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Assessing Resting Energy Expenditure in Overweight and Obese Adolescents in a Clinical Setting: Validity of a Handheld Indirect Calorimeter

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Shared first authorship

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Category of Study: Translational Research

RUNNING HEAD: REE in Overweight Adolescents

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ABSTRACT

Background: Accurately determining energy requirements is key for nutritional management of pediatric obesity. Recently, a portable handheld indirect calorimeter, MedGem (MG) has become available to measure resting energy expenditure (REE). Our work aims to determine the clinical validity and usefulness of MG to measure REE in overweight and obese adolescents.

Methods: 39 overweight and obese adolescents (16M: 23F, 15.2±1.9y, BMI percentile: 98.6±2.2 %) and 15 normal weight adolescents (7M: 8F, Age 15.2±2.0y, BMI percentile: 39.2±20.9%) participated. REE was measured with both MG and standard indirect calorimeter (VMax) in random order.

Results: MG REE (1600±372 kcal/d) was lower than VMax REE (1727±327 kcal/) in the overweight and obese adolescents. Bland Altman analysis (MG –VMax) showed a mean bias of -127 kcal/day (95%CI = -72 to-182 kcal/day, p<0.001), and a proportional bias existed such that lower measured REE by VMax was underestimated by MG, and higher measured REE by VMax were overestimated by MG.

Conclusions: MG systematically underestimates REE in the overweight and adolescent population, thus the MG portable indirect calorimeter is not recommended for routine use. Considering that it is a systematic underestimation of REE, MG may be clinically acceptable, only if used with caution.
INTRODUCTION

Childhood obesity is a major public health issue around the world and has a huge impact on the health of millions of young people. In Canada, nearly one out of three children and youth are overweight or obese (1). Energy balance is an important physiological aspect of obesity and sustainable weight loss may be achieved by creating a negative energy balance. To achieve an appropriate weight for age, the Canadian Obesity Guidelines suggests that a negative energy balance be achieved by a well-balanced, calorie-reduced diet and increased physical activity (2). The Academy of Nutrition and Dietetics recommends the use of indirect calorimeters to determine caloric targets for weight loss in obese youth (3). Both of these recent guidelines acknowledge the importance of understanding or measuring accurate energy requirements as part of a comprehensive pediatric nutrition assessment for weight loss in this population.

Resting energy expenditure (REE) represents 60-75% of daily total energy expenditure (TEE) (4), and can be quantified using traditional open-circuit indirect calorimeter carts or metabolic carts and is primarily based on measurement of oxygen consumption (VO\textsubscript{2}) and carbon dioxide (VCO\textsubscript{2})(5, 6). This method has long been established as the reference standard for measuring REE (7, 8). However, metabolic carts are expensive, require careful calibration, are less mobile and require longer testing time (9, 10).

Portable handheld devices such as the MedGem (MG), have become available as an alternative to the traditional calorimeter (11). MG measures VO\textsubscript{2} and calculates VCO\textsubscript{2}, based on an assumed constant respiratory quotient (RQ) of 0.85, where RQ is usually derived as a ratio of VCO\textsubscript{2}/VO\textsubscript{2}. The validity of the portable indirect calorimeter has been assessed in adults, both healthy (12 -14) and unhealthy (e.g.eating disorders, cancer) (15 -19) with varying results. In obese adults earlier studies have reported that the portable indirect calorimeters overestimate REE, when compared to the standard indirect calorimeters compared (20 - 22). In healthy...
normal weight children Fields et al (23) showed that hand held calorimeters overestimate REE, while Nieman et al (24) showed that the hand held calorimeter provides reliable REE measures compared to a Douglas bag method, a measure of pulmonary ventilation and respiratory gas exchange that provides an estimation of energy expenditure. Whether the portable indirect calorimeters are reliable and valid to use in an adolescent overweight and obese population is unknown. Our goal was to validate the use of MG in our pediatric Endocrinology Clinic and Type 2 Diabetes clinic. Thus, the primary objective in the current study is to compare measured REE from the handheld indirect calorimeter (MG) against a standard indirect calorimeter (VMax) in overweight and obese adolescents. Because energy requirements are often estimated using predictive equations (26, 27) in a clinical setting, our secondary objective is to compare measured REE against predicted REE in the overweight and obese adolescents. We hypothesized that the measured REE will be comparable between the portable and standard indirect calorimeters, and that the predicted REE using equations will not be comparable to the standard indirect calorimeter measured REE.

