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1 **Introducing eccentric cycling during a multidisciplinary weight loss intervention might**
2 **prevent adolescents with obesity from increasing their food intake: the TEXTOO study**

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27
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33
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41 (IUF) and the National Academy of Medicine.

42 **Abstract**

43
44 **Purpose.** The present study compared the appetite responses to an inpatient eccentric *versus*
45 concentric cycling training programs in adolescents with obesity.

46 **Methods.** 24 adolescents with obesity (12-16yrs; Tanner 3-4) followed a 12-week
47 multidisciplinary intervention (Phase1), after which they were randomized to concentric (CON)
48 or eccentric (ECC) training for 12 weeks (Phase2). Assessment of anthropometrics, body
49 composition (DXA), aerobic power (VO_{2max}), energy (EI) and macronutrient intake, food
50 reward, and subjective appetite were performed at baseline, and after Phase1 (T1) and Phase2
51 (T2).

52 **Results.** Body mass, BMI, and fat mass (FM%) decreased in both groups ($p < 0.001$). FM%
53 reduction was greater in ECC at T2 (-9.9%). EI did not change in either group at T1, but was
54 greater at T2 relative to T1 in CON only ($p < 0.001$, +22%). There was no correlation between
55 the change in body mass, FM%, fat-free mass and EI. Hunger ($p = 0.002$) and desire to eat
56 ($p = 0.001$) were higher in CON vs. ECC with no time effects nor interactions. Prospective food
57 consumption increased in both groups with no group effect nor interaction. Satiety was not
58 different between groups or over time. In ECC, preference for high-fat foods increased
59 ($p = 0.03$), and preference ($p = 0.004$) and implicit wanting ($p = 0.016$) for sweet foods decreased.

60 **Conclusion.** Eccentric cycling as part of an inpatient multidisciplinary weight-loss intervention
61 might help prevent increased *ad libitum* energy intake compared to concentric exercise training
62 in adolescents with obesity, potentially through distinct effects of the food reward system.

63
64
65 **Key words.** Eccentric cycling, Appetite, Energy intake, Food reward, Pediatric obesity

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81 **Introduction**

82 With one out of five children having obesity in Europe (WHO. 2016), pediatric obesity is an
83 alarming public health concern. This is particularly concerning since about 83% of adolescents
84 with overweight and obesity will remain so in adulthood (Herman et al. 2009) and will develop
85 associated metabolic disorders that might increase their morbidity and reduce their life
86 expectancy (Lifshitz 2008). Developing effective weight loss strategies is of high importance.
87 Multidisciplinary interventions combining physical activity, nutritional guidelines and
88 psychological support are recommended among youth with obesity (Boff et al. 2017; Cardel et
89 al. 2019). Such interventions have been shown effective, at least in the short term, leading to
90 significant improvements in body mass and body composition (Zolotarjova et al. 2018),
91 physical fitness (Aguer et al. 2010), cardio-metabolic profile (Caranti et al. 2007), and health-
92 related quality of life (Fonvig et al. 2017), among others factors.

93 More recent studies have also investigated the potential appetite and eating behavior
94 responses to such weight loss interventions, including subjective appetite sensations, energy
95 intake, macronutrient preferences and food preferences (for review see (Schwartz et al. 2016;
96 Thivel et al. 2019)). Most of the available literature seems to indicate that physical activity-
97 based weight loss interventions favor a reduction in food intake among adolescents with obesity
98 (Schwartz et al. 2016). However, these data are based on self-reported measures, which have
99 been shown to provide under-reported food intake, especially among adolescents with obesity
100 (Burrows et al. 2010). Some studies using objective food intake measurements using *ad libitum*
101 test meals however report increased daily intake in both adults (Myers et al. 2019) and
102 adolescents (Miguet et al. 2019) with obesity. Interestingly, Miguet and colleagues found a
103 significant increase in energy intake after the second phase (5 months) of a 10-month
104 multidisciplinary intervention conducted among adolescents with obesity, compared with the
105 first five months (Miguet et al. 2019). This increase in energy intake could explain the

106 attenuated reduction in body mass and fat mass percentage observed during the second phase
107 of similar interventions (Miguet et al. 2019; Carnier et al. 2013; Julian et al. 2019). Julian and
108 collaborators recently compared the efficacy of eccentric (ECC) *versus* concentric (CON)
109 cycling of similar metabolic loads as part of an inpatient multidisciplinary weight loss program
110 among adolescents with obesity, placing the exercise training in this second phase of the
111 intervention (Julian et al. 2019). According to their results, both exercise interventions favored
112 improved body mass, body composition and aerobic capacity. Interestingly, these
113 improvements were greater in ECC, particularly for fat mass, fat-free mass, lower-limb muscle
114 strength as well as insulin resistance (Julian et al. 2019).

115 Due to its lower metabolic demand compared with CON exercise when performed at
116 the same mechanical power (Julian et al. 2018), ECC exercise seems particularly suitable for
117 patients with chronic diseases and limited functional and cardiorespiratory fitness. Importantly,
118 the use of eccentric cycling exercise should be prioritized over classical eccentric weight-lifting
119 exercise among patients with obesity, particularly youth. Indeed, cycling exercise, as a non-
120 weight-bearing exercise modality, limits the high solicitation of bone and joints imposed by
121 classical weight-lifting, thus reducing potential pain and injuries. Additionally, eccentric
122 cycling favors the use of a larger muscle mass, larger time under tension and a better control of
123 the imposed mechanical load, optimizing its quantification and the effects of the intervention.

