



# **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/](http://www.swansea.ac.uk/iss/researchsupport/cronfa-support/ ) 

## **Abstract**

 *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 activity levels (PAL) or how these are accrued in those with CF. *Methods:* Physical activity was measured by a hip-worn accelerometer for seven consecutive days by eighteen children 6 (10 boys;  $12.4 \pm 2.8$  years) with mild to moderate CF and eighteen age- and sex-matched 7 controls (10 boys;  $12.5 \pm 2.7$  years). **Results:** Both CF and healthy children demonstrated similar PAL and patterns of accumulation across the intensity spectrum, with higher levels of 9 PA during weekdays in both groups. FEV<sub>1</sub> was predicted by high-light PA in CF compared to low-light PA in healthy children. *Conclusion:* These findings highlight weekends and light PA as areas warranting further research for the development of effective intervention strategies to increase PA in the youth CF population.

#### **Introduction**

 Cystic Fibrosis (CF) is the most prevalent lethal autosomal recessive disease in the Caucasian 15 population.<sup>1</sup> Mutations in the cystic fibrosis transmembrane regulator (CFTR) gene lead to malfunctioning or absent CFTR proteins, impairing mucosal clearance mechanisms. As such, CF is characterised by excessive viscous secretions in almost all organs, particularly the lungs, resulting in recurring infections, inflammation, airflow obstruction, and ultimately progressive functional decline. Whilst there remains no cure, advances in the treatment for patients with CF have resulted in an increased median life expectancy from 8 years in 1974, to 31 years in 2005 and 41 years in 2012. 2 

 Whilst physical activity (PA) has been associated with numerous physiological and 24 psychosocial benefits for healthy children,<sup>3</sup> there are additional health benefits for patients with CF. These include slower lung function decline,<sup>4</sup> reduced hospital admissions,<sup>5</sup> 26 improved quality of life and nutritional status,<sup>6</sup> improved bone mineral density,<sup>7</sup> and 27 enhanced airway clearance<sup>8</sup> and ion channel function, which could lead to improved mucus 28 hydration and clearance.<sup>9</sup> PA could be imperative for ultimate survival in patients with severe 29 lung deterioration, given the strong positive relationship with aerobic capacity.<sup>10,11</sup> However, 30 recent research suggests that as little as  $2.1\%$  of children with  $CF^{12}$  meet the government PA guidelines of at least 60 minutes of moderate-to-vigorous-intensity physical activity (MVPA) 32 every day.<sup>13</sup> Whilst regular PA is increasingly important in the care of patients with CF,<sup>14</sup> there is a dearth of research and indeed little consensus on physical activity levels (PAL) in children and adolescents with CF. Moreover, little is known as to whether beneficial outcomes may be achieved with engagement of PA at different intensities, which would be critical information for interventions and on going care.

 Variations in PAL reported in the existing literature may be due to methodological inconsistencies. Earlier research employing self-reported measures found that children aged 7-17 years with CF participated in less very strenuous (> 6 METs) PA relative to healthy 41 controls, even when patients had well-preserved lung function.<sup>10</sup> Conversely, Selvadurai and 42 colleagues<sup>6</sup> reported no significant differences between CF patients and age- and sex- matched controls in similarly-aged children (9-17 years), using uniaxial accelerometry. 44 Advancing previous research, which only reported total counts<sup>6</sup>, Aznar et al.<sup>12</sup> utilised 45 Evenson<sup>15</sup> cut-points to find that 6-17 year old children with CF engaged in significantly less MVPA and vigorous physical activity (VPA) but demonstrated higher total and light physical 47 activity (TPA and LPA, respectively). Yet Jantzen et al.<sup>16</sup> found similar PAL in CF patients across the age and intensity spectrum, but less engagement in strenuous activities for school- aged children (6-13 years) compared to healthy controls. Interestingly, when extreme values were removed, no relationship was present between strenuous PA and percentage predicted  $\rm FEV_1.^{16}$ 