RESULTS
Participant Characteristics

Forty-one overweight, and fifteen normal weight (BMI between 15th and 85th percentile) children were recruited. Because two overweight participants had difficulties breathing in the supine position and their data did not reach steady state, data are presented for thirty-nine overweight children (23F:16M) and fifteen normal-weight (8F:7M) children. (Table 1). Weight and BMI (mean ± SD) were significantly higher (p<0.001) in the overweight group (88.1 ± 20.2 kg, 32.3 ± 5.8 kg/m²) than in the normal-weight group (51.3 ± 5.8 kg, 19.4 ± 1.9 kg/m²). Mean BMI of all children in the overweight group was ≥ 99th percentile for age and gender. Body fat
percentage was significantly higher in the overweight children when compared to the normal-weight children (46.5 ± 4.5% vs. 28.3± 7.1%, \(p<0.001\)). Fat free mass (FFM) was also significantly higher in the overweight children when compared to the normal-weight children (48.1 ± 7.5 kg vs. 36.8 ± 6.3 kg, \(p<0.001\)).

**REE Measurement by VMax and MG**

On average REE measured by MG was ~8% lower when compared to REE measured by VMax (1600 ± 404 kcal/d vs. 1727 ± 349 kcal/d, \(p<0.001\)) in the overweight group (Table 2). Similarly, on average REE measured by MG was ~16% lower than REE measured by VMax ((1181 ± 230 kcal/d vs.1370 ± 234kcal/d, \(p<0.001\)) in the normal-weight group. 29 of the 39 children (74%) had REE measured by MG within the clinically acceptable a priori set at ±250kcal/day. This is based on the calculation that a 250 kcal/d change in energy intake would lead to a half-pound weight loss in a week, as previously applied by Curtin et al (28), which is in line with recommendations by the American Academy of Pediatrics for treatment of and overweight and obese adolescents (29).

**Correlation Analysis**

Pearson correlation coefficient (\(r=0.91, p<0.001, 95\% CI = 0.83 - 0.95\)) showed a strong linear relationship between REE measured using MG and VMax in the overweight group (Figure 1). There was also a strong linear relationship (\(r=0.83, p<0.001, 95\% CI= 0.54, 0.94\)) between REE measured using MG and VMax in the normal-weight group (Figure 2).

**Bland Altman Analysis**

Mean overall difference in values obtained with the two different methods (bias) and limits of agreement (95% confidence limits for the bias) were determined. In the overweight children, there was a mean bias of -127±168 kcal/day between measured REE using MG when
compared to REE measured using VMax (Figure 3). The upper and lower 95% limits of agreement were 203 and -457 kcal/day. In addition to the fixed systematic bias, there was a proportional bias in the measured REE differences between the MG and VMax in the overweight group, such that lower measured REE by VMax was underestimated by MG, and higher measured REE by VMax were overestimated by MG. In the normal-weight children there was a mean bias of -190 ±136 kcal/day between measured REE using MG and VMax (Figure 4). The upper and lower 95% limits of agreement were 79 and -458 kcal/day. There was no proportional bias observed in the measured REE by MG in the normal-weight children.

Comparison of measure REE using MG against predictive equations

In the overweight/obese group predictive equations overestimate REE by ~30% (Table 3). The Molnar equation had the least mean bias of 2% when compared with the metabolic cart (VMax) REE predicted using the WHO/FAO/UNU equation showed an overestimation by up to 41%.

