



Cronfa - Swansea University Open Access Repository

This is an author produced version of a paper published in: Journal of Physical Activity and Health

Cronfa URL for this paper: http://cronfa.swan.ac.uk/Record/cronfa34944

Paper:

Mackintosh, K., Ridgers, N., Evans, R. & McNarry, M. (2017). Physical Activity and Sedentary Time Patterns in Children and Adolescents with Cystic Fibrosis and Age- and Sex-Matched Healthy Controls. *Journal of Physical Activity and Health*, 1-24.

http://dx.doi.org/10.1123/jpah.2017-0011

This item is brought to you by Swansea University. Any person downloading material is agreeing to abide by the terms of the repository licence. Copies of full text items may be used or reproduced in any format or medium, without prior permission for personal research or study, educational or non-commercial purposes only. The copyright for any work remains with the original author unless otherwise specified. The full-text must not be sold in any format or medium without the formal permission of the copyright holder.

Permission for multiple reproductions should be obtained from the original author.

Authors are personally responsible for adhering to copyright and publisher restrictions when uploading content to the repository.

http://www.swansea.ac.uk/iss/researchsupport/cronfa-support/

1 Abstract

2 **Background:** Regular physical activity (PA) is increasingly recognised as important in the care of patients with Cystic Fibrosis (CF) but there is a dearth of evidence regarding physical 3 activity levels (PAL) or how these are accrued in those with CF. *Methods:* Physical activity 4 5 was measured by a hip-worn accelerometer for seven consecutive days by eighteen children 6 (10 boys: 12.4 ± 2.8 years) with mild to moderate CF and eighteen age- and sex-matched 7 controls (10 boys; 12.5 ± 2.7 years). **Results:** Both CF and healthy children demonstrated 8 similar PAL and patterns of accumulation across the intensity spectrum, with higher levels of PA during weekdays in both groups. FEV₁ was predicted by high-light PA in CF compared to 9 low-light PA in healthy children. *Conclusion:* These findings highlight weekends and light 10 PA as areas warranting further research for the development of effective intervention 11 strategies to increase PA in the youth CF population. 12

13 Introduction

Cystic Fibrosis (CF) is the most prevalent lethal autosomal recessive disease in the Caucasian 14 population.¹ Mutations in the cystic fibrosis transmembrane regulator (CFTR) gene lead to 15 16 malfunctioning or absent CFTR proteins, impairing mucosal clearance mechanisms. As such, CF is characterised by excessive viscous secretions in almost all organs, particularly the 17 lungs, resulting in recurring infections, inflammation, airflow obstruction, and ultimately 18 progressive functional decline. Whilst there remains no cure, advances in the treatment for 19 patients with CF have resulted in an increased median life expectancy from 8 years in 1974, 20 to 31 years in 2005 and 41 years in 2012.² 21

22

Whilst physical activity (PA) has been associated with numerous physiological and 23 psychosocial benefits for healthy children,³ there are additional health benefits for patients 24 with CF. These include slower lung function decline,⁴ reduced hospital admissions,⁵ 25 improved quality of life and nutritional status,⁶ improved bone mineral density,⁷ and 26 enhanced airway clearance⁸ and ion channel function, which could lead to improved mucus 27 hydration and clearance.⁹ PA could be imperative for ultimate survival in patients with severe 28 lung deterioration, given the strong positive relationship with aerobic capacity.^{10,11} However, 29 recent research suggests that as little as 2.1% of children with CF¹² meet the government PA 30 guidelines of at least 60 minutes of moderate-to-vigorous-intensity physical activity (MVPA) 31 every day.¹³ Whilst regular PA is increasingly important in the care of patients with CF,¹⁴ 32 there is a dearth of research and indeed little consensus on physical activity levels (PAL) in 33 children and adolescents with CF. Moreover, little is known as to whether beneficial 34 outcomes may be achieved with engagement of PA at different intensities, which would be 35 critical information for interventions and on going care. 36

37

38 Variations in PAL reported in the existing literature may be due to methodological inconsistencies. Earlier research employing self-reported measures found that children aged 39 7-17 years with CF participated in less very strenuous (> 6 METs) PA relative to healthy 40 controls, even when patients had well-preserved lung function.¹⁰ Conversely, Selvadurai and 41 colleagues⁶ reported no significant differences between CF patients and age- and sex-42 43 matched controls in similarly-aged children (9-17 years), using uniaxial accelerometry. Advancing previous research, which only reported total counts⁶, Aznar et al.¹² utilised 44 Evenson¹⁵ cut-points to find that 6-17 year old children with CF engaged in significantly less 45 MVPA and vigorous physical activity (VPA) but demonstrated higher total and light physical 46 activity (TPA and LPA, respectively). Yet Jantzen et al.¹⁶ found similar PAL in CF patients 47 across the age and intensity spectrum, but less engagement in strenuous activities for school-48 aged children (6-13 years) compared to healthy controls. Interestingly, when extreme values 49 were removed, no relationship was present between strenuous PA and percentage predicted 50 FEV₁.¹⁶ 51

52

A potential limitation of earlier studies is the lack of age- and population-specific cut-points, 53 although, in healthy populations, it is pertinent to note that Trost and colleagues¹⁷ supported 54 the use of Evenson cut-points. Arguably, the relative intensity for children and adolescents 55 with CF could be greater and therefore light physical activity may be more beneficial to their 56 health in comparison to their healthy counterparts. However, with the exception of Aznar and 57 colleagues,¹² and to some extent Jantzen et al.,¹⁶ the majority of studies did not consider PA 58 across the spectrum.^{6,10,18} As such, the identification of patients with CF participating in more 59 LPA and TPA¹² may warrant further investigation. Specifically, previous research has 60 suggested that time spent in low-light physical activity (low-LPA) and high-light physical 61 activity (high-LPA) may have some favourable independent health benefits.¹⁹ Additionally, a 62