52

53 A potential limitation of earlier studies is the lack of age- and population-specific cut-points, 54 although, in healthy populations, it is pertinent to note that Trost and colleagues<sup>17</sup> supported 55 the use of Evenson cut-points. Arguably, the relative intensity for children and adolescents 56 with CF could be greater and therefore light physical activity may be more beneficial to their 57 health in comparison to their healthy counterparts. However, with the exception of Aznar and 58 colleagues,<sup>12</sup> and to some extent Jantzen et al.,<sup>16</sup> the majority of studies did not consider PA 59 across the spectrum.<sup>6,10,18</sup> As such, the identification of patients with CF participating in more 60 LPA and  $TPA<sup>12</sup>$  may warrant further investigation. Specifically, previous research has 61 suggested that time spent in low-light physical activity (low-LPA) and high-light physical 62 activity (high-LPA) may have some favourable independent health benefits.<sup>19</sup> Additionally, a  sedentary lifestyle has been shown to contribute to the progression of both functional and 64 physical impairment in CF populations,<sup>20</sup> yet little research has objectively assessed time spent being sedentary, nor the accumulation of PA or sedentary time. Indeed, the majority of 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. 68 Gabel et al.<sup>21</sup> reported sedentary bouts of  $\geq$ 5 minutes to be detrimentally associated with C-69 reactive protein in healthy children, whereas PA bouts of  $\geq 1$  minute, which have previously 70 been used to identify sporadic bouts of PA, are reported to be associated with lower BMI.<sup>22</sup> Identifying patterns of accumulation in youth with CF, and those patterns that may be associated with functional gains, is important for advancing the design and evaluation of future interventions in this population.

 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 PA children and adolescents with and without CF. Therefore, the purpose of this study was to 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 ascertain whether such parameters could predict lung function. We hypothesised that PA intensity and duration would be significantly lower in patients with CF and be a significant predictor of disease severity (i.e., lung function).

#### **Methods**

*Participants*

86 In total, 36 participants (12.6  $\pm$  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-

88 moderate CF, confirmed by a sweat chloride > 60 mmol·l<sup>-1</sup> and genotyping (8  $\Delta$ F508 Homozygote, 10 ΔF508 Heterzygote; 4 CF-related liver disease) were recruited from an outpatient CF clinic in South Wales (United Kingdom). Patients were included in the study if 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); unstable non-pulmonary comorbidities or acute infections warranted exclusion. Eighteen age- and sex-matched non-clinical children were recruited from local schools to act as a healthy comparison group. Ethical approval was granted by the Bromley NHS research ethics 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 continue prescribed medications as usual throughout the duration of their study involvement.

#### *Measurements*

 At their routine visits to the clinic, participants forced vital capacity (FVC) and forced expiratory volume in 1s (FEV1) were assessed using flow-volume loop spirometry (Vitalograph, UK)). The best of three consistent exhalations (<5% variability) was recorded. All lung function measurements were expressed as a percentage predicted normal, using 105 appropriate reference data.<sup>23</sup> Furthermore, body mass (Seca 220; Hamburg, Germany), stature and sitting stature (Seca 220; Hamburg, Germany) were measured to the nearest 0.01 kg and 0.01 m, respectively. Waist circumference was measured to the nearest 0.01 m using a non- 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 to attend one laboratory session at Swansea University for all measurements to be undertaken. All participants were provided with a hip-mounted ActiGraph GT3X+ 112 accelerometer (ActiGraph LLC, Pensacola, FL) to assess habitual PAL over seven 113 consecutive days.