124 It has also been suggested that ECC exercise could impact mechanisms of appetite
125 control, but this remains to be elucidated (Julian et al. 2018). ECC exercise might indeed have
126 particular effects on the physiological pathways involved in the control of energy intake, mainly
127 through its specific effects on fat-free mass and fat mass (Julian et al. 2018). Fat-free mass and
128 fat mass are both involved in the control of food intake, mainly through their impact on resting
129 metabolic rate as well as insulin and leptin sensitivity (Blundell et al. 2012). Interestingly, ECC
130 exercise has shown to increase brain-derived neurotrophic factor (BDNF) concentrations in

131 both animal models and humans, which is a strong central regulator of energy intake (Hu and
132 Russek 2008; Matthews et al. 2009; Yarrow et al. 2010). In addition to such physiological
133 factors, exercise in general has also been shown to affect energy intake through hedonic
134 pathways (Miguet et al. 2018; McNeil et al. 2015), but this remains to be examined in response
135 to ECC exercise. Although Paschalis and colleagues did not observe any changes in total
136 energy and macronutrient intake in response to an 8-week ECC program compared to CON,
137 their study only included healthy young women (Paschalis et al. 2011). To our knowledge, the
138 appetite and eating behavior responses to chronic ECC exercise among patients with obesity,
139 particularly children and adolescents, remain unexplored. Therefore, the aim of the present
140 work was to compare the effect of an inpatient ECC *versus* CON cycling training program on
141 energy intake, appetite sensations and food reward in adolescents with obesity.

142

143 **Methods**

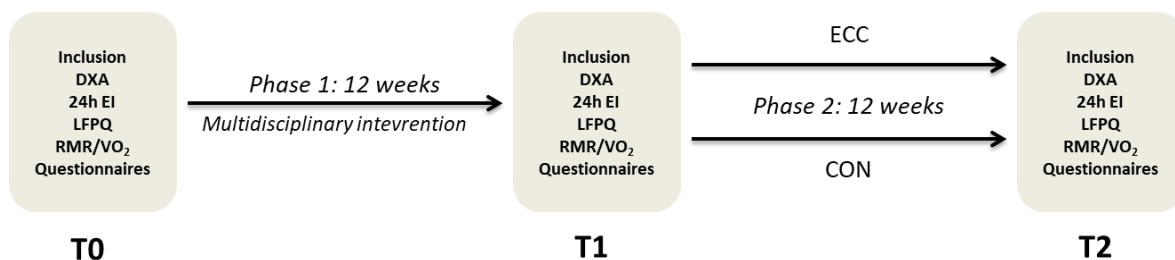
144 **Subjects**

145 Twenty-four adolescents with obesity (12 males and 12 females; Tanner Stages 3-4) were
146 recruited from the Pediatric Obesity Center (Tzanou, la Bourboule, France). To be included,
147 the adolescents had to meet the following criteria: 1) age 12-16 years, 2) having obesity (not
148 overweight) with body mass index [BMI] > 90th percentile, according to the international cut-
149 off points (Cole et al. 2000), 3) being free of any medication that could affect energy
150 metabolism; 4) not being regular tobacco or alcohol consumers, and 5) free of any
151 contraindications to exercise. Detailed information sheets were provided to the adolescents and
152 their legal representatives and written informed consent was obtained. This study received
153 ethical approval (*CPP Est IV*, 2016-A00043-48; ClinicalTrials.gov: NCT02925572).

154

155 **Study design**

156 This is a secondary analysis of the TEXTOO study and the entire design has been previously
 157 described (Julian, et al. 2019). The main aim of the TEXTOO study was to compare the effect
 158 of an inpatient ECC or CON cycling exercise weight loss intervention inducing the same
 159 metabolic load, on body composition and muscle strength in adolescents with obesity. Briefly,
 160 after a full medical examination to control for their ability to realize the study, the adolescents
 161 followed 12 weeks of inpatient classic medical care (Phase 1) (Prado et al. 2009). They were
 162 then randomized and assigned, using randomized blocks, to 12 weeks of concentric (CON; 6
 163 girls and 6 boys) or eccentric (ECC; 6 girls and 6 boys) cycling training (Phase 2).
 164 Anthropometric and body composition measurements, maximal incremental exercise tests and
 165 a full appetite and eating behavior assessment day (daily energy and macronutrient intake, food
 166 reward, and appetite sensations) were performed at baseline (T0), after Phase 1 (T1) and after
 167 the 12 weeks of CON or ECC training (Phase 2, T2). The study design is illustrated in Figure
 168 1.
 169



170
 171 **Figure 1.** Design of the study. T0, baseline; T1, end of phase 1; T2, end of phase 2; CON, Concentric cycling;
 172 ECC, Eccentric cycling; DXA, Dual X-ray absorptiometry; EI, Energy Intake; LFPQ, Leeds Food Preference
 173 Questionnaire; RMR, Resting Metabolic Rate.

174
 175

176 **Anthropometric measurements and body composition**

177 Body mass and height were respectively recorded to the nearest 0.1 kg and 0.5 cm using a
 178 digital scale and a standard wall-mounted stadiometer (SECA, les Mureaux, France). The

179 adolescents were required to be wearing light clothes and be bare-footed. BMI was calculated
180 as body mass (kg) divided by height squared (m²) and was reported in the age- and sex-
181 dependent international reference curves to get the BMI percentile. Fat mass (FM) and fat-free
182 mass (FFM) were assessed, by dual-energy X-ray absorptiometry (DXA) following
183 standardized procedures (QDR4500A scanner, Hologic, Waltham, MA, USA).

184

185 **Maximal oxygen uptake**

186 An incremental exercise test was performed on a traditional concentric cycling ergometer
187 (Ergoselect 100 P, MSE medical, France) that was different from the one used for the CON
188 cycling training (Optibike Med 600, MSE Medical, France). The initial power was set at 30W
189 during 3 minutes, followed by 15W increments every minute until exhaustion. The
190 experimenters strongly encouraged the adolescents during the test to favor a maximal effort. At
191 least two of the following criteria had to be reached for the test to be defined as maximal: heart
192 rate above 90% of the theoretical maximum heart rate, respiratory exchange ratio
193 ($RER = \dot{V}CO_2/\dot{V}O_2 > 1.1$; $\dot{V}O_2$ plateau. The theoretical maximal heart rate was calculated using
194 equations developed in children ($208 - 0.7 * \text{age}$) (Tanaka et al. 2001). Cardiac electrical activity
195 was continuously monitored (Ultima SeriesTM, Saint Paul, MN) and combined with breath-by-
196 breath gas exchange measurement (BreezeSuite Software, Saint Paul, MN) to determine $\dot{V}O_2$
197 and $\dot{V}CO_2$. $\dot{V}O_{2\text{peak}}$ was defined as the average of the last 30s of exercise before exhaustion
198 (Rowland 1996). All the adolescents enrolled met these criteria at each time point of the study.