63 sedentary lifestyle has been shown to contribute to the progression of both functional and physical impairment in CF populations,²⁰ yet little research has objectively assessed time 64 spent being sedentary, nor the accumulation of PA or sedentary time. Indeed, the majority of 65 66 physical activity research to date has focused on the total volume of PA rather than the manner in which this activity is accumulated with regards to bout frequency and duration. 67 Gabel et al.²¹ reported sedentary bouts of >5 minutes to be detrimentally associated with C-68 reactive protein in healthy children, whereas PA bouts of ≥ 1 minute, which have previously 69 been used to identify sporadic bouts of PA, are reported to be associated with lower BMI.²² 70 Identifying patterns of accumulation in youth with CF, and those patterns that may be 71 associated with functional gains, is important for advancing the design and evaluation of 72 future interventions in this population. 73

74

75 In order for effective interventions aimed at improving PAL in children and adolescents to be developed, it is important to further understand current levels, intensities and accumulation of 76 77 PA children and adolescents with and without CF. Therefore, the purpose of this study was to 78 investigate PA and sedentary time patterns of children and adolescents with CF, in comparison to age- and sex-matched healthy controls. Furthermore, the study sought to 79 ascertain whether such parameters could predict lung function. We hypothesised that PA 80 intensity and duration would be significantly lower in patients with CF and be a significant 81 predictor of disease severity (i.e., lung function). 82

83

84 Methods

85 *Participants*

In total, 36 participants (12.6 ± 2.7 years; 18 CF) were invited to take part in the study. Descriptive characteristics are shown in Table 1. Eighteen patients (10 boys) with mild-to-

moderate CF, confirmed by a sweat chloride > 60 mmol·l⁻¹ and genotyping (8 Δ F508 88 Homozygote, 10 Δ F508 Heterzygote; 4 CF-related liver disease) were recruited from an 89 outpatient CF clinic in South Wales (United Kingdom). Patients were included in the study if 90 91 they were aged 6 - 17 years old, had no increase in symptoms or weight loss two weeks prior to testing, and had a stable lung function (within 10% of best in the preceding six months); 92 93 unstable non-pulmonary comorbidities or acute infections warranted exclusion. Eighteen ageand sex-matched non-clinical children were recruited from local schools to act as a healthy 94 comparison group. Ethical approval was granted by the Bromley NHS research ethics 95 96 committee (REC reference: 13/LO/1907) and written informed consent and assent were obtained from parents/guardians and patients, respectively. All patients were instructed to 97 continue prescribed medications as usual throughout the duration of their study involvement. 98

99

100 Measurements

At their routine visits to the clinic, participants forced vital capacity (FVC) and forced 101 102 expiratory volume in 1s (FEV1) were assessed using flow-volume loop spirometry (Vitalograph, UK)). The best of three consistent exhalations (<5% variability) was recorded. 103 All lung function measurements were expressed as a percentage predicted normal, using 104 appropriate reference data.²³ Furthermore, body mass (Seca 220; Hamburg, Germany), stature 105 and sitting stature (Seca 220; Hamburg, Germany) were measured to the nearest 0.01 kg and 106 107 0.01 m, respectively. Waist circumference was measured to the nearest 0.01 m using a non-108 elastic anthropometric tape (Seca Ltd., Birmingham, UK) at the narrowest point between the bottom of the ribs and the iliac crest. Healthy age- and sex-matched counterparts were asked 109 to attend one laboratory session at Swansea University for all measurements to be 110 undertaken. All participants were provided with a hip-mounted ActiGraph GT3X+ 111

accelerometer (ActiGraph LLC, Pensacola, FL) to assess habitual PAL over sevenconsecutive days.

114

ActiGraph monitors, shown to have acceptable reliability and validity in paediatric 115 populations,²⁴ sampled raw data at 100Hz. Data were downloaded using ActiLife software 116 (v6.10.4; ActiGraph, Pensacola, FL), processed into 15s epochs and reduced using a 117 customised Excel macro. Sustained periods of 20 minutes of consecutive zero's were used to 118 define non-wear time, which has been found to result in an almost identical wear time and a 119 smaller difference between sedentary time and sitting time estimates (assessed using 120 activPAL; PAL Technologies, Glasgow, Scotland) compared with a 60 minute definition in 121 children.²¹ Sedentary time was defined as $<100 \text{ counts} \cdot \text{min}^{-1}$, shown to be a good estimate of 122 free-living sitting.²⁵ Time spent in MPA (4-5.99 METs) and VPA (≥6 METs) was determined 123 using age-specific cut-points,²⁶ which demonstrated comparable accuracy to Evenson cut-124 points.¹⁷ A threshold of 4 METs was used to define MPA, as brisk walking has been 125 associated with this energy cost in calibration studies.^{27,28} MPA and VPA were summed to 126 create MVPA. The rest of the time was classified as either low light-intensity physical 127 activity (low-LPA; 100-799 counts min⁻¹) or high light-intensity physical activity (high-128 LPA; 800-<4 METs). The 800 counts min⁻¹ threshold was selected as this published 129 sedentary cut-point captures both sedentary time and static light-intensity activities such as 130 standing,²⁵ and has been found to have differential associations with cardiometabolic 131 biomarkers in adolescents.¹⁹ A valid day was defined as ≥ 9 hours day⁻¹, which has been 132 previously used in clinical populations.²⁹ To be included in the analyses, children were 133 required to have worn the ActiGraph for at least three days, which has been shown to have a 134 reliability coefficient of 0.7.³⁰ PAL are reported for overall, weekday and weekend days 135 separately. 136



Patterns of sedentary time and PA accumulation were also calculated. Breaks in sedentary time were defined as the number of times that the accelerometer exceeded 25 counts per 15s epoch following a 15s epoch of <25 counts per epoch.³¹ The frequency and duration of time spent in sedentary (\geq 5 min),²¹ and low-LPA, high-LPA, MPA and VPA (\geq 1 minutes) were also determined.²² No interruptions to these bouts were permitted.