114

115 ActiGraph monitors, shown to have acceptable reliability and validity in paediatric 116 populations,  $24$  sampled raw data at 100Hz. Data were downloaded using ActiLife software 117 (v6.10.4; ActiGraph, Pensacola, FL), processed into 15s epochs and reduced using a 118 customised Excel macro. Sustained periods of 20 minutes of consecutive zero's were used to 119 define non-wear time, which has been found to result in an almost identical wear time and a 120 smaller difference between sedentary time and sitting time estimates (assessed using 121 activPAL; PAL Technologies, Glasgow, Scotland) compared with a 60 minute definition in 122 children.<sup>21</sup> Sedentary time was defined as <100 counts·min<sup>-1</sup>, shown to be a good estimate of 123 free-living sitting.<sup>25</sup> Time spent in MPA (4-5.99 METs) and VPA ( $\geq$ 6 METs) was determined 124 using age-specific cut-points,<sup>26</sup> which demonstrated comparable accuracy to Evenson cut-125 points.<sup>17</sup> A threshold of 4 METs was used to define MPA, as brisk walking has been 126 associated with this energy cost in calibration studies.<sup>27,28</sup> MPA and VPA were summed to 127 create MVPA. The rest of the time was classified as either low light-intensity physical 128 activity (low-LPA; 100-799 counts·min<sup>-1</sup>) or high light-intensity physical activity (high-129 LPA; 800-<4 METs). The 800 counts·min<sup>-1</sup> threshold was selected as this published 130 sedentary cut-point captures both sedentary time and static light-intensity activities such as 131 standing, $25$  and has been found to have differential associations with cardiometabolic 132 biomarkers in adolescents.<sup>19</sup> A valid day was defined as ≥9 hours·day<sup>-1</sup>, which has been 133 previously used in clinical populations.<sup>29</sup> To be included in the analyses, children were 134 required to have worn the ActiGraph for at least three days, which has been shown to have a 135 reliability coefficient of 0.7.<sup>30</sup> PAL are reported for overall, weekday and weekend days 136 separately.



 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 140 epoch following a 15s epoch of  $\langle 25 \rangle$  counts per epoch.<sup>31</sup> The frequency and duration of time 141 spent in sedentary ( $\geq$ 5 min),<sup>21</sup> and low-LPA, high-LPA, MPA and VPA ( $\geq$ 1 minutes) were 142 also determined. $^{22}$  No interruptions to these bouts were permitted.

#### *Data Analysis*

 Gaussian distribution was confirmed by the Shapiro-Wilks test. Following this, the participant groups and weekday vs. weekend day were compared using a multivariate ANCOVA with group as a fixed factor and day as a repeated measure, controlling for wear 148 time. A stepwise linear regression was used to analyse the association between  $FEV<sub>1</sub>$  and PA intensity levels and patterns, adjusting for predefined potential confounders (age, sex, stature, mass and wear time). To explore differences between the groups in terms of those that met current government guidelines for PA (i.e., average of ≥60 minutes of MVPA/day), a Chi- square test was used. All statistical analyses were conducted using PASW Statistics 21 153 (SPSS, Chicago, IL). All data are presented as means  $\pm$  SD. Statistical significance was 154 accepted when  $P \le 0.05$ .

#### **Results**

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

- 161 However, those with CF presented with a significantly lower percentage of predicted  $FEV<sub>1</sub>$ and FEV:FVC when described in both absolute and relative to predicted terms.
- 

 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, 167 participants achieved  $4.5 \pm 1.2$  and  $1.8 \pm 0.6$  valid weekdays and weekend days, respectively. 

 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 171 (Table 2). There was a trend for greater time spent in LPA in CF patients  $(222.7 \pm 12.8 \text{ vs.})$  $207.3 \pm 12.4$  mins;  $P > 0.05$ ), although this failed to reach significance. There were significant differences between weekday and weekend day PA with regards to total LPA 174 (229.3  $\pm$  52.4 vs. 203.8  $\pm$  50.6 mins, respectively; *P* < 0.05), MPA (45.1  $\pm$  21.5 vs. 36.6  $\pm$ 175 27.9 mins, respectively;  $P < 0.05$ ) and MVPA (62.4  $\pm$  32.1 vs. 51.2  $\pm$  39.9 mins, respectively;  $P < 0.05$ ), with greater levels of activity achieved during weekdays than weekend days in both groups.

 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.

 Healthy controls and CF patients demonstrated similar patterns of physical activity accumulation (Table 3). However, different patterns were evident during weekday and 186 weekend days, with weekdays characterised by a greater frequency and duration of LPA and 187 MPA bouts and a lower duration of sedentary bouts compared to weekend days.