199

200 **Resting metabolic rate**

201 Resting metabolic rate (RMR) was assessed in the morning under fasted state using indirect
202 calorimetry (K4b², Cosmed, Rome, Italy). The gas analyzer was calibrated in accordance with
203 the manufacturer's recommendations before each evaluation. Participants were asked to stay at

204 rest in a supine position in a thermoneutral room (22–25°C ambient temperature) for 45 min
205 before starting the measurement. After reaching a steady state, $\dot{V}O_2$ and $\dot{V}CO_2$ standardized for
206 barometric pressure, temperature, and humidity were recorded every minute for 20–45 min and
207 averaged over the entire measurement period. Resting metabolic rate (in kcal/day) and RER
208 were then calculated (Weir 1949).

209

210 **Eating behavior traits**

211 The Dutch Eating Behavior Questionnaire (DEBQ) was used to assess the adolescents' eating
212 behavior traits (Van Strien et al. 1986). Three domains were studied: restrained eating
213 (individuals' efforts to limit their food consumption to control their body mass or to generate
214 weight loss; 10 items), emotional eating (excessive eating in response to negative moods; 13
215 items) and external eating (eating in response to food-related stimuli, regardless of the internal
216 state of satiety or hunger; 10 items). Items were answered using six-point Likert scales (never
217 (1), seldom (2), sometimes (3), often (4), very often (5) and not relevant (0)). Their answers
218 were coded according to the recommendations given by Lluch et al. (Lluch et al. 2000). Mean
219 values were calculated for each domain: higher scores indicating greater endorsement of the
220 eating behavior. Scores equal to or above three were used to determine adolescents as
221 restrained, emotional and/or external eaters (Antinori 2004). The French version of this
222 questionnaire has been validated (Lluch et al. 2000).

223

224 **Food consumption**

225 Daily food consumption was assessed using *ad libitum* buffet meals at baseline (T0), after Phase
226 1 (T1) and after Phase 2 (T2). As previously detailed (Miguet, et al. 2019), the participants were
227 offered a calibrated breakfast at 08:00am, prepared according to the recommendations for their
228 age (≈ 500 kcal) (Murphy and Poos 2002). *Ad libitum* buffet-type meals were served at lunch

229 (12:00h) and dinner (19:00h). The adolescent's food preferences and eating habits were
230 considered for these buffets. Disliked and preferred items, as well as items liked but not usually
231 consumed were excluded to avoid over-, under- and occasional/opportunistic intake (Thivel et
232 al. 2016). The lunch buffet included: beef steaks, pasta, mustard, yogurt, cheese, fruits as well
233 as apple sauce and bread. For dinner, the adolescents were offered: ham/turkey, mashed
234 potatoes, beans, yogurt, cheese, apple sauce, fruits and bread. Each item was offered in
235 abundance and water was freely available. Participants were instructed to eat until feeling
236 comfortably satiated and extra food was available if needed. Food items were weighed by the
237 investigators before and after consumption, and the macronutrient composition (proportion of
238 carbohydrate, protein and fat) and total EI (kcal) were obtained using the Bilnut 4.0 software.
239 Breakfast, lunch and dinner meals were summed to obtain total daily energy intake. The
240 participants were asked not to engage in moderate to vigorous physical activities between meals
241 and mainly engaged in sedentary activities not involving screens, such as reading, homework,
242 or board games.

243

244 **Appetite sensations**

245 Visual analogue scales (VAS; 150-mm) were used to assess subjective appetite sensations
246 (hunger, fullness, desire to eat and prospective food consumption) at regular interval during the
247 day: before and after breakfast, lunch and dinner, and 30 and 60 minutes after breakfast and
248 lunch. The questions were: i) "How hungry do you feel?", ii) "How full do you feel?", iii)
249 "Would you like to eat something?", iv) "How much do you think you can eat?" (using French
250 translations of these questions). The participants were asked to rate their answer on a 150-mm
251 scale, from "not at all" to "a lot". This method has been previously validated (Flint et al. 2000).

252

253 **Food reward**

254 The Leeds Food Preference Questionnaire (LFPQ) was used to measure food preference and
255 food reward (Dalton and Finlayson 2014). The adolescents completed the LFPQ in a rested
256 **fasted state prior to the lunch meal**. They were presented with series of pictures of individual
257 food items common in the diet. The pictures were selected from a validated database to be
258 predominantly high (>50% energy) or low (<20% energy) in fat while showing similar
259 familiarity, protein content, palatability and being suitable for the population under study.
260 Explicit liking and wanting were measured by participants rating the extent to which they like
261 each item (“How pleasant would it be to taste this food now?”) and want each item (“How much
262 do you want to eat this food now?”). The pictures were presented individually, in a randomized
263 order and the participants were asked to rate them on a 100-mm VAS. Implicit wanting and
264 relative food preference were assessed using a forced choice task in which the food pictures
265 were paired so that every image from each of the four food types was compared to every other
266 type over 96 trials (food pairs). The adolescents were asked to respond **as fast and as precisely**
267 as possible to indicate the food they want to eat the most at that time (“Which food do you most
268 want to eat now?”). Reaction times (milliseconds) for all responses were covertly recorded and
269 used to compute mean response times for each food type after adjusting for frequency of
270 selection, in order to evaluate implicit wanting. To measure food choice as a marker of food
271 preference, the mean frequency of selection for each food type was recorded. The responses
272 obtained from the LFPQ were used to calculate mean scores for high-fat, low-fat, sweet or
273 savory food types (and different fat-taste combinations). Fat bias scores were calculated as the
274 difference between the high-fat scores and the low-fat scores, with positive values indicating
275 greater liking, wanting or preference for high-fat relative to low-fat foods and negative values
276 indicating greater liking, wanting or preference for low-fat relative to high-fat foods. Sweet bias
277 scores were calculated as the difference between the sweet and savory scores, with positive
278 values indicating greater liking, wanting or preference for sweet relative to savory foods and

279 negative values indicating greater liking, wanting or preference for savory relative to sweet
280 foods. The LFPQ has been used widely in adults (Dalton and Finlayson 2014) and youth
281 (Miguet et al. 2018). A French version of this LFPQ has been used, as previously reported
282 (Miguet et al. 2018).