143

144 Data Analysis

Gaussian distribution was confirmed by the Shapiro-Wilks test. Following this, the 145 participant groups and weekday vs. weekend day were compared using a multivariate 146 ANCOVA with group as a fixed factor and day as a repeated measure, controlling for wear 147 time. A stepwise linear regression was used to analyse the association between FEV₁ and PA 148 intensity levels and patterns, adjusting for predefined potential confounders (age, sex, stature, 149 mass and wear time). To explore differences between the groups in terms of those that met 150 151 current government guidelines for PA (i.e., average of ≥60 minutes of MVPA/day), a Chisquare test was used. All statistical analyses were conducted using PASW Statistics 21 152 (SPSS, Chicago, IL). All data are presented as means \pm SD. Statistical significance was 153 accepted when $P \leq 0.05$. 154

155

156 **Results**

157 No significant differences were observed between boys and girls with regards to 158 anthropometrics or lung function, with the exception of maturity offset, which was 159 significantly greater in boys (Table 1). Consequently, all data were pooled for subsequent 160 analyses. The healthy and CF groups did not differ in anthropometrical characteristics.

- However, those with CF presented with a significantly lower percentage of predicted FEV₁
 and FEV:FVC when described in both absolute and relative to predicted terms.
- 163

A total of four (2 healthy controls; 2 patients with CF) participants did not fulfil the wear time criteria for valid accelerometry data and were therefore excluded from further analyses. Those excluded did not differ in anthropometrics or lung function to those retained. Overall, participants achieved 4.5 ± 1.2 and 1.8 ± 0.6 valid weekdays and weekend days, respectively.

169 CF patients and healthy controls engaged in similar levels of PA across the intensity spectrum, irrespective of whether weekday, weekend day or overall days were considered 170 (Table 2). There was a trend for greater time spent in LPA in CF patients (222.7 \pm 12.8 vs. 171 207.3 ± 12.4 mins; P > 0.05), although this failed to reach significance. There were 172 significant differences between weekday and weekend day PA with regards to total LPA 173 $(229.3 \pm 52.4 \text{ vs. } 203.8 \pm 50.6 \text{ mins, respectively; } P < 0.05)$, MPA $(45.1 \pm 21.5 \text{ vs. } 36.6 \pm 1.5 \text{ vs. } 36.5 \pm 1.5 \text{ vs. } 36.5$ 174 27.9 mins, respectively; P < 0.05) and MVPA (62.4 ± 32.1 vs. 51.2 ± 39.9 mins, respectively; 175 P < 0.05), with greater levels of activity achieved during weekdays than weekend days in 176 both groups. 177

178

Overall, 44.4% (n=8) vs. 38.9% (n=7) in the healthy and CF groups, respectively, met the current guidelines for MVPA. Fewer children met the guidelines on weekend days (44.4% vs. 30.6%; *P* < 0.05). The percentage meeting government guidelines did not differ between CF and healthy children during week or weekend days.

183

Healthy controls and CF patients demonstrated similar patterns of physical activityaccumulation (Table 3). However, different patterns were evident during weekday and

- weekend days, with weekdays characterised by a greater frequency and duration of LPA andMPA bouts and a lower duration of sedentary bouts compared to weekend days.
- 188

Linear regression revealed that FEV₁ was predicted by height and LPA when both groups were pooled for analysis ($F_{(2,31)} = 62.93$, P < 0.001; $R^2 = 0.80$). More specifically, when LPA was split into low-LPA and high-LPA, height and low-LPA significantly predicted FEV₁ ($F_{(2,31)} = 68.07$, P < 0.001; $R^2 = 0.82$). When the groups were considered independently, the intensity of LPA that predicted FEV₁ differed, with FEV₁ predicted by height and high-LPA in CF patients ($F_{(2,14)} = 79.60$, P < 0.001; $R^2 = 0.92$) compared to height and low-LPA in healthy controls ($F_{(2,14)} = 24.31$, P < 0.001; $R^2 = 0.78$).

196

197 Discussion

198 Children with CF and age- and sex-matched healthy controls did not differ in overall PAL or 199 the pattern in which these levels were accrued. Interestingly, despite these similarities, FEV₁ 200 was dependent on LPA levels in both CF patients and their healthy counterparts, although the 201 intensity within LPA differed across the groups. Finally, we observed significant decreases in 202 PAL during weekends, with increased sedentary time and decreased frequency and duration 203 of LPA and MPA bouts, irrespective of disease status.

