Moore CD, Craven BC, Thabane L, Laing AC, Frank-Wilson AW, Kontulainen
471 SA, et al. Lower-extremity muscle atrophy and fat infiltration after chronic spinal
472 cord injury. *J Musculoskelet Neuronal Interact.* 2015;15(1):32-41.
- 473 5. Buchholz AC, McGillivray CF, Pencharz PB. Physical activity levels are low in
474 free-living adults with chronic paraplegia. *Obes Res.* 2003;11(4):563-70.
- 475 6. Bauman WA, Spungen AM, Zhong YG, Rothstein JL, Petry C, Gordon SK.
476 Depressed serum high-density-lipoprotein cholesterol levels in veterans with
477 spinal-cord injury. *Paraplegia.* 1992;30(10):697-703.

- 478 7. Gater D. Pathophysiology of Obesity After Spinal Cord Injury. *Top Spinal Cord*
479 *Inj Rehabil.* 2007;12(4):20-34.
- 480 8. Bauman WA, Spungen AM. Coronary heart disease in individuals with spinal
481 cord injury: assessment of risk factors. *Spinal Cord.* 2008;46(7):466-76.
- 482 9. Harris JA, Benedict, F.G. *A biometric study of basal metabolism in man.*
483 Washington, DC: Carnegie Institute of Washington; 1919. p. 1-280.
- 484 10. Mifflin MD, Stjeor ST, Hill LA, Scott BJ, Daugherty SA, Koh YO. A new
485 predictive equation for resting energy-expenditure in healthy-individuals. *Am J*
486 *Clin Nutr.* 1990;51(2):241-7.
- 487 11. Schofield WN. Predicting basal metabolic rate, new standards and review of
488 previous work. *Hum Nutr Clin Nutr.* 1985;39 Suppl 1:5-41.
- 489 12. Nevin AN, Steenson J, Vivanti A, Hickman IJ. Investigation of measured and
490 predicted resting energy needs in adults after spinal cord injury: a systematic
491 review. *Spinal Cord.* 2016;54(4):248-53.
- 492 13. Aquilani R, Boschi F, Contardi A, Pistarini C, Achilli MP, Fizzotti G, et al.
493 Energy expenditure and nutritional adequacy of rehabilitation paraplegics with
494 asymptomatic bacteriuria and pressure sores. *Spinal Cord.* 2001;39(8):437-41.
- 495 14. Cunningham JJ. Body-composition as a determinant of energy-expenditure - a
496 synthetic review and a proposed general prediction equation. *Am J Clin Nutr.*
497 1991;54(6):963-9.
- 498 15. Gorgey AS, Chiodo AE, Zemper ED, Hornyak JE, Rodriguez GM, Gater DR.
499 Relationship of Spasticity to Soft Tissue Body Composition and the Metabolic
500 Profile in Persons With Chronic Motor Complete Spinal Cord Injury. *J Spinal*
501 *Cord Med.* 2010;33(1):6-15.

- 502 16. Chun SM, Kim HR, Shin HI. Estimating the Basal metabolic rate from fat free
503 mass in individuals with motor complete spinal cord injury. *Spinal Cord*.
504 2017;55(9):844-7.
- 505 17. Nelson KM, Weinsier RL, Long CL, Schutz Y. Prediction of resting energy-
506 expenditure from fat-free mass and fat mass. *Am J Clin Nutr*. 1992;56(5):848-56.
- 507 18. Lazzer S, Bedogni G, Lafortuna CL, Marazzi N, Busti C, Galli R, et al.
508 Relationship Between Basal Metabolic Rate, Gender, Age, and Body
509 Composition in 8,780 White Obese Subjects. *Obesity*. 2010;18(1):71-8.
- 510 19. McMurray RG, Soares J, Caspersen CJ, McCurdy T. Examining Variations of
511 Resting Metabolic Rate of Adults: A Public Health Perspective. *Med Sci Sports*
512 *Exerc*. 2014;46(7):1352-8.
- 513 20. Gorgey AS, Dolbow DR, Dolbow JD, Khalil RK, Castillo C, Gater DR. Effects
514 of spinal cord injury on body composition and metabolic profile - Part I. *J Spinal*
515 *Cord Med*. 2014;37(6):693-702.
- 516 21. Gorgey AS, Gater DR. Regional and relative adiposity patterns in relation to
517 carbohydrate and lipid metabolism in men with spinal cord injury. *Appl Physiol*
518 *Nutr Metab*. 2011;36(1):107-14.
- 519 22. Compher C, Frankenfield D, Keim N, Roth-Yousey L, Evidence Analysis
520 Working G. Best practice methods to apply to measurement of resting metabolic
521 rate in adults: a systematic review. *J Am Diet Assoc*. 2006;106(6):881-903.
- 522 23. McLaughlin JE, King, G.A., Howley, E.T., Bassett, D.R Jr., Ainsworth, B.E.
523 Validation of the COSMED K4b2 portable metabolic system. *Int J Sports Med*.
524 2001;22(4):280 - 4.
- 525 24. Weir JBD. New Methods for calculating metabolic rate with special reference
526 to protein metabolism. *J Physiol*. 1949;109(1-2):1-9.