283

284 **Exercise training**

285 **Phase 1.** As previously described by Miguet and colleagues (Miguet, et al. 2019), the first 12
286 weeks of the inpatient multidisciplinary intervention combined physical activity, nutritional
287 education and psychological support. The adolescents were asked to perform four 60-min
288 physical activity sessions per week including aerobic exercise, strength training, aquatic and
289 leisure-time activities (e.g. handball). They also received two hours of physical education per
290 week as part of their classical school curriculum. Nutritional education classes were organized
291 by a dietitian twice per month (one hour each time) and the adolescents received individualized
292 psychological support once per month (one hour). A diet respecting the adolescents' energy
293 requirements, based on age and sex recommendation, was prescribed to the adolescents
294 (Murphy and Poos 2002).

295 **Phase 2.** Details regarding this second phase of the intervention have been previously published
296 (Julian, et al. 2019). The adolescents were randomly assigned to a 12-week ECC or CON
297 exercise intervention during Phase 2. Both ECC and CON interventions were composed of three
298 ergometer exercise sessions per week, for a total of 36 sessions. The training program started
299 with two weeks of acclimatization where the intensity of the exercise and duration of the
300 sessions were progressively increased in order to protect subjects from delayed onset muscle
301 soreness (DOMS). Importantly, the CON and ECC training sessions were calibrated to induce
302 the same energy expenditure. ECC training was conducted using commercial ECC motor-
303 driven cycling ergometers (Cyclus2 Eccentric Recumbent, RBM elektronik-automation, MSE

304 Medical, France) and CON training was carried out on CON cycling ergometers (Optibike Med
305 600, MSE Medical, France). In addition to these specific sessions, the adolescents maintained
306 their two hours of physical education per week as well as a session dedicated to collective
307 games. Participants in both groups received the same dietary counseling throughout the
308 interventional period, although no guidelines for nutritional energy restriction were given in
309 order to investigate the sole impact of the training program. For more details regarding the
310 protocol used in Phase 2, please see Julian and colleagues (Julian et al. 2019).

311

312 **Statistical analysis**

313 Stata Software version 13 (StataCorp, College Station, TX, US) was used for statistical
314 analyses. The tests were two-sided, with a type I error set at 5%. Continuous data were
315 expressed as mean \pm standard deviation (SD) or median [interquartile range] according to
316 statistical distribution. Normality was checked using the Shapiro-Wilk test. Random-effects
317 models for correlated data were used to assess time effects (T0, T1 and T2) and group effects
318 (CON and ECC), and time by group interaction, considering between and within subject
319 variability (subject as random-effect). The normality of residuals from these models was
320 examined using the Shapiro-Wilk test. Logarithmic transformation were used if needed to
321 achieve the normality of dependent outcomes. Then, multivariable analyses were conducted
322 adjusting for i) age, ii) sex, iii) fat mass or BMI changes between baseline and T1 to compare
323 changes in the ECC and CON groups independently of the preceding modifications in fat mass
324 or body mass. Similar multivariable analyses based on models for repeated measures were
325 conducted to compare appetite sensations (daily hunger, prospective food consumption, satiety
326 and desire to eat) between groups across the time. Moreover, concerning the comparisons of
327 baseline values, quantitative variables were compared between groups using Student t-tests or
328 Mann-Whitney tests when assumptions of t-test were not met. The homoscedasticity was

329 analyzed using the Fisher-Snedecor's test. Categorical parameters were compared between
330 groups using chi-squared or Fisher's exact tests. The relationships between quantitative
331 variables were examined via correlation coefficients (Pearson or Spearman, according to
332 statistical distribution) applying Sidak's type I error correction due to multiple comparisons.

333

334 **Results**

335 Of the 24 adolescents who enrolled in the study (12 per group), 10 completed the whole protocol
336 in the ECC group (5 girls and 5 boys) and 11 in the CON group (6 girls and 5 boys). Adolescents
337 left the inpatient program for family and academic reasons, but none dropped out for reasons
338 associated with the program itself. Based on the DEBQ, 26.1% of the whole sample were
339 restrained eaters at baseline against 73.9% unrestrained, 47.8% at T1 (vs. 52.2% unrestrained)
340 and 52.0% at T2 (vs. 48.0% unrestrained).

341

342 As detailed in Table 1, body mass significantly decreased in both groups ($p < 0.001$) with
343 significant time x group effects at T1 ($p = 0.027$) and T2 ($p = 0.009$). Although the absolute body
344 mass changes between T1 and T2 (phase 2) were not significantly different between CON (-
345 4.2%) and ECC (-4.5%; $p = 0.74$), it was significantly different when adjusted on the T0-T1 body
346 mass changes ($p = 0.04$). BMI significantly decreased in both groups during phase 1 and phase
347 2 ($p < 0.001$) with no differences between groups. FM (%) significantly decreased in both groups
348 in phase 1 and phase 2 ($p < 0.001$). Although no group effect nor interaction were observed,
349 change in FM (%) during phase 2 was significantly higher in the ECC group (-9.9%) compared
350 with CON (-6.7%) when controlling for the change observed during phase 1 ($p < 0.004$). FFM
351 significantly increased between T0 and T1 ($p = 0.002$) in the CON group but remained
352 statistically unchanged between T1 and T2, and when considering the whole intervention

353 (between T0 and T2). No changes in FFM were observed in ECC and no group effect was
354 observed.