204

In agreement with some,^{6,18,32} but not all,^{10,12} previous studies, no significant difference was observed in the PAL of children with and without CF, although a considerably higher proportion of our CF population met recommended guidelines compared to previous research.¹² Given the numerous additional health benefits for patients with CF,⁴⁻⁹ over and above the physiological and psychosocial benefits of regular PA identified in healthy children,³ these findings highlight the need for strategies to increase PA in this population. Indeed, the importance of PA has been recognised by the European Cystic Fibrosis Society (ECFS) and recent Cochrane Reviews,^{14,33} which advocate the cost-effectiveness and beneficial effects of PA for promoting quality of life in patients with CF. However, information regarding PA behaviours in CF is limited and although PA as a treatment is becoming increasingly valued by CF clinical teams,³⁴ it remains underutilized in routine CF management ³⁵. Furthermore, there is a paucity of evidence-based guidance regarding the optimal combination of intensity and duration to elicit health benefits.

218

Further controversy surrounds the relationship between CF and the intensity of PA 219 undertaken, including with regards to the direction of causality. In earlier studies, Nixon et 220 al.¹⁰ suggested that, even when lung function was preserved, children with CF engaged in 221 significantly less VPA relative to healthy peers, whereas Selvadurai et al.⁶ and Britto et al.¹⁸ 222 found no differences in the intensity undertaken, with Britto et al.¹⁸ reporting VPA 223 participation to decline with age irrespective of disease status or severity. In contrast to the 224 present findings, Aznar et al.¹² and Jantzen et al.³² have previously reported lower total daily 225 VPA in children with CF. Moreover, Aznar et al.¹² also found a greater engagement in daily 226 TPA and LPA, the latter in agreement with the current study. Whilst the reason(s) for this 227 lack of consensus are likely to be multi-faceted, certain methodological differences should be 228 noted. Specifically, whilst a similar age range has been used in the majority of studies, ^{6,12,18,32} 229 pooling of data from boys and girls^{12,32} and a failure to account for maturity^{16,18,32} or disease 230 severity^{12,18} limits further inter-study comparisons. Indeed, Selvadurai et al.⁶ reported 231 significant influences of maturity and sex on PAL in those with CF and their healthy 232 counterparts. Caution is required when interpreting the PAL reported in previous studies that 233 have used long measurement epochs^{12,16} or questionnaires,^{6,10} with concerns raised regarding 234 the validity of questionnaire-derived PA estimates in chronic conditions such as CF,^{16,36} 235

which are susceptible to several forms of bias. In light of the highly sporadic nature of children's PA,^{37,38} with the median duration of high-intensity bouts suggested to be only 3s and 95% lasting less than 15s,^{38,39} the use of 15s epochs in this and previous studies may have influenced the findings, with VPA potentially miscategorised as MPA. Whilst the present study utilised this method to increase inter-study comparability of the results, future studies are suggested to use 1s epochs in accord with recommendations for the accurate assessment of PA intensity.⁴⁰

243

244 Alternatively, or additionally, discrepancies between accelerometry studies may be related to the cut-points used to delineate activity intensities. As there is a lack of age- and population-245 specific cut-points developed and validated for CF populations, each study has utilised 246 different cut-points, which has implications in the estimation of the time spent in different 247 activities.⁴¹ The impact of cut-point selection may be especially relevant in clinical 248 populations in whom it could be argued that the relative intensity of a given count rate is 249 250 higher than in their healthy counterparts. Whilst emphasising the need for disease-specific cut-points to be developed, this notion also highlights that the higher LPA reported here and 251 elsewhere in CF children may be clinically meaningful. Indeed, it has previously been 252 reported that time spent in low-LPA and high-LPA may have some favourable independent 253 health benefits¹⁹ but the minimum PA intensity and volume required to confer health benefits 254 255 remains to be elucidated. The present study further supports the potential importance of low-LPA and high-LPA by demonstrating these factors to significantly predict lung function 256 (FEV₁) in healthy and CF children, respectively. Further work is warranted to investigate 257 whether targeting increases in low-LPA and high-LPA rather than increases in MVPA per se, 258 may have beneficial health outcomes in this population, particularly given the high 259 correlation between LPA and sedentary time ¹⁹. Increasing LPA through interventions may be 260

a more feasible and constructive first step for the large proportion of patients not meeting
 current PA guidelines.¹⁹

263

Despite the increasing attention on sedentary behaviour as an independent risk factor for 264 cardiometabolic disease in children and youth,⁴² there is a lack of data regarding sedentary 265 behaviours in the CF population. In accord with Aznar et al.,¹² we found no difference in the 266 time spent sedentary by children with CF and their healthy counterparts. Whilst not the focus 267 of the present study, no relationships were found between sedentary behaviour and disease 268 severity, although the limited sample size should be considered when interpreting these 269 findings. Future studies should explore the potential relationship and interactions between 270 sedentary behaviour, PA and health in CF patients using objective measures and novel 271 272 statistical approaches to allow the optimal combination of these independent factors to be identified. Indeed, a growing body of evidence in healthy children suggests that the specific 273 type of sedentary behaviour (e.g., television viewing, computer use), rather than being 274 sedentary *per se*, may be an important determinant of health.^{43,44} 275

276

Emerging evidence suggests that the pattern in which PAL and sedentary time are accrued 277 may be an important determinant with regards to health. In healthy children, sedentary bouts 278 have been associated with C-reactive protein²¹ and HDL cholesterol.⁴⁵ However, in contrast, 279 Carson and Janssen⁴⁶ found that patterns of sedentary behavior were not related to cardio-280 metabolic risk factors in 6-19 year olds. Therefore, whether differences in the pattern of 281 sedentary time and PA have implications for health, particularly when TPA is similar, 282 remains to be resolved. The present study revealed no significant differences between the 283 groups with regards to the frequency or duration of sedentary or PA bouts, although there was 284 a trend for longer high-LPA bouts in the CF children. We did, however, observe significant 285

