

ORIGINAL ARTICLE

Comorbidity and physical activity in people with paraplegia: a descriptive cross-sectional study

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Study Design: Descriptive cross-sectional study.

Setting: The study was conducted in the Spinal Cord Injury Unit of the University Vall d'Hebron Hospital and in the Physical Education and Sports Department of the University of Valencia.

Objectives: The aim of this study was to quantify the presence of comorbidities in spinal cord injury (SCI) subjects who did or did not perform regular physical activity (PA) and to identify the relationship between PA and the level of comorbidity.

Methods: The sample consisted of patients with complete motor SCI (T2–T12), who were fitted with an accelerometer attached to the non-dominant wrist for a period of 1 week. The clinical and blood analytic variables were selected by an expert panel.

Results: In the exploratory analysis, we have found differences in the total number of pathologies between active and inactive patients, with fewer total pathologies in the active patient group. An association was found between the PA level and diabetes mellitus ($\chi^2_1 = 3.96$; $P = 0.047$; $\phi = 0.25$). We also observed an association between the cardioprotector level of high-density lipoprotein