355

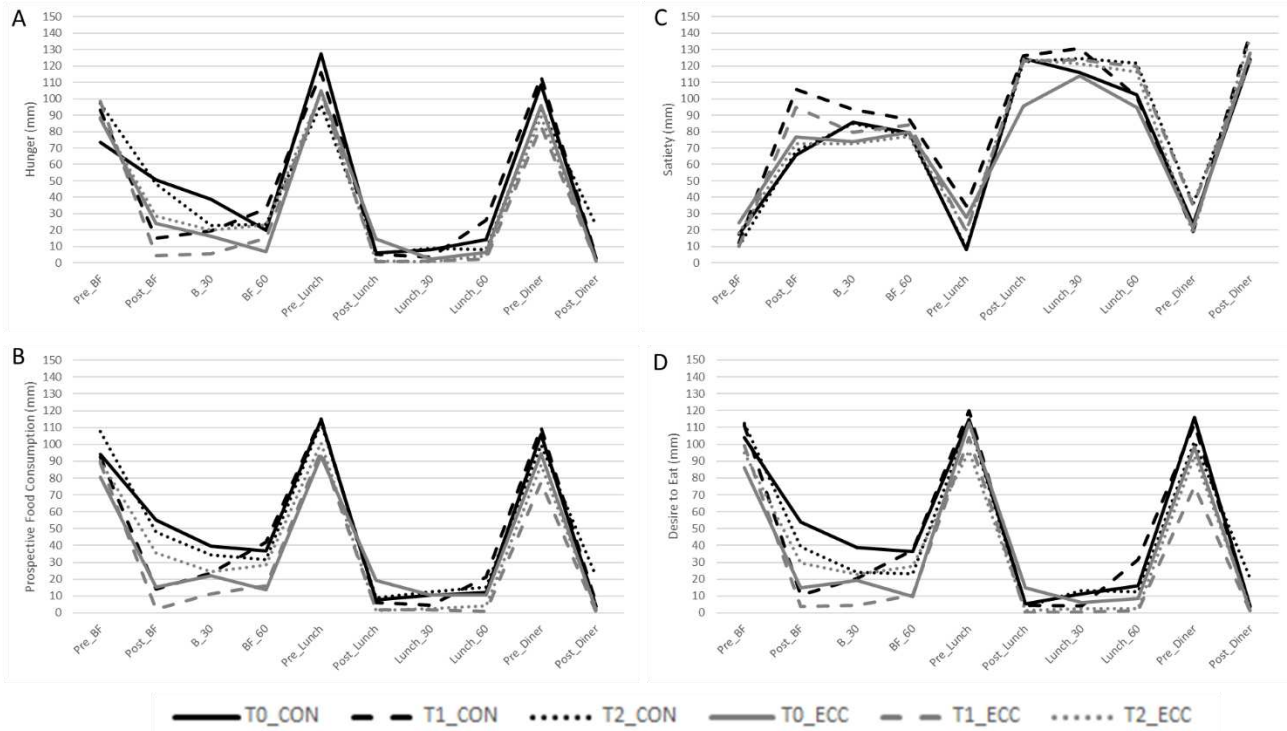
356 Table 2 presents the results pertaining to energy and macronutrient intake. Total daily EI
357 significantly increased in both groups during the whole intervention (CON: $p < 0.001$; ECC:
358 $p = 0.0136$) with no group effect nor interaction. EI did not significantly change in both groups
359 between T0 and T1, but significantly increased during phase 2 in the CON group ($p < 0.001$,
360 $+22\%$) but not in the ECC group ($p = 0.13$; $+8\%$). Table 2 also details the results regarding the
361 absolute and relative (percentage of total energy ingested) intake of each macronutrient.

362 There was no correlation between the changes in body mass, FM (%), FFM and EI between T0
363 and T2, T0 and T1, and T1 and T2. No differences were found between initially restrained and
364 unrestrained eaters in terms of body mass, FM (%) and EI changes between T0 and T1. FFM
365 increased significantly more in the initially restrained adolescents between T0 and T2 ($+3.3\%$
366 vs. -0.25% , $p = 0.0032$) and between T0 and T1 ($+3.6\%$ vs. $+0.84\%$; $p = 0.0075$). Body mass, FM
367 (%), FFM and EI changes during phase 2 were not significantly different between adolescents
368 categorized as restrained or unrestrained at T1.

369

370 Figure 2 illustrates the results for appetite sensations. While fasting hunger and satiety were not
371 different between groups at T0, desire to eat and prospective food consumption were
372 significantly higher in the CON group (17 ± 3 and 17 ± 7 mm, respectively) compared with the
373 ECC group (10 ± 5 and 10 ± 5 mm; $p = 0.04$ and $p = 0.02$, respectively). A group effect was found
374 for overall hunger and desire to eat ($p = 0.002$ and $p = 0.001$, respectively) with CON having
375 higher appetite ratings than ECC. There were no time effects nor interactions. A significant
376 time effect was found for prospective food consumption in both CON ($p = 0.03$; T0 vs. T1

377 p=0.03; T1 vs. T2 p=0.02) and ECC (p=0.04; T1 vs. T2 p=0.001) with no group effects nor
 378 interactions. Satiety was not different between groups or time points.
 379



380
 381 **Figure 2.** Changes in appetite sensations throughout the assessment day during the
 382 intervention: Hunger (A); Prospective food consumption (B); Satiety (C); and Desire to eat
 383 (D). Pre-BF, Pre-breakfast; Post-BF, Post-breakfast; B_30, 30 minutes post-breakfast; B_60, 60 minutes post-
 384 breakfast; Lunch-30, 30 minutes post-lunch; Lunch-60, 60 minutes post-lunch; T0_CON, Concentric group at
 385 baseline; T0_ECC, Eccentric group at baseline; T1_CON, Concentric group at the end of phase 1; T1_ECC, Eccentric
 386 group at the end of phase 1; T2_CON, Concentric group at the end of phase 2; T2_ECC, Eccentric group at the end of
 387 phase 2
 388
 389

390 The LFPQ results can be found in Table 3. Relative fat preference, implicit wanting for fat,
 391 explicit wanting for sweet and explicit liking for fat and sweet were not different between
 392 groups and did not show any time or interaction effects. While none of the LFPQ dimensions
 393 showed a time effect in the CON group, time effects were observed in the ECC group. In ECC,