DISCUSSION

Results from our study indicate that the handheld indirect calorimeter (MG) significantly underestimates REE (mean bias, which represents a systematic error, of -127kcal/day) when compared to the standard indirect calorimeter (VMax), in an adolescent overweight population. A proportional bias also existed where the underestimation decreased with a higher measured REE. The limits of agreement, which reflects random error and represent the range of values in which the agreement between methods will lie for 95% of the sample, were +203 to -457kcal/day. These results suggest that the MG is not universally acceptable for routine use in the pediatric weight management practice. Considering that the systematic effect is an underestimate, if the MG is to be used in clinical settings for overweight and adolescent weight
management, caution needs to be exercised. Frequent monitoring of the caloric recommendation should be adopted for clinic patients.

To the best of our knowledge, our study is the first to compare MG to a standard indirect calorimeter in an adolescent overweight population and adds to the current body of related literature, where MG was tested against traditional indirect calorimetry or predictive equations in other populations. Earlier studies show conflicting results regarding REE comparisons between the portable and standard indirect calorimeters: in a larger sample of non-obese children (n=100), 10-13y, Fields et al (23) found MG to overestimate REE by 8% when compared with a traditional indirect calorimeter. On the other hand, two studies in adults reported that MG underestimated REE in stable patients on home nutrition support (17) and patients with cancer (19). In the study with patients on home nutrition support, a difference of ~130kcal/day between measurements of MedGem and a standard indirect calorimeter with limits of agreement beyond the clinically acceptable ±250kcal/day was observed (17). In the study with cancer patients (19), where the clinical acceptability was a priori set at <5% difference, less than half of the patients had measured REE by the handheld device within clinically acceptable limits. Thus, in general the MG measured REE in most populations does not compare well with the standard indirect calorimeter. One of the reasons for this lack of comparison could be due to the fact that the MG assumes a constant RQ of 0.85, and measures only VO\textsubscript{2} (11). In the present study a lower measured VO\textsubscript{2}, and an assumed RQ of 0.85 used to calculate VCO\textsubscript{2} combined to underestimate the REE consistently in both the overweight and obese children, and the healthy children.

REE is an essential component of total energy expenditure (TEE) as it represents up to 75% of the total energy needs. Thus, a reasonable measure of REE is important to estimate energy intake goals for weight management in overweight/obese individuals. Although REE can
be determined accurately using the traditional indirect calorimeters, its usage is limited due to the cost, maintenance and the expertise required in operating the device (5). Therefore in a clinical setting, energy requirements are estimated using predictive equations. Many studies have validated a variety of equations to estimate resting energy needs in the overweight adolescent population, although most equations are derived from a healthy weight cohort of children (26). Thus, predictive equations derived from normal weight populations may overestimate REE up to 25% in some cases (5, 30) in overweight and obese population. Body weights can be 20% or more than the ideal weight in obese individuals, and can introduce an additional value in the estimation of caloric needs. Although some studies have suggested the use of ideal body weight or adjusted body weight, (31), it is controversial and not widely followed. Among the predictive equations the Molnar equation (27) and Harris Benedict equation (32) compared reasonably well in the present study, and the FAO/WHO/UNU (33) equation provided significantly higher REE values in the obese and overweight population. The Molnar equation was developed in an adolescent population (27), and thus provides a reasonable alternative when directly measuring REE is not an option. Our results are in agreement with Hofsteenge et al (26) who found that the Molnar equation most accurately predicted REE when compared to measured REE by standard indirect calorimeter in 121 overweight and obese adolescents. Thus, while it is accepted that the standard indirect calorimeter is still the preferred method to estimate REE in adolescents, in clinical practice where the measurement is not available, the Molnar equation seems to predict REE accurately to make diet prescriptions.

Although the MG is likely to underestimate REE and has limits of agreement beyond a priori set at ±250 kcal/day in our findings, majority of the children (74%) were within the ±250 kcal/day. In addition the MG can offer an objective measure of REE as a starting caloric goal in
diet prescription. There is reasonable evidence that providing empowerment to vulnerable teenagers, such as obese/overweight children in our study with an objective measure of a caloric goal based on a personalized test, can provide additional motivation to make small changes in their dietary intake (34). These small changes in nutrition (energy intake) in addition to physical activity behaviors can prevent weight gain (35). Thus, an individually measured REE provides an additional tool to aid in the weight management strategy, as shown earlier in overweight US Air Force personnel (36). This aspect of an objective measurement of REE based diet and nutrition counseling, may provide an additional tool in the treatment and management of overweight and obese adolescents, and needs to be explored further.