188

189 Linear regression revealed that  $FEV_1$  was predicted by height and LPA when both groups 190 were pooled for analysis  $(F_{(2,31)} = 62.93, P < 0.001; R^2 = 0.80)$ . More specifically, when LPA 191 was split into low-LPA and high-LPA, height and low-LPA significantly predicted  $FEV<sub>1</sub>$ 192  $(F_{(2,31)} = 68.07, P < 0.001; R^2 = 0.82)$ . When the groups were considered independently, the 193 intensity of LPA that predicted  $FEV_1$  differed, with  $FEV_1$  predicted by height and high-LPA 194 in CF patients ( $F_{(2,14)} = 79.60$ ,  $P < 0.001$ ;  $R^2 = 0.92$ ) compared to height and low-LPA in 195 healthy controls  $(F_{(2,14)} = 24.31, P < 0.001; R^2 = 0.78)$ .

196

## 197 **Discussion**

 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,  $FEV<sub>1</sub>$  was dependent on LPA levels in both CF patients and their healthy counterparts, although the intensity within LPA differed across the groups. Finally, we observed significant decreases in PAL during weekends, with increased sedentary time and decreased frequency and duration of LPA and MPA bouts, irrespective of disease status.

204

205 In agreement with some,  $6,18,32$  but not all,  $10,12$  previous studies, no significant difference was 206 observed in the PAL of children with and without CF, although a considerably higher 207 proportion of our CF population met recommended guidelines compared to previous 208 research.<sup>12</sup> Given the numerous additional health benefits for patients with  $CF<sub>1</sub><sup>4-9</sup>$  over and 209 above the physiological and psychosocial benefits of regular PA identified in healthy 210 children,<sup>3</sup> these findings highlight the need for strategies to increase PA in this population.

211 Indeed, the importance of PA has been recognised by the European Cystic Fibrosis Society 212 (ECFS) and recent Cochrane Reviews,  $14,33$  which advocate the cost-effectiveness and 213 beneficial effects of PA for promoting quality of life in patients with CF. However, 214 information regarding PA behaviours in CF is limited and although PA as a treatment is 215 becoming increasingly valued by CF clinical teams, $34$  it remains underutilized in routine CF 216 management  $35$ . Furthermore, there is a paucity of evidence-based guidance regarding the 217 optimal combination of intensity and duration to elicit health benefits.

218

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

 which are susceptible to several forms of bias. In light of the highly sporadic nature of 237 children's PA,  $37,38$  with the median duration of high-intensity bouts suggested to be only 3s 238 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 242 of PA intensity.

 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- specific cut-points developed and validated for CF populations, each study has utilised different cut-points, which has implications in the estimation of the time spent in different 248 activities.<sup>41</sup> The impact of cut-point selection may be especially relevant in clinical populations in whom it could be argued that the relative intensity of a given count rate is 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 elsewhere in CF children may be clinically meaningful. Indeed, it has previously been 253 reported that time spent in low-LPA and high-LPA may have some favourable independent 254 health benefits<sup>19</sup> but the minimum PA intensity and volume required to confer health benefits 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 (FEV<sub>1</sub>) in healthy and CF children, respectively. Further work is warranted to investigate whether targeting increases in low-LPA and high-LPA rather than increases in MVPA *per se,* may have beneficial health outcomes in this population, particularly given the high 260 correlation between LPA and sedentary time . Increasing LPA through interventions may be  a more feasible and constructive first step for the large proportion of patients not meeting 262 current PA guidelines.<sup>19</sup>

 Despite the increasing attention on sedentary behaviour as an independent risk factor for 265 cardiometabolic disease in children and youth, there is a lack of data regarding sedentary 266 behaviours in the CF population. In accord with Aznar et al.,  $^{12}$  we found no difference in the 267 time spent sedentary by children with CF and their healthy counterparts. Whilst not the focus of the present study, no relationships were found between sedentary behaviour and disease severity, although the limited sample size should be considered when interpreting these findings. Future studies should explore the potential relationship and interactions between sedentary behaviour, PA and health in CF patients using objective measures and novel 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 type of sedentary behaviour (e.g., television viewing, computer use), rather than being 275 sedentary *per se*, may be an important determinant of health.<sup>43,44</sup>