394 preference for high-fat relative to low-fat foods increased at T2 relative to T0 ($p=0.02$) and T1
395 ($p=0.02$). Preference for sweet relative to savory foods decreased at T1 ($p=0.04$) and T2
396 ($p=0.001$) relative to T0 (i.e. negative values indicate higher preference for savory vs. sweet
397 foods, thus preference for savory foods increased). Implicit wanting for sweet relative to savory
398 foods decreased at T1 ($p=0.02$) and T2 ($p=0.008$) relative to T0 (i.e. negative values indicate
399 greater wanting for savory vs. sweet foods). Group effects were observed for the relative
400 preference for fat ($p=0.058$) and explicit wanting for fat ($p=0.009$), with CON having greater
401 preference and explicit wanting for high-fat foods than ECC. Group x time interactions were
402 observed at T1 and T2 for the preference for sweet food ($p=0.007$ and $p=0.002$, respectively),
403 with CON increasing and ECC decreasing. There was also a group x time interaction for the
404 implicit wanting for sweet food ($p=0.002$) at T2, with CON increasing and ECC decreasing.

405

406 **Discussion**

407 While there is a constant need to improve our weight loss strategies, the present study compared
408 the effect of a 12-week eccentric *versus* concentric cycling intervention on energy intake,
409 appetite, and food reward in adolescents with obesity. According to our results, 12 weeks of
410 eccentric cycling as part of a multidisciplinary intervention might prevent adolescents from the
411 increased energy intake observed in response to similar concentric training matched for
412 metabolic load.

413 The TEXTOO study is, to our knowledge, the first study to compare such eccentric
414 *versus* concentric cycling interventions of similar metabolic load in adolescents with obesity
415 (Julian et al. 2019). As previously reported, (Julian et al. 2019), significant body mass, BMI
416 and fat mass percentage decreases were observed in both groups during both phase 1 (classical
417 multidisciplinary program) and phase 2 (ECC vs. CON), with a greater body mass reduction in
418 the ECC group. This is of particular importance since body mass and fat mass losses have been

419 shown to be attenuated during the second phase of multidisciplinary interventions that mainly
420 use concentric exercise, in similar populations (Julian et al. 2019; Miguet et al. 2019; Carnier
421 et al. 2013). In their recent study, Miguet and colleagues observed minor body mass and body
422 composition improvements during the second half of a 10-month multidisciplinary weight loss
423 program in adolescents with obesity (Miguet et al. 2019). Interestingly, this was accompanied
424 by a significant increase in EI, particularly in restrained adolescents (Miguet et al. 2019).
425 According to the present results, *ad libitum* daily EI was increased in both groups when
426 considering the entire intervention; however, a significant increase in only the CON group was
427 found during phase 2 (+22% increase from T1) whereas it did not change significantly in the
428 ECC group. Similarly, while protein and fat intake increased in both groups during the
429 intervention (Phase 1 and 2), it did not change significantly in the ECC group during phase 2.
430 ECC training seems to have prevented the adolescents from increasing their food consumption
431 during the second phase of the current weight loss intervention.

432 Since the control of EI has been shown to be associated with FM and FFM (Blundell et
433 al. 2003; Thivel et al. 2019), it could be suggested that the observed EI differences between
434 ECC and CON are explained by the differential effect of the two interventions on body
435 composition. However, no correlations were found between the changes in body mass, FM (%)
436 and FFM and the EI changes in phase 1 and phase 2, despite slightly higher body mass and
437 FFM at baseline in ECC compared with CON (marginally significant at $p=0.09$ and $p=0.08$,
438 respectively), that were not maintained at T1 and T2. While this absence of association between
439 body composition and EI in the present study may be due to the low sample size, other factors
440 related to the hedonic and neurocognitive control of EI could explain our results (Thivel et al.
441 2019). The role of the level of cognitive restriction has been recently proposed to explain the
442 appetite and EI responses to weight loss in adolescents with obesity (Miguet et al. 2019). Sim
443 et al. have for instance described that physical activity can lead to counterproductive responses

444 in restrained eaters by favoring increased food intake (Sim et al. 2018). More recently,
445 cognitively restrained adolescents with obesity were found to have a greater increase in EI in
446 response to a 10-month weight loss intervention based on concentric exercise, compared with
447 unrestrained eaters (Miguet et al. 2019). Although we did not find EI differences between
448 restrained and unrestrained adolescents when considering our whole sample, 66.6% of the
449 adolescents composing the CON sample were found restrained at T2 against only 27.2% in the
450 ECC sample, which might explain the observed lower energy intake in this latter group.
451 However, this must be interpreted with caution due to the relatively small size of our sample.

452 In addition to the level of cognitive restriction, food reward has been identified as a key
453 determinant of the control of energy intake (Finlayson and Dalton 2012; Thivel et al. 2019).
454 We found only one study that examined the effect of exercise on food reward in children and
455 adolescents with obesity, suggesting that an acute 16-minute bout of high intensity interval
456 exercise might favor reduced reward for fat and sweet foods compared with a resting condition
457 in 12 to 15 year-old adolescents with obesity (Miguet et al. 2018). In adults with obesity,
458 Martins and collaborators compared the food reward responses to 12 weeks of isocaloric high-
459 intensity interval training *versus* moderate intensity continuous training, not observing any
460 changes over time in both groups (Martins et al. 2017). The present study is to our knowledge
461 the first to investigate the effect of chronic exercise on food reward in adolescents with obesity.
462 Interestingly, our results did not demonstrate a clear effect of exercise per se on the dimensions
463 of food reward. However, CON showed a greater explicit wanting for high-fat foods compared
464 to ECC; and CON experienced an increase in implicit wanting and preference for sweet food
465 during training, while ECC tended towards savoury food during training. These differences in
466 food reward overall and in response to the exercise interventions may explain the greater EI
467 observed in CON during the second phase of the intervention. Preference for high-fat foods
468 may have led to more energy dense food selection during the test meal buffet. The differences

469 in taste preference in ECC and CON are more difficult to interpret as the test foods in the buffet
470 were predominantly savoury, and savoury foods are typically found in the form of entrees, sides
471 and salty snacks that comprise a large proportion of total energy in the diet. An increase in
472 reward for sweet-fat foods has previously been shown to explain overconsumption in women
473 (Dalton and Finlayson 2014), however added salt in savory foods has also been shown to
474 increase energy intake independent of fat content (Bolhuis et al. 2016). Future research should
475 employ test meal designs that include a greater range of sweet and savoury foods varying in
476 fat/energy density.