In conclusion, the results from the current study showed that MG underestimates traditional indirect calorimetry measured REE systematically in overweight and obese adolescents, and cannot be recommended for routine use. The Molnar equation of predicting REE provides similar estimates to measured REE by a standard indirect calorimeter in this vulnerable population, and can be recommended, when traditional indirect calorimeters are not available. The possible application of a portable device in the day-to-day practice of weight management interventions for obese adolescents, especially in the outpatient setting is quite attractive. Obesity management in adolescents should be individually tailored and several factors should be taken into account, including sex, age, the degree of obesity, individual health risks, psycho-behavioral and metabolic characteristics. A handheld indirect calorimeter (MG) provides a practical alternative to traditional indirect calorimetry. Future studies are necessary to determine whether individually measured REE using handheld devices contributes to adolescent overweight/obese patient education, and improves effectiveness of weight loss recommendations.
METHODS

Participants

Forty-one children aged 12-18y who were overweight as defined by BMI between 15th and 85th percentile were recruited from the BC Children’s Hospital outpatient clinics and by study advertisements in clinical areas of the hospital. Fifteen children with similar age and normal weight were recruited from the community to establish study day protocol, and trouble shoot problems with usage of the hand held indirect calorimeter. All potential participants interested in the study underwent a phone prescreening, during which time a research assistant answered questions, and explained the study objectives and procedures. Written informed consent was obtained from all participants and legal representative before study start. All research protocols were approved by the University of British Columbia / Children’s and Women’s Health Centre of British Columbia Research Ethics Board (UBC C&W REB).

Study day procedures

The same trained technicians (PW, GM) carried out all REE measurements in a quiet and temperature-controlled (22°C) room in the Clinical Research and Evaluation Unit, BC Children’s Hospital. Children arrived for the study after an overnight fast (10-12h), including abstention from all beverages except water. All children reported limited physical activity in the 8h period prior to testing. Basic demographic information was collected using a questionnaire. Anthropometric measurements including body weight measured to the nearest 0.1kg with an electronic scale, and height without shoes was measured to the nearest 0.1cm with a stadiometer. Participants voided prior to arrival for the study. REE was determined using the hand-held indirect calorimeter (MG) and standard indirect calorimeter (VMax) on the same day for each participant in a random order.
REE measurement using standard indirect calorimeter (VMax)

Open-circuit indirect calorimetry (VMax Encore VE29n, Viasys Healthcare systems, Mississauga, ON, Canada) was performed using a clear ventilated canopy hood to measure oxygen consumption (VO₂) and carbon dioxide production (VCO₂) for 25 min (5). The VMax was calibrated daily using a certified calibration syringe for flow. Gas sensors for CO₂ and O₂ were calibrated prior to each measurement with standardized gases. Children remained in supine position, awake and motionless prior to and throughout the testing period to ensure valid and reproducible results (5). All children had their head slightly raised in order to minimize breathing irregularities, if any. The flow rate of room air drawn through the canopy was adjusted manually to maintain the fraction of CO₂ expired at optimal range (0.6 – 0.9%). Data from the first 5 min of the testing period were excluded to allow participant acclimatization (5). Steady state was defined as intervals of time (a minimum of 10 minutes), during which average VO₂ and VCO₂ varied less than 10%, and the respiratory quotient (RQ, VCO₂/VO₂) varied ≤ 5%. The Vmax software, which utilizes the modified Weir equation, was used to compute REE (5).