- 527 25. Gorgey AS, Dolbow DR, Gater DR. A Model of Prediction and Cross-Validation
528 of Fat-Free Mass in Men With Motor Complete Spinal Cord Injury. *Arch Phys*
529 *Med Rehabil.* 2012;93(7):1240-5.
- 530 26. Energy and protein requirements. Report of a joint FAO/WHO/UNU Expert
531 Consultation. World Health Organization technical report series. 1985;724:1-206.
- 532 27. Bauman WA, Spungen AM, Wang J, Pierson RN. The relationship between
533 energy expenditure and lean tissue in monozygotic twins discordant for spinal
534 cord injury. *J Rehabil Res Dev.* 2004;41(1):1-8.
- 535 28. Hayes M, Chustek M, Wang ZM, Gallagher D, Heshka S, Spungen A, et al.
536 DXA: Potential for creating a metabolic map of organ-tissue resting energy
537 expenditure components. *Obes Res.* 2002;10(10):969-77.
- 538 29. Bauman WA, Cirnigliaro CM, La Fontaine MF, Jensen AM, Wecht JM,
539 Kirshblum SC, et al. A Small-Scale Clinical Trial to Determine the Safety and
540 Efficacy of Testosterone Replacement Therapy in Hypogonadal Men with Spinal
541 Cord Injury. *Horm Metab Res.* 2011;43(8):574-9.
- 542 30. Yilmaz B, Yasar E, Goktepe AS, Onder ME, Alaca R, Yazicioglu K, et al. The
543 relationship between basal metabolic rate and femur bone mineral density in men
544 with traumatic spinal cord injury. *Arch Phys Med Rehabil.* 2007;88(6):758-61.
- 545 31. Collins EG, Gater D, Kiratli J, Butler J, Hanson K, Langbein WE. Energy Cost of
546 Physical Activities in Persons with Spinal Cord Injury. *Med Sci Sports Exerc.*
547 2010;42(4):691-700.
- 548 32. Sharp TA, Bell ML, Grunwald GK, Schmitz KH, Sidney S, Lewis CE, et al.
549 Differences in resting metabolic rate between white and African-American young
550 adults. *Obes Res.* 2002;10(8):726-32.

- 551 33. Janssen I, Heymsfield SB, Wang ZM, Ross R. Skeletal muscle mass and
552 distribution in 468 men and women aged 18-88 yr. *J Appl Physiol*.
553 2000;89(1):81-8.
- 554 34. van der Ploeg GE, Gunn SM, Withers RT, Modra AC, Keeves JP, Chatterton BE.
555 Predicting the resting metabolic rate of young Australian males. *Eur J Clin Nutr*.
556 2001;55(3):145-52.
- 557 35. Johnstone AM, Rance KA, Murison SD, Duncan JS, Speakman JR. Additional
558 anthropometric measures may improve the predictability of basal metabolic rate
559 in adult subjects. *Eur J Clin Nutr*. 2006;60(12):1437-44.
- 560 36. Gorgey AS, Caudill C, Sistrun S, Khalil RE, Gill R, Castillo T, et al. Frequency
561 of Dietary Recalls, Nutritional Assessment, and Body Composition Assessment
562 in Men With Chronic Spinal Cord Injury. *Arch Phys Med Rehabil*.
563 2015;96(9):1646-53.
- 564 37. Staudenmayer J, Zhu W, Catellier DJ. Statistical considerations in the analysis of
565 accelerometry-based activity monitor data. *Med Sci Sports Exerc*. 2012;44(1
566 Suppl 1):S61-7.
- 567 38. Levi R, Hultling C, Seiger A. The Stockholm spinal-cord injury study .2.
568 Associations between clinical patient characteristics and post-acute medical
569 problems. *Paraplegia*. 1995;33(10):585-94.
- 570 39. Stoquart GG, Detrembleur C, Nielens H, Lejeune TM. Efficiency of work
571 production by spastic muscles. *Gait & Posture*. 2005;22(4):331-7.
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576 **Figure Legend**