 Emerging evidence suggests that the pattern in which PAL and sedentary time are accrued may be an important determinant with regards to health. In healthy children, sedentary bouts 279 have been associated with C-reactive protein<sup>21</sup> and HDL cholesterol.<sup>45</sup> However, in contrast, 280 Carson and Janssen<sup>46</sup> found that patterns of sedentary behavior were not related to cardio- metabolic risk factors in 6-19 year olds. Therefore, whether differences in the pattern of sedentary time and PA have implications for health, particularly when TPA is similar, remains to be resolved. The present study revealed no significant differences between the groups with regards to the frequency or duration of sedentary or PA bouts, although there was 285 a trend for longer high-LPA bouts in the CF children. We did, however, observe significant  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 by greater time spent sedentary with a lower frequency and duration of LPA and MPA bouts. Since children potentially have more control over weekend free-time, it could be postulated 290 that intra-individual differences may be most evident on weekend days.<sup>47</sup> Indeed, the greater 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 is likely to be minimal across seven days of objective PA assessment. Nonetheless, these findings highlight the importance of considering different strategies to target week and weekend day PA promotion.

 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 which PA is accrued in those with CF, it is important to note certain limitations. Firstly, the 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 relevant and generalizable conclusions. Given the small sample size, the results of the present linear regression should be considered exploratory; larger studies looking at the patterning of 304 PA across the disease spectrum would be invaluable in the future. It is pertinent to note that 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 307 days. Given that PA is suggested to differ between weekdays and weekends in healthy<sup>48,49</sup> 308 and CF youth<sup>12</sup> this may have influenced the current findings. The integration of postural assessment may have provided greater insights into specific sedentary behaviours, such as sitting. Furthermore, the lack of consistency in how bouts are defined (i.e., bout and

311 interruption length) limits cross-study comparisons;<sup>50</sup> the durations utilised in the present study were informed by research in healthy populations regarding bout and interruption 313 durations.<sup>21,22</sup> Finally, the cross-sectional design of the present study also limits the ability to make casual inferences regarding the relationships and their directionality.

## **Conclusions**

 In conclusion, the present study has demonstrated that there are no differences between CF children and age- and sex-matched healthy controls with regards to overall PAL or the 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 significant predictor of lung function in both healthy children and those with CF, although the 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 development of effective intervention strategies to increase PA in the youth CF population.

#### **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.

## **Funding Source**

No funding was received to support this study.

#### **References**

- 1. Quinton PM. Cystic fibrosis: A disease in electrolyte transport. *FASEB Journal*. 1990;4:2709-2717.
- 2. Cystic Fibrosis Foundation. Patient Registry: Annual Report 20122012.
- 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.
- 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.
- 5. Wilkes DL, Schneiderman-Walker J, Corey M, et al. Longterm effect of habitual
- physical activity on lung health in patients with cystic fibrosis. *Pediatr Pulmonol*.
- 2007:358-359.
- 6. 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.
- 7. 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.
- 8. McIlwaine M. Chest physical therapy, breathing techniques and exercise in children with CF. *Paediatr Respir Rev*. 2007;8:8-16.
- 9. 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.
- 10. Nixon PA, Orenstein DM, Kelsey SF. Habitual physical activity in children and adolescents with cystic fibrosis. *Med Sci Sport Exerc*. 2001;33:30-35.
- 11. 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.
- 12. 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
- Active, Stay Active: A report on physical activity from the four home countries' Chief
- Medical Officers. London: Department of Health 2011.
- 14. Radtke T, Nolan SJ, Hebestreit H, et al. Physical exercise training for cystic fibrosis. *Cochrane Database Syst Rev*. 2015.
- 15. 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.
- 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.
- 17. 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.
- 19. Carson V, Ridgers ND, Howard BJ, et al. Light-intensity physical activity and cardiometabolic biomarkers in US adolescents. *PLoS One*. 2013;8:e71417.
- 20. Schneiderman JE, Wilkes DL, Atenafu EG, et al. Longitudinal relationship between
- physical activity and lung health in patients with cystic fibrosis. *Eur Respir J*.
- 2014;43:817-823.
- 21. Gabel L, Ridgers ND, Della Gatta PA, et al. Associations of sedentary time patterns and TV viewing time with inflammatory and endothelial function biomarkers in children. *Pediatr Obes*. 2016;11:194-201.
- 22. Mark AE, Janssen I. Influence of bouts of physical activity on overweight in youth. *Am J Prev Med*. 2009;36:416-421.
- 23. 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.
- 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;30:629- 633.
- 25. 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.
- 26. Freedson P, Pober D, Janz KF. Calibration of Accelerometer Output for Children.
- *Med Sci Sport Exerc*. 2005;37(11):S523-S530.
- 27. 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.
- 28. Mackintosh KA, Ridley K, Stratton G, et al. Energy Cost of Free-Play Activities in