REE measurement using portable indirect calorimeter (Medgem, MG)

The hand-held indirect calorimeter used in this study was the MedGem (Microlife, Golden, CO). The device consists of an oxygen sensor and flow sensor to measure VO2, and REE is estimated using the Weir equation (5) under the assumption of a constant RQ (11). All participants rested in a relaxed seated position for 10 min prior to measurements. Participants stayed awake, upright and motionless throughout the testing period (8-12 min). The device was calibrated placing it on a flat surface before each measurement, as per manufacturer instructions. The nose clip was placed on the participant’s nose to close the nostrils and prevent breathing
through the nose and the mouthpiece was positioned in the mouth. Care was taken to ensure a firm seal around the mouthpiece. Each child was permitted to test the mouthpiece prior to testing to ensure comfort of nose clip and to check for air leaks. Blanket rolls were used to prop up the arms of the children to minimize discomfort or unnecessary movements. Children held the MedGem device in place until it beeped, indicating end of measurement.

**REE prediction using equations**

At BC Children’s Hospital (BCCH), The Harris Benedict equation (32), WHO/FAO/UNU equation (33) and the Molnar equation (26) (not in order of preference) are the three most commonly used to predict energy needs in the overweight/obese population. Hence, for the purpose of this study, REE was calculated using the above predictive equations, to compare against the measured REE using MG and VMax.

**Body composition measurement**

All children had their body composition analyzed using Bioelectrical Impedance Analysis (BIA) immediately following REE measurements with the standard indirect calorimeter, while still lying in a supine position. This allowed for stabilization of fluid volume. Children were positioned with limbs apart. Two electrodes were placed on the hand and foot as previously described (37). Body composition was determined using a bioelectrical impedance analyzer (Quantum IV, RJL Systems, Detroit, MI) at a frequency of 50 kHz with a constant current of 0.8 mA to obtain impedance (38). Three measurements on each child, not to vary by more than 1 Ω, were taken (37). Total body water (TBW) was estimated using the resistance index (RI=Height²/Resistance) and age/sex specific equations (39, 40) were used to calculate fat mass (FM) and fat free mass (FFM).
Statistical analysis

Results are represented as mean ± SD. Paired student t test was used to assess systematic differences between REE measured by VMax and MG. Pearson’s correlation analysis was used to detect linear agreement between REE measured using VMax and MG. Since correlation between measures does not provide information about the differences between the two measures, Bland-Altman analysis (41 - 44) was used to assess the degree of agreement, mean bias and proportional bias between the two indirect calorimeter assessments of REE. Mean bias determines the quantitative difference between the two measurements, and the proportional bias determines whether the magnitude of the measurement affects the quantitative difference. In order to interpret Bland-Altman analysis, a ±250kcal/day in REE measured by the two techniques was set a priori as a clinically significant difference. This is based on the calculation that a 250 kcal/d change in energy intake would lead to a half-pound weight loss in a week, as previously applied by Curtin et al (28), which is in line with recommendations by the American Academy of Pediatrics for treatment of and overweight and obese adolescents (3, 29).

Data was analyzed using SPSS version 17 (SPSS Inc, Chicago, IL) and Bland-Altman plots were constructed using Med-Calc version 9.3.1 (MedCalc Software, Ostend, Belgium).

ACKNOWLEDGEMENTS

We would like to thank Jennifer Kwan for help with study coordination.
REFERENCES


FIGURE LEGENDS

Figure 1. Correlation between MG and VMax REE measurements in the overweight group

Pearson r = 0.91, *p<0.001, n=39

Figure 2. Correlation between MG and VMax REE measurements in the normal-weight group

Pearson r = 0.83, *p<0.001, n=15

Figure 3. Bland-Altman plot showing the mean bias and limits of agreement between MG and VMax in overweight group

The solid line represents mean bias (-127.1 kcal/d, a systematic underestimation in MG REE) and dashed lines represent +/- 2 standard deviations (Limits of Agreement). The dotted line represents the linear trend of the differences between the two measures depicting proportional bias (*p<0.042, n=39)

Figure 4. Bland-Altman plot showing the mean bias and limits of agreement between MG and VMax in normal-weight group

The solid line represents mean bias (-189.7 kcal/d, a systematic underestimation in MG REE) and dashed lines represent +/- 2 standard deviations (Limits of Agreement). The dotted line represents the linear trend of the differences between the two measures depicting no proportional bias (p=0.923, n=15)
### Table 1. Subject characteristics\(^a\)