477 It is also important to acknowledge that potential underlying group differences that may
478 have contributed to the effects found on EI. Although hunger and desire to eat were significantly
479 higher in CON (group effect), no time effect nor interaction were found, minimizing their
480 potential role in the EI difference observed between groups. This unlikely effect of subjective
481 appetite on the EI difference between CON and ECC is reinforced by the lack of group and
482 time effects in satiety and the increase in prospective food consumption from T0 to T2 in both
483 groups. This confirms once more the potentially uncoupling effect of physical exercise on
484 energy intake and appetite sensations that has been previously described (Thivel and Chaput
485 2014). Furthermore, we found that savory foods within the LFPQ food stimuli were preferred
486 over sweet foods at all time points during the intervention (negative sweet bias values). Whether
487 this is a characteristic of the sample or due to the food stimuli used within the task is unknown,
488 as well as whether this could have influenced our outcomes. However, it should be noted that
489 the task was performed in the hungry state prior to a meal, when preferences for savory foods
490 are typically greater than sweet foods (Finlayson et al. 2008). Future studies using food images
491 that have been culturally adapted for the French population may shed some light on this
492 observation (Oustric et al. 2019).

493 The present results must be interpreted in light of some limitations. First, although the
494 TEXTOO study was sufficiently powered for the analyses of the energy intake outcomes, this
495 is a secondary analysis of a larger study which aimed to assess the effect of ECC versus CON
496 training on muscle strength in adolescents with obesity. Similarly, despite our statistical
497 analyses being adjusted for the sex of the adolescents, and Julian et al. previously reporting
498 similar adherence rates and perceived exertion between boys and girls and between groups
499 (ECC vs. CON) (Julian et al. 2019), larger studies should be conducted to better examine any
500 differences in the responses to ECC cycling training between boys and girls. Moreover, we used
501 *ad libitum* buffet meals as an objective measure of energy intake instead of self-reported diaries.
502 Although self-reported diaries have been shown to underestimate food intake, particularly in
503 such a population (Burrows et al. 2010), placing the adolescents in a context of food abundance
504 might not represent their normal eating behaviors, especially during a weight loss intervention.
505 Moreover, while the adolescents were used to eating a mid-afternoon snack during their
506 inpatient program, they were not allowed to eat in between meals during our nutritional
507 evaluation days, which might have slightly affected our results. It would have also been of
508 interest to conduct follow-up assessments to analyze the potential long-term responses, but this
509 was not possible for practical reasons.

510 To conclude, the present study suggests that the appetite and eating behavior responses
511 of adolescents with obesity to an inpatient multidisciplinary weight loss intervention might be
512 different depending on the modality of the exercise used. Introducing eccentric cycling during
513 a multidisciplinary weight loss intervention, previously shown to have a greater effect on body
514 composition over a classical concentric exercise training, might help prevent increased *ad*
515 *libitum* energy intake observed in response to the concentric intervention in these adolescents;
516 potentially through distinct effects of the food reward system. Further larger longitudinal

517 studies are required to confirm these results and consider the potential inter-individual
518 variability.

519

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Table 1. Anthropometric characteristics and body composition changes during the intervention in CON and ECC

	T0		T1		T2		Time effect	T0 vs. T1	T0 vs. T2	T1 vs. T2	Group effect	Interaction TxG	
	Mean	SD	Mean	SD	Mean	SD						T1	T2
<i>Body mass (kg)</i>													
CON	81.3	13.3	75.5	11.6	72.3	11.8	<0.0001	<0.0001	<0.0001	0.0038	0.145	0.027	0.009
ECC	93.9	20.3	82.8	17.4	78.8	16.0	<0.0001	<0.0001	<0.0001	0.0129			
<i>p</i>	0.09		0.32		0.22								
<i>BMI (kg/m²)</i>													
CON	31.8	3.8	29.4	3.6	27.6	4.0	<0.0001	<0.0001	<0.0001	<0.0001	0.305	0.147	0.144
ECC	34.8	5.5	30.8	4.9	29.0	4.5	<0.0001	<0.0001	<0.0001	0.0005			
<i>p</i>	0.16		0.54		0.53								
<i>Fat mass (%)</i>													
CON	38.0	4.7	31.8	4.7	29.8	6.3	<0.0001	<0.0001	<0.0001	0.0051	0.43	0.54	0.19
ECC	38.0	5.0	30.3	5.1	27.5	6.6	<0.0001	<0.0001	<0.0001	0.0031			
<i>p</i>	0.89		0.35		0.32								
<i>Fat-free mass (kg)</i>													
CON	48.5	9.0	49.5	8.4	48.6	8.4	0.0088	0.002	0.086	0.14	0.085	0.23	0.40
ECC	55.4	9.9	55.2	10.3	54.6	10.6	0.37	0.17	0.59	0.38			
<i>p</i>	0.08		0.12		0.13								

Data presented as mean ± standard deviation; T0, baseline; T1, end of phase 1; T2, end of phase 2; Interaction TxG, Interaction Time x Group; CON, Concentric group; ECC, Eccentric group; SD, Standard deviation.