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578 **Figure 1:** Bland-Altman plots depicting mean bias (solid line) and 95% limits of
579 agreement (dashed lines) of estimated relative to criterion basal metabolic rate
580 measured by indirect calorimetry for prediction Model 1 (FFM alone; **A**), 2 (FFM
581 plus anthropometrics and circumferences; **B**), 3 (FFM plus anthropometrics; **C**) and 4
582 (anthropometrics alone; **D**). Bias represents predicted-criterion BMR. Abbreviations:
583 BMR, basal metabolic rate.

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585 **Figure 2:** Scatterplot displaying BMR prediction error for each of the pre-existing
586 equations (absolute, **A**; percentage, **C**) and generated Models (absolute, **B**;
587 percentage, **D**). Mean error for each equation is displayed as a thick black bar, with
588 individual data points also shown (open circles). The highlighted areas (grey) are for
589 equations that utilize fat free mass (FFM) to predict BMR, with the dashed line
590 representing zero prediction error. Absolute error (accounting for under and over-
591 prediction) mean \pm SD is displayed for each equation above the Figures. # Significant
592 difference between predicted and criterion BMR ($P < 0.001$). Abbreviations: BMR,
593 basal metabolic rate.

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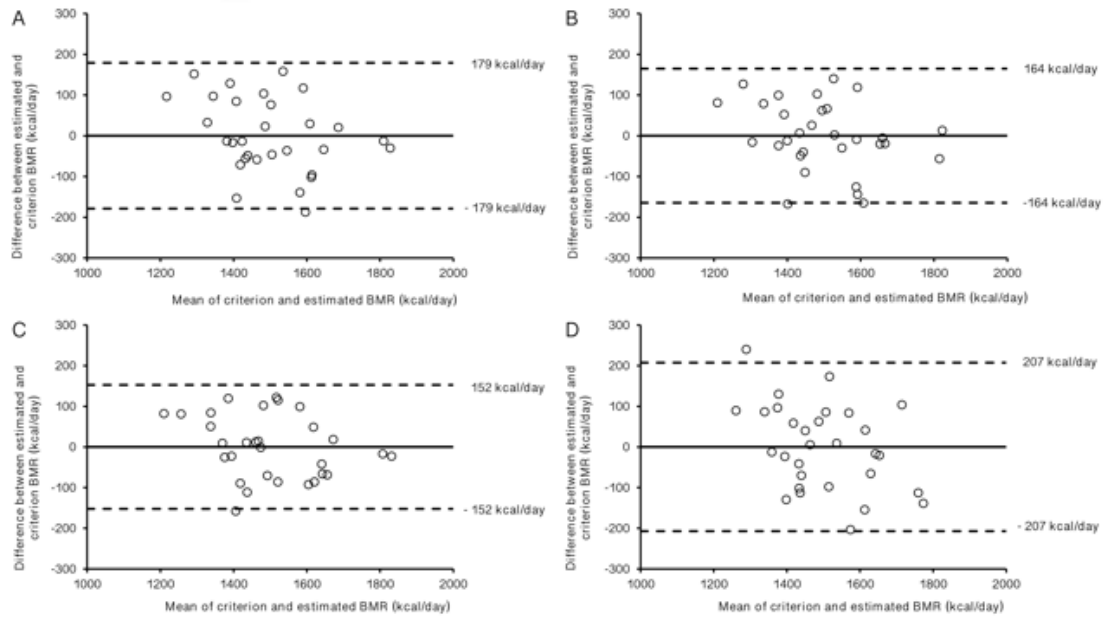
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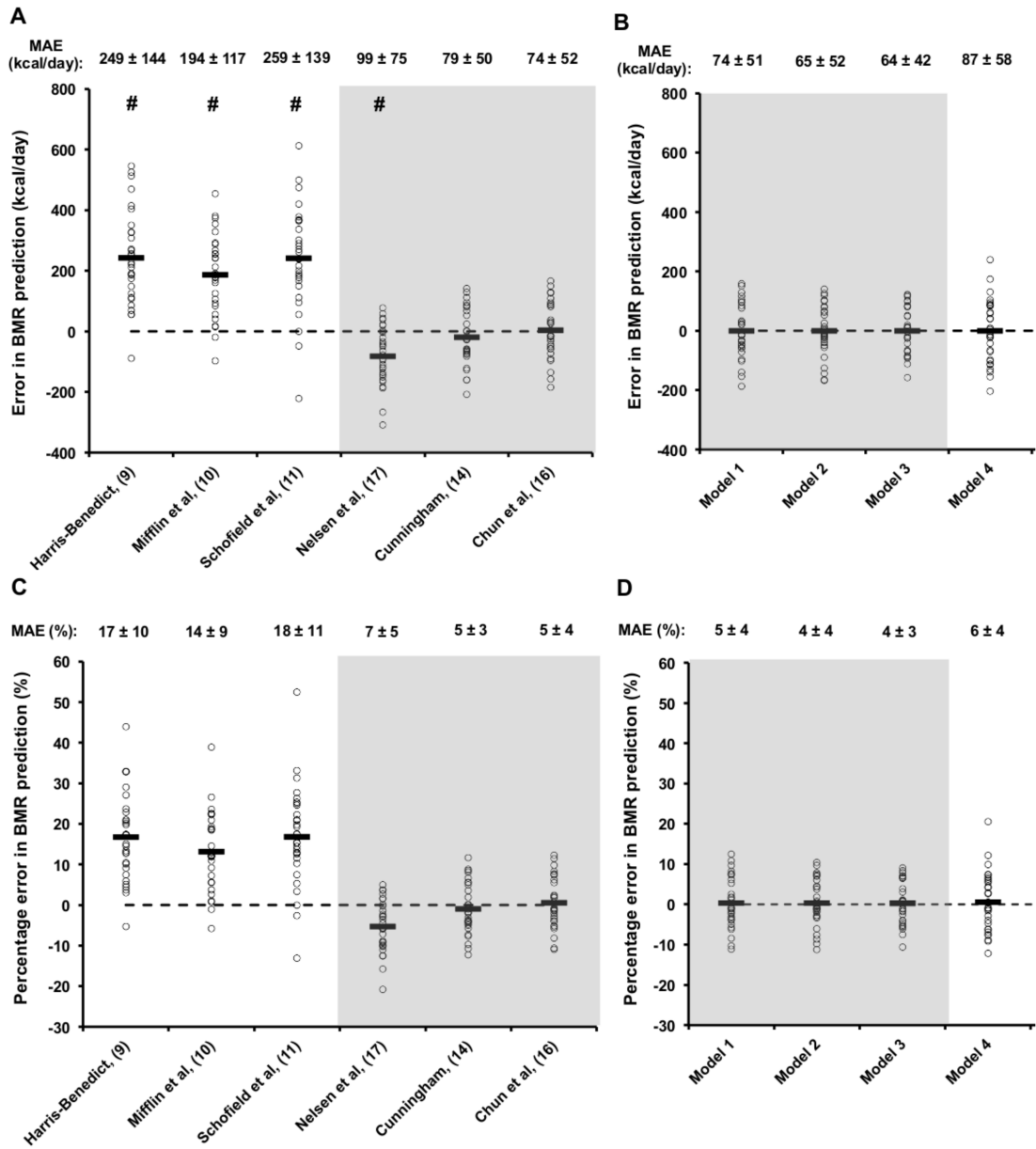
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602 **Figure 1**

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605 **Figure 2**

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613 **Table 1: Basal metabolic rate prediction equations**

Equation author	BMR prediction equation
<i>Weight, height and age</i>	
Harris-Benedict (9)	$= 66.4730 + (13.7516 \times \text{weight}) + (5.0033 \times \text{height}) - (6.7550 \times \text{age})$
Mifflin-St. Jeor (10)	$= 10 \times \text{weight} + 6.25 \times \text{height} - 5 \times \text{age} + 5$
Schofield (11)	$= 15.057 \times \text{weight} + 692.2$ (age, 18 – 30 years)
	$= 11.472 \times \text{weight} + 873.1$ (age, 30 – 60 years)
	$= 11.711 \times \text{weight} + 587.7$ (age, > 60 years)
<i>FFM and FM</i>	
Nelson <i>et al.</i> , (17)	$= 25.80 \times \text{FFM} + 4.04 \times \text{FM}$
Cunningham (14)	$= 370 + 21.6 \times \text{FFM}$
Chun <i>et al.</i> , (16) <i>SCI-specific</i>	$= (24.5 \times \text{FFM} + 244.4)$

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615 Abbreviations: **BMR**, basal metabolic rate; **FFM**, fat free mass; **FM**, fat mass.