286 differences in the pattern of PA and sedentary behaviours during weekdays and weekend days, which were similar across the groups. Specifically, weekend days were characterized 287 by greater time spent sedentary with a lower frequency and duration of LPA and MPA bouts. 288 289 Since children potentially have more control over weekend free-time, it could be postulated that intra-individual differences may be most evident on weekend days.⁴⁷ Indeed, the greater 290 291 PAL during weekdays may, at least in part, be attributable to participation in Physical Education lessons and/or extra mural sports teams, although the effect of such isolated events 292 is likely to be minimal across seven days of objective PA assessment. Nonetheless, these 293 findings highlight the importance of considering different strategies to target week and 294 weekend day PA promotion. 295

296

297 Although the present study had numerous strengths, such as the objective measurement of PA, precisely matched healthy counterparts, and the novel consideration of the pattern in 298 which PA is accrued in those with CF, it is important to note certain limitations. Firstly, the 299 300 sample size was limited and, consequently, as was the range of disease severities included, although relative to the overall CF population, we believe that the present results provide 301 relevant and generalizable conclusions. Given the small sample size, the results of the present 302 linear regression should be considered exploratory; larger studies looking at the patterning of 303 PA across the disease spectrum would be invaluable in the future. It is pertinent to note that 304 305 whilst three or more days of valid PA data were required for inclusion in the analyses, no stipulations were made regarding the breakdown of these days between week and weekend 306 days. Given that PA is suggested to differ between weekdays and weekends in healthy^{48,49} 307 and CF youth¹² this may have influenced the current findings. The integration of postural 308 assessment may have provided greater insights into specific sedentary behaviours, such as 309 sitting. Furthermore, the lack of consistency in how bouts are defined (i.e., bout and 310

interruption length) limits cross-study comparisons;⁵⁰ the durations utilised in the present study were informed by research in healthy populations regarding bout and interruption durations.^{21,22} Finally, the cross-sectional design of the present study also limits the ability to make casual inferences regarding the relationships and their directionality.

315

316 Conclusions

In conclusion, the present study has demonstrated that there are no differences between CF 317 children and age- and sex-matched healthy controls with regards to overall PAL or the 318 319 manner in which these intensities are accrued, with significantly lower PA and greater sedentary behaviours during the weekend. Furthermore, the present study found LPA to be a 320 significant predictor of lung function in both healthy children and those with CF, although the 321 322 relevant intensity of LPA differed with high-LPA most important in those with CF. These findings therefore highlight weekends and LPA as areas warranting further research for the 323 development of effective intervention strategies to increase PA in the youth CF population. 324

325

326 Acknowledgments

We would like to thank Michele Barry and Julie Clarke from the Department of Child Health at Morriston Hospital, Swansea, for their assistance in conducting this study and the patients for their participation.

330

331 Funding Source

332 No funding was received to support this study.

333

334 **References**

- Quinton PM. Cystic fibrosis: A disease in electrolyte transport. *FASEB Journal*.
 1990;4:2709-2717.
- 2. Cystic Fibrosis Foundation. Patient Registry: Annual Report 20122012.
- 338 3. Janssen I, Leblanc AG. Systematic review of the health benefits of physical activity
- and fitness in school-aged children and youth. *Int J Behav Nutr Phys Act.* 2010;7:40.
- 340 4. Schneiderman-Walker J, Pollock SL, Corey M, et al. A randomized controlled trial of
- a 3-year home exercise program in cystic fibrosis. *J Pediatr*. 2000;136:304-310.
- 342 5. Wilkes DL, Schneiderman-Walker J, Corey M, et al. Longterm effect of habitual
- 343 physical activity on lung health in patients with cystic fibrosis. *Pediatr Pulmonol*.
- **344** 2007:358-359.
- Selvadurai HC, Blimkie CJ, Cooper PJ, et al. Gender differences in habitual activity
 in children with cystic fibrosis. *Arch Dis Child*. 2004;89:928-933.
- 3477.Buntain HM, Greer RM, Schluter PJ, et al. Bone mineral density in Australian
- children, adolescents and adults with cystic fibrosis: a controlled cross sectional
 study. *Thorax*. 2004;59:149-155.
- McIlwaine M. Chest physical therapy, breathing techniques and exercise in children
 with CF. *Paediatr Respir Rev.* 2007;8:8-16.
- Hebestreit A, Kersting U, Basler B, et al. Exercise inhibits epithelial sodium channels
 in patients with cystic fibrosis. *Am J Respir Crit Care Med*. 2001;164:443-446.
- Nixon PA, Orenstein DM, Kelsey SF. Habitual physical activity in children and
 adolescents with cystic fibrosis. *Med Sci Sport Exerc.* 2001;33:30-35.
- Nixon PA, Orenstein DM, Kelsey SF, et al. The Prognostic Value of Exercise Testing
 in Patients with Cystic-Fibrosis. *N Engl J Med.* 1992;327:1785-1788.