<table>
<thead>
<tr>
<th>Variable</th>
<th>Overweight (n=39)</th>
<th>Normal-weight (n=15)</th>
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<tbody>
<tr>
<td>Age (y)</td>
<td>15.2 ± 1.8 (12.4-18.8)</td>
<td>15.5 ± 2.0 (12.6-18.4)</td>
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<td>Sex (f/m)</td>
<td>23/16</td>
<td>8/7</td>
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<tr>
<td>Height (cm)</td>
<td>164.4 ± 8.7 (141-189.3)</td>
<td>162.2 ± 6.9 (152.3-174.5)</td>
</tr>
<tr>
<td>Weight (kg)</td>
<td>88.1 ± 20.2* (55.5-147.9)</td>
<td>51.3 ± 5.8 (41.7-66.2)</td>
</tr>
<tr>
<td>BMI (kg/m(^2))</td>
<td>32.4 ± 5.8* (23.4-53.7)</td>
<td>19.4 ± 1.9 (15.8-22.4)</td>
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<td>FFM (kg)</td>
<td>46.6 ± 8.9 (30.2-71.3)</td>
<td>36.8 ± 6.3 (29.3-52.9)</td>
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<td>Body Fat (%)</td>
<td>46.1 ± 4.5* (36.4-57.1)</td>
<td>28.3 ± 7.1 (18.6-38.1)</td>
</tr>
</tbody>
</table>

\(^a\)Values are mean ± SD (range)

\(^*\)Significantly (p<0.001) different compared to normal-weight group
Table 2. Measured resting energy expenditure (REE) and VO₂ using MedGem (MG) and Standard Indirect calorimeter (VMax)\(^{a,b}\)

<table>
<thead>
<tr>
<th></th>
<th>Overweight (n=39)</th>
<th>Normal-weight (n=15)</th>
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</thead>
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<tr>
<td>REE (kcal/day)</td>
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<tr>
<td>VMax</td>
<td>1727 ± 349</td>
<td>1370 ± 234</td>
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<tr>
<td>MedGem (MG)</td>
<td>1600 ± 404(^*)</td>
<td>1181 ± 230(^*)</td>
</tr>
<tr>
<td>VO₂ (ml/min)</td>
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<td></td>
</tr>
<tr>
<td>VMax</td>
<td>245 ± 50</td>
<td>194 ± 32</td>
</tr>
<tr>
<td>MedGem (MG)</td>
<td>231 ± 58</td>
<td>170 ± 31</td>
</tr>
</tbody>
</table>

\(^a\)Values are mean ± SD

\(^b\)No comparisons were made between Overweight and normal-weight group

\(^*\)Significantly (p<0.001) different when compared to REE from VMax
Table 3. Predicted resting energy expenditure (REE) compared to measured resting energy expenditure a

<table>
<thead>
<tr>
<th>Predictive Equation</th>
<th>Overweight (n=39)</th>
<th>Normal-weight (n=15)</th>
</tr>
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<tbody>
<tr>
<td>Measured REE (VMax)</td>
<td>1727 ± 349</td>
<td>1370 ± 234</td>
</tr>
<tr>
<td>Measured REE (MG)</td>
<td>1600 ± 404</td>
<td>1181 ± 230</td>
</tr>
<tr>
<td>Harris Benedict</td>
<td>1741 ± 248</td>
<td>1438 ± 116*</td>
</tr>
<tr>
<td>WHO/FAO/UNU</td>
<td>1890 ± 359*</td>
<td>1459 ± 150*</td>
</tr>
<tr>
<td>Molnar</td>
<td>1758 ± 325</td>
<td>n/a</td>
</tr>
</tbody>
</table>

aValues are mean ± SD

*significantly (p<0.01) higher than REE using MedGem (MG)

**significantly (p<0.001) higher than REE using MedGem (MG)

n/a- equation derived for use in obese population
Figure 1
Figure 3
Figure 4