Table 2. Changes in energy and macronutrient intake during Phase 1 and Phase 2 in CON and ECC

		T0		T1		T2		Time effect	T0 vs. T1	T0 vs. T2	T1 vs. T2	Group effect	Interaction TxG	
		Mean	SD	Mean	SD	Mean	SD						T1	T2
Energy intake (kcal)														
	CON	1856	253	1936	309	2396	431	<0.0001	0.49	<0.0001	0.0004	0.72	0.76	0.16
	ECC	1932	539	2080	515	2232	522	0.0136	0.15	0.003	0.13			
	<i>p</i>	0.8		0.45		0.78								
Protein (g)														
	CON	96.8	18.2	108.0	15.8	130.9	25.5	<0.0001	0.086	<0.0001	0.0006	0.93	0.82	0.07
	ECC	101.3	29.4	113.5	33.3	116.8	30.4	0.0074	0.019	0.003	0.5235			
	<i>p</i>	0.88		0.42		0.39								
Fat (g)														
	CON	60.4	15.4	71.4	16.3	84.8	21.4	0.0016	0.088	<0.0001	0.0463	0.93	0.65	0.26
	ECC	60.2	20.5	76.0	21.9	76.9	22.0	0.0001	<0.0001	<0.0001	0.8284			
	<i>p</i>	0.62		0.57		0.11								
CHO (g)														
	CON	236.4	34.2	215.4	38.3	274.3	60.3	0.0017	0.189	0.034	0.0004	0.63	0.65	0.55
	ECC	244.4	67.9	234.0	59.8	263.7	68.4	0.1977	0.535	0.249	0.0761			
	<i>p</i>	0.72		0.41		0.58								
Protein (%)														
	CON	20.8	2.0	22.5	2.6	22.0	2.5	0.2187	0.082	0.304	0.4913	0.34	0.52	0.54
	ECC	21.0	1.7	21.7	2.9	21.0	2.7	0.5774	0.374	0.971	0.3548			
	<i>p</i>	0.98		0.88		0.41								
Fat (%)														
	CON	29.2	5.4	33.1	4.7	31.8	5.1	0.0426	0.014	0.073	0.5440	0.80	0.80	0.95
	ECC	28.1	3.6	32.9	4.0	31.3	5.7	0.0003	<0.0001	0.008	0.1819			
	<i>p</i>	0.83		0.66		0.06								
CHO (%)														
	CON	51.0	3.9	44.8	6.1	45.8	5.6	<0.0001	<0.0001	<0.0001	0.6733	0.80	0.60	0.34
	ECC	50.6	4.5	45.1	4.2	46.9	5.5	0.0052	0.001	0.033	0.2940			
	<i>p</i>	0.77		0.51		0.06								

Data presented as mean ± standard deviations; T0, Baseline; T1, end of phase 1; T2, end of phase 2; Interaction TxG, Interaction Time x Group; CON, Concentric group; ECC, Eccentric group; SD, Standard deviation.

Table 3. Changes in relative preference, implicit wanting, explicit wanting and explicit liking for high vs. low fat foods (fat bias) and sweet vs. savory foods (sweet bias) in CON and ECC

	T0		T1		T2		Time effect	T0 vs. T1	T0 vs. T2	T1 vs. T2	Group effect	Interaction TxG	
	Mean	SD	Mean	SD	Mean	SD						T1	T2
Relative preference fat bias (no. of choices)													
<i>CON</i>	6.8	5.5	11.2	6.4	9.7	8.2	0.24	0.10	0.33	0.49	0.058	0.23	0.44
<i>ECC</i>	3.2	9.3	3.2	7.2	7.1	6.3	0.03	0.96	0.02	0.02			
Relative preference sweet bias (no. of choices)													
<i>CON</i>	-18.4	6.7	-13.9	7.2	-14.9	7.1	0.17	0.06	0.23	0.51	0.87	0.007	0.002
<i>ECC</i>	-10.8	10.3	-15.0	8.6	-17.9	6.7	0.0049	0.04	0.001	0.2			
Implicit wanting fat bias (a.u.)													
<i>CON</i>	19.6	15.6	23.9	19.6	13.0	32.3	0.35	0.71	0.33	0.16	0.36	0.95	0.06
<i>ECC</i>	9.1	24.7	11.5	18.1	17.0	14.4	0.15	0.65	0.06	0.15			
Implicit wanting sweet bias (a.u.)													
<i>CON</i>	-48.9	22.3	-41.9	23.0	-31.8	30.7	0.18	0.5	0.07	0.22	0.92	0.07	0.002
<i>ECC</i>	-31.2	28.3	-43.3	18.6	-47.0	20.0	0.016	0.02	0.008	0.60			
Explicit wanting fat bias (mm)													
<i>CON</i>	10.6	7.1	7.5	8.2	10.8	12.1	0.55	0.38	0.95	0.33	0.009	0.5	0.67
<i>ECC</i>	3.0	14.7	3.7	10.1	0.6	4.1	0.76	0.87	0.58	0.48			
Explicit wanting sweet bias (mm)													
<i>CON</i>	-17.6	15.0	-18.4	16.6	-15.1	14.5	0.89	0.80	0.84	0.64	0.38	0.25	0.22
<i>ECC</i>	-19.1	22.0	-10.5	20.3	-7.3	11.1	0.17	0.18	0.07	0.59			
Explicit liking fat bias (mm)													
<i>CON</i>	11.4	9.1	9.8	13.3	8.0	12.1	0.81	0.70	0.51	0.77	0.14	0.47	0.91
<i>ECC</i>	4.4	15.3	6.9	11.0	0.8	4.6	0.33	0.52	0.39	0.13			
Explicit liking sweet bias (mm)													
<i>CON</i>	-20.8	13.4	-18.1	16.9	-13.3	11.7	0.41	0.66	0.19	0.36	0.38	0.85	0.86

ECC -17.3 20.6 -13.4 14.1 -9.2 13.6 0.36 0.50 0.15 0.43

Data presented as mean \pm standard deviation; T0, Baseline; T1, end of phase 1; T2, end of phase 2; Interaction TxG, Interaction Time x Group; CON, Concentric group; ECC, Eccentric group; SD, Standard deviation; a.u., arbitrary unit. Note: Positive values indicate greater liking, wanting or relative preference for high-fat relative to low-fat foods (fat bias) or sweet relative to savory foods (sweet bias).