- Aznar S, Gallardo C, Fiuza-Luces C, et al. Levels of moderate–vigorous physical
 activity are low in Spanish children with cystic fibrosis: A comparison with healthy
 controls. *J Cyst Fibros*. 2014;13:335-340.
- 13. Department of Health Physical Activity Health Improvement and Protection. Start
- 362 Active, Stay Active: A report on physical activity from the four home countries' Chief
- 363 Medical Officers. London: Department of Health 2011.
- Radtke T, Nolan SJ, Hebestreit H, et al. Physical exercise training for cystic fibrosis. *Cochrane Database Syst Rev.* 2015.
- Evenson KR, Catellier DJ, Gill K, et al. Calibration of two objective measures of
 physical activity for children. *J Sport Sci.* 2008;24:1557-1565.
- 368 16. Jantzen A, Opoku-Pare M, Bieli C, et al. Perspective on cystic fibrosis and physical
 activity: Is there a difference compared to healthy individuals? *Pediatr Pulmonol*.
 2016.
- Trost SG, Wong WK, Pfeiffer KA, et al. Artificial neural networks to predict activity
 type and energy expenditure in youth. *Med Sci Sport Exerc*. 2012;44:1801-1809.
- 18. Britto MT, Garrett JM, Konrad TR, et al. Comparison of physical activity in
- adolescents with cystic fibrosis versus age-matched controls. *Pediatr Pulmonol*.
 2000;30:86-91.
- Carson V, Ridgers ND, Howard BJ, et al. Light-intensity physical activity and
 cardiometabolic biomarkers in US adolescents. *PLoS One*. 2013;8:e71417.
- 37820.Schneiderman JE, Wilkes DL, Atenafu EG, et al. Longitudinal relationship between
- physical activity and lung health in patients with cystic fibrosis. *Eur Respir J*.
- 380 2014;43:817-823.

- 381 21. Gabel L, Ridgers ND, Della Gatta PA, et al. Associations of sedentary time patterns
 382 and TV viewing time with inflammatory and endothelial function biomarkers in
 383 children. *Pediatr Obes*. 2016;11:194-201.
- 384 22. Mark AE, Janssen I. Influence of bouts of physical activity on overweight in youth.
 385 *Am J Prev Med.* 2009;36:416-421.
- Stanojevic S, Wade A, Cole TJ, et al. Spirometry centile charts for young Caucasian
 children: the Asthma UK Collaborative Initiative. *Am J Respir Crit Care Med*.
 2009;180:547-552.
- 389 24. Trost SG, Ward DS, Moorehead SM, et al. Validity of the Computer Science and
- Application (CSA) activity monitor in children. *Med Sci Sport Exerc*. 1998;30:629633.
- Ridgers ND, Salmon J, Ridley K, et al. Agreement between activPAL and ActiGraph
 for assessing children's sedentary time. *Int J Behav Nutr Phys Act*. 2012;9:15.
- 39426.Freedson P, Pober D, Janz KF. Calibration of Accelerometer Output for Children.
- 395 *Med Sci Sport Exerc*. 2005;37(11):S523-S530.
- Trost SG, Loprinzi PD, Moore R, et al. Comparison of accelerometer cut-points for
 predicting activity intensity in youth. *Med Sci Sport Exerc*. 2011;43:1360-1368.
- 398 28. Mackintosh KA, Ridley K, Stratton G, et al. Energy Cost of Free-Play Activities in
- 399 10- to 11-Year-Old Children. *J Phys Act Health*. 2016;13:S71-74.
- 400 29. Ryan JM, Forde C, Hussey JM, et al. Comparison of Patterns of Physical Activity and
- 401 Sedentary Behavior Between Children With Cerebral Palsy and Children With
- 402 Typical Development. *Physical Therapy*. 2015;95:1609-1616.
- 40330.Mattocks C, Ness AR, Leary SD, et al. Use of accelerometers in a large field-based
- 404 study of children: Protocols, design issues, and effects on precision. *J Phys Activ*
- 405 *Health*. 2008;5:S98-S111.

- 406 31. Healy GN, Wijndaele K, Dunstan DW, et al. Objectively measured sedentary time,
- 407 physical activity, and metabolic risk: the Australian Diabetes, Obesity and Lifestyle
 408 Study (AusDiab). *Diabetes Care*. 2008;31:369-371.
- Jantzen A, Opoku-Pare M, Ruf K, et al. Cystic fibrosis and physical activity: Is there
 a significant difference to healthy individuals? *Eur Respir J*. 2014;44.
- 411 33. Bradley JM, Moran F. Physical training for cystic fibrosis. *Cochrane Database Syst*412 *Rev.* 2008.
- 413 34. Stevens D, Oades PJ, Armstrong N, et al. A survey of exercise testing and training in
 414 UK cystic fibrosis clinics. *J Cyst Fibros*. 2010;9:302-306.
- 415 35. Williams CA, Saynor ZL, Tomlinson OW, et al. Cystic fibrosis and physiological
 416 responses to exercise. *Expert Rev Respir Med.* 2014;8:751-762.
- 417 36. Ruf KC, Fehn S, Bachmann M, et al. Validation of activity questionnaires in patients
 418 with cystic fibrosis by accelerometry and cycle ergometry. *BMC Med Res Methodol*.
 419 2012;12.
- 420 37. Rowlands AV, Eston RG, Ingledew DK. Measurement of Physical Activity in
- 421 Children with Particular Reference to the use of Heart Rate and Pedometry. *Sports*422 *Med.* 1997;24:259-272.
- 42338.Bailey RC, Olson J, Pepper SL, et al. The level and tempo of children's physical
- 424 activities: an observational study. *Med Sci Sport Exerc*. 1995;27:1033-1041.
- 42539.Baquet G, Stratton G, Van Praagh E, et al. Improving physical activity assessment in
- 426 prepubertal children with high-frequency accelerometry monitoring: A
- 427 methodological issue. *Prev Med.* 2007;44:143-147.
- 428 40. Rowland A, Eston RG. The measurement and interpretation of children's physical
 429 activity. *J Sport Sci Med.* 2007;6:270-276.