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627 **Table 2: Participant characteristics**

Characteristic	Mean ± SD	Range (minimum – maximum)
<i>Age (years)</i>	35 ± 11	19 - 61
<i>Body mass (kg)</i>	74.5 ± 14.1	52.3 – 106.3
<i>Height (m)</i>	1.78 ± 0.05	1.69 – 1.87
<i>Race</i>	11 African American (37%) 19 white (63%)	
<i>Body fat (%)</i>	30.6 ± 10.2	14.8 – 48.2
<i>Fat mass (kg)</i>	22.9 ± 11.3	8.7 – 47.5
<i>Bone mineral content (kg)</i>	2.95 ± 0.39	2.09 – 3.66
<i>Fat free mass (kg)</i>	51.3 ± 5.7	41.4 – 64.7
<i>Level of injury</i>	9 Tetraplegic (30%) 21 Paraplegic (70%)	C5 – C7 T4 – L1
<i>TSI (years)</i>	9 ± 9	1 - 34
<i>AIS</i>	20 A (67%) 8 B (27%) 2 C (6%)	
<i>BMR (Kcal/day)</i>	1499 ± 162	1169 - 1843
<i>RER</i>	0.83 ± 0.04	0.74 – 0.90

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629 Abbreviations: **AIS, American Spinal Injury Association Impairment Scale;**
 630 **BMR, basal metabolic rate; RER, respiratory exchange ratio; TSI, time since**
 631 **injury.**

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643 **Table 3: The association (*r*) between independent predictive traits (injury and**
 644 **demographic characteristics, body composition components and anthropometric**
 645 **measurements) and criterion basal metabolic rate**

Demographic and injury characteristics		Body composition		Anthropometric measurements	
Age (yrs)	0.04	DXA-FFM (kg)	0.83†	Body mass (kg)	0.56†
LOI	0.22	DXA- FM (kg)	0.30	Height (cm)	0.42*
TSI (yrs)	0.06	DXA-BMC (kg)	0.48†	Supine waist circumference (cm)	0.41*
		Predicted FFM (kg)	0.56†	Supine abdominal circumference (cm)	0.37*
				Supine hip circumference (cm)	0.32
				Supine thigh circumference (cm)	0.27
				Sitting calf circumference (cm)	0.47†
				Supine SAD (cm)	0.30
				Supine TAD (cm)	0.29

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647 Abbreviations: **BMC, bone mineral content; DXA, dual-energy x-ray**
 648 **absorptiometry; FFM, fat free mass; LM, lean mass; LOI, level of injury; SAD,**
 649 **sagittal abdominal diameter TAD, transverse abdominal diameter; TSI, time**
 650 **since injury.**

651 *** P < 0.05, † P < 0.01**

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656 **Table 4: Generated basal metabolic rate prediction models using fat free mass**
 657 **and anthropometric measurements**
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Model name	BMR (kcal/day) prediction algorithm	R²	SEE (kcal/day)
1. FFM alone	= 23.469 × FFM (kg) + 294.330	0.69	93
2. FFM plus circumferences and diameters	= 23.995 × FFM (kg) + 6.189 × SAD (cm) + 6.384 × TAD (cm) – 6.948 × THIGH CIRC (cm) + 275.211	0.73	90
3. FFM plus anthropometrics	= 19.789 × FFM (kg) + 5.156 × weight + 8.090 × height – 15.301 × calf (cm) – 860.546	0.77	84
4. Anthropometrics alone	= 13.202 × height (cm) + 11.329 × weight (kg) – 16.729 × TAD (cm) – 1185.445	0.57	112

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 660 Abbreviations: **BMR, basal metabolic rate; FFM, fat free mass; SAD, sagittal**
 661 **abdominal diameter; SEE, standard error of the estimate; TAD, transverse**
 662 **abdominal diameter; THIGH CIRC, thigh circumference.**