10- to 11-Year-Old Children. *J Phys Act Health*. 2016;13:S71-74.

- 29. Ryan JM, Forde C, Hussey JM, et al. Comparison of Patterns of Physical Activity and
- Sedentary Behavior Between Children With Cerebral Palsy and Children With
- Typical Development. *Physical Therapy*. 2015;95:1609-1616.
- 30. Mattocks C, Ness AR, Leary SD, et al. Use of accelerometers in a large field-based
- study of children: Protocols, design issues, and effects on precision. *J Phys Activ*
- *Health*. 2008;5:S98-S111.
- 31. Healy GN, Wijndaele K, Dunstan DW, et al. Objectively measured sedentary time,
- physical activity, and metabolic risk: the Australian Diabetes, Obesity and Lifestyle Study (AusDiab). *Diabetes Care*. 2008;31:369-371.
- 32. 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.
- 33. Bradley JM, Moran F. Physical training for cystic fibrosis. *Cochrane Database Syst Rev*. 2008.
- 34. Stevens D, Oades PJ, Armstrong N, et al. A survey of exercise testing and training in UK cystic fibrosis clinics. *J Cyst Fibros*. 2010;9:302-306.
- 35. Williams CA, Saynor ZL, Tomlinson OW, et al. Cystic fibrosis and physiological responses to exercise. *Expert Rev Respir Med*. 2014;8:751-762.
- 36. Ruf KC, Fehn S, Bachmann M, et al. Validation of activity questionnaires in patients with cystic fibrosis by accelerometry and cycle ergometry. *BMC Med Res Methodol*. 2012;12.
- 37. Rowlands AV, Eston RG, Ingledew DK. Measurement of Physical Activity in
- Children with Particular Reference to the use of Heart Rate and Pedometry. *Sports Med*. 1997;24:259-272.
- 38. Bailey RC, Olson J, Pepper SL, et al. The level and tempo of children's physical activities: an observational study. *Med Sci Sport Exerc*. 1995;27:1033-1041.
- 39. Baquet G, Stratton G, Van Praagh E, et al. Improving physical activity assessment in
- prepubertal children with high-frequency accelerometry monitoring: A
- methodological issue. *Prev Med*. 2007;44:143-147.
- 40. Rowland A, Eston RG. The measurement and interpretation of children's physical activity. *J Sport Sci Med*. 2007;6:270-276.



- 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.

462 **Table 1.** Participant characteristics

- 458
- 459
- 460 **Tables**
- 461

# **Total Cystic Fibrosis Controls**



463 Mean  $\pm$  S.D. PHV, peak height velocity; FVC, forced vital capacity; FEV<sub>1</sub>, forced expiratory volume

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

465

# 466 **Table 2.** Physical activity data by group



## Physical Activity Levels in Cystic Fibrosis Youth



467 Means  $\pm$  SD. Low-LPA, low light physical activity; High-LPA, high light physical activity; MPA, 468 moderate physical activity; VPA, vigorous physical activity; VVPA, very vigorous physical activity; 469 MVPA, moderate-to-vigorous physical activity. \* Significant difference between week- and weekend 470 day within group 471 472 473

- 474
- 475



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



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