430	41.	Banda JA, Haydel KF, Davila T, et al. Effects of Varying Epoch Lengths, Wear Time
431		Algorithms, and Activity Cut-Points on Estimates of Child Sedentary Behavior and
432		Physical Activity from Accelerometer Data. PLoS One. 2016;11:e0150534.
433	42.	Saunders TJ, Chaput J-P, Tremblay MS. Sedentary Behaviour as an Emerging Risk
434		Factor for Cardiometabolic Diseases in Children and Youth. Can J Diabetes.
435		2014;38:53-61.
436	43.	Tremblay MS, LeBlanc AG, Kho ME, et al. Systematic review of sedentary behaviour
437		and health indicators in school-aged children and youth. Int J Behav Nutr Phys Act.
438		2011;8:1-22.
439	44.	Robinson S, Daly RM, Ridgers ND, et al. Screen-Based Behaviors of Children and
440		Cardiovascular Risk Factors. J Pediatr. 2015;167:1239-1245.
441	45.	Cliff DP, Jones RA, Burrows TL, et al. Volumes and bouts of sedentary behavior and
442		physical activity: Associations with cardiometabolic health in obese children. Obesity.
443		2014;22:E112-E118.
444	46.	Carson V, Janssen I. Volume, patterns, and types of sedentary behavior and cardio-
445		metabolic health in children and adolescents: a cross-sectional study. BMC Public
446		Health. 2011;11:1-10.
447	47.	Stone MR, Rowlands AV, Eston RG. Characteristics of the activity pattern in normal
448		weight and overweight boys. Prev Med. 2009;49:205-208.
449	48.	Evenson KR, Wen F, Hales D, et al. National youth sedentary behavior and physical
450		activity daily patterns using latent class analysis applied to accelerometry. Int J Behav
451		Nutr Phys Act. 2016;13:55.
452	49.	Fairclough SJ, Boddy L, M, Mackintosh KA, et al. Weekday and weekend sedentary
453		time and physical activity in differentially active children. J Sci Med Sport.
454		2015;444-449.

19

- 455 50. Chinapaw MJM, de Niet M, Verloigne M, et al. From Sedentary Time to Sedentary
- 456 Patterns: Accelerometer Data Reduction Decisions in Youth. *PLoS One*.
- 457 2014;9:e111205.
- 458
- 459
- 460 Tables
- 461

462 **Table 1.** Participant characteristics

	Total	Cystic Fibrosis	Controls
n	36	18	18
Age (yrs)	12.6 ± 2.7	12.4 ± 2.8	12.5 ± 2.7
Stature (m)	1.48 ± 0.14	1.46 ± 0.14	1.51 ± 0.13
Mass (kg)	44.24 ± 12.99	41.16 ± 12.51	47.52 ± 13.04
Waist circumference (m)	0.67 ± 0.08	0.66 ± 0.07	0.67 ± 0.09
BMI (kg·m ²)	19.6 ± 3.4	18.8 ± 2.8	20.5 ± 3.8
Maturity offset (yrs from PHV)	-1.28 ± 3.00	-1.04 ± 2.42	-1.54 ± 3.57
FVC (% predicted)	84 ± 15	83 ± 12	85 ± 18
FEV ₁ (% predicted)	85 ± 14	80 ± 9	$89\pm17^{*}$

463 Mean \pm S.D. PHV, peak height velocity; FVC, forced vital capacity; FEV₁, forced expiratory volume

464 in 1 second. * significant difference between control and Cystic Fibrosis

465

Table 2. Physical activity data by group

	Total	CF	Control
	<i>n</i> = 32	<i>n</i> = 16	<i>n</i> = 16
Overall			
Sedentary time (min·day ⁻¹)	545.4 ± 76.0	539.2 ± 64.6	551.3 ± 87.0
Low-LPA (min·day ⁻¹)	141.4 ± 34.9	144.3 ± 30.9	138.7 ± 39.1
High-LPA (min·day ⁻¹)	72.5 ± 23.9	77.5 ± 20.6	67.7 ± 26.3
MPA (min·day ⁻¹)	40.5 ± 22.6	39.5 ± 23.3	41.3 ± 22.6
VPA (min·day ⁻¹)	12.6 ± 10.6	13.1 ± 12.4	12.1 ± 9.0
VVPA (min·day ⁻¹)	2.7 ± 3.4	2.7 ± 3.9	2.7 ± 2.9
MVPA (min·day ⁻¹)	55.7 ± 32.7	55.3 ± 38.0	56.1 ± 27.9
Week days			
Sedentary time (min·day ⁻¹)	542.8 ± 84.1	532.1 ± 69.2	552.3 ± 96.5
Low-LPA (min·day ⁻¹)	146.2 ± 36.1	149.7 ± 27.0	143.2 ± 43.2
High-LPA (min·day ⁻¹)	76.9 ± 26.3	83.8 ± 22.0	70.8 ± 29.0
MPA (min·day ⁻¹)	43.8 ± 22.0	43.7 ± 22.8	43.8 ± 21.9
VPA (min·day ⁻¹)	13.7 ± 11.2	14.7 ± 12.7	12.8 ± 10.0
VVPA (min·day ⁻¹)	2.7 ± 3.3	3.0 ± 4.0	2.4 ± 2.5
MVPA (min·day ⁻¹)	60.2 ± 32.4	61.5 ± 38.0	59.0 ± 27.6
Weekend days			
Sedentary time (min·day ⁻¹)	555.7 ± 91.3	554.9 ± 94.5	556.5 ± 90.1
Low-LPA (min·day ⁻¹)	135.7 ± 35.7	140.3 ± 38.6	130.9 ± 32.9
High-LPA (min·day ⁻¹)	66.0 ± 22.7	69.2 ± 22.4	62.6 ± 23.1
MPA (min·day ⁻¹)	36.0 ± 27.7	$34.5\pm27.1^*$	$37.6 \pm 29.1^{*}$
VPA (min·day ⁻¹)	11.1 ± 12.5	11.3 ± 14.0	11.0 ± 11.1

Physical Activity Levels in Cystic Fibrosis Youth

VVPA (min·day ⁻¹)	3.1 ± 5.7	2.4 ± 3.8	3.8 ± 7.2
MVPA (min·day ⁻¹)	50.3 ± 39.6	$48.2\pm41.4^*$	$52.5 \pm 38.9^{*}$

Means ± SD. Low-LPA, low light physical activity; High-LPA, high light physical activity; MPA,
moderate physical activity; VPA, vigorous physical activity; VVPA, very vigorous physical activity;
MVPA, moderate-to-vigorous physical activity. * Significant difference between week- and weekend
day within group
471
472
473

- 474
- 475

	Total	CF	Control
	<i>n</i> = 32	<i>n</i> = 16	<i>n</i> = 16
Overall			
Frequency SED	28 ± 7	27 ± 7	28 ± 7
Duration SED (mins)	271.2 ± 83.5	263.7 ± 86.2	278.3 ± 82.6
Number SED Breaks	301.3 ± 57.2	303.6 ± 56.2	299.0 ± 59.7
Frequency LPA	62 ± 18	66 ± 16	59 ± 20
Duration LPA (mins)	98.4 ± 33.7	105.5 ± 28.7	91.6 ± 37.3
Frequency Low-LPA	22 ± 8	23 ± 6	21 ± 9
Duration Low-LPA (mins)	27.0 ± 9.8	28.2 ± 7.3	26.0 ± 11.8
Frequency High-LPA	9 ± 7	11 ± 7	8 ± 8
Duration High-LPA (mins)	13.9 ± 14.1	16.6 ± 14.7	11.3 ± 13.3
Frequency MPA	7 ± 4	7 ± 4	7 ± 5
Duration MPA (mins)	11.8 ± 12.1	10.1 ± 6.3	13.3 ± 15.8
Frequency VPA	3 ± 3	3 ± 4	3 ± 3
Duration VPA (mins)	5.2 ± 5.9	5.3 ± 7.4	5.1 ± 4.1
Weekdays			
Frequency SED	27 ± 8	26 ± 7	28 ± 9
Duration SED (mins)	262.4 ± 88.8	251.6 ± 88.3	272.1 ± 90.6
Number SED Breaks (mins)	309.0 ± 54.8	311.8 ± 45.4	306.6 ± 63.2
Frequency LPA	65 ± 21	69 ± 17	61 ± 24
Duration LPA (mins)	103.5 ± 39.6	112.1 ± 32.1	95.8 ± 44.7
Frequency Low-LPA	23 ± 9	23 ± 6	22 ± 11
Duration Low-LPA (mins)	27.9 ± 11.3	28.8 ± 7.7	27.1 ± 14.0
Frequency High-LPA	10 ± 8	12 ± 8	9 ± 9

Table 3. Patterns of PA accumulation on week days, weekend days and overall (average day)

Duration High-LPA (mins)	15.7 ± 17.7	18.8 ± 19.6	12.9 ± 15.8
Frequency MPA	8 ± 4	8 ± 4	8 ± 5
Duration MPA (mins)	13.0 ± 12.0	11.4 ± 6.3	14.4 ± 15.5
Frequency VPA	3 ± 3	3 ± 4	3 ± 3
Duration VPA (mins)	5.3 ± 6.1	5.8 ± 7.5	4.9 ± 4.6
Weekend days			
Frequency SED	29 ± 8	29 ± 9	29 ± 8
Duration SED (mins)	286.8 ± 97.1	278.4 ± 108.6 *	295.7 ± 85.8 *
Number SED Breaks	293.0 ± 66.1	299.6 ± 78.8	286.0 ± 51.1
Frequency LPA	59 ± 16	62 ± 17 *	56 ± 15 *
Duration LPA (mins)	90.8 ± 26.2	$97.0\pm27.8~^{\#}$	84.2 ± 23.4 $^{\#}$
Frequency Low-LPA	21 ± 8	22 ± 8	19 ± 7
Duration Low-LPA (mins)	25.6 ± 10.3	27.5 ± 10.7	23.6 ± 9.9
Frequency High-LPA	8 ± 6	9 ± 6 [#]	6 ± 6 [#]
Duration High-LPA (mins)	10.2 ± 8.8	12.7 ± 9.1 $^{\#}$	6.1 ± 6.0 $^{\#}$
Frequency MPA	6 ± 6	5 ± 5 [#]	6 ± 7 [#]
Duration MPA (mins)	9.8 ± 14.7	$8.3\pm9.8~^*$	11.4 \pm 18.8 *
Frequency VPA	3 ± 4	3 ± 4	3 ± 4
Duration VPA	5.3 ± 7.6	4.9 ± 8.3	5.7 ± 7.1

477 Mean \pm SD. SED, sedentary behaviour; Low-LPA, low light physical activity; High-LPA, high light 478 physical activity; LPA, light physical activity; MPA, moderate physical activity; VPA, vigorous 479 physical activity. * Significant difference within condition between weekday and weekend P < 0.05; # 480 Significant difference within condition between weekday and weekend P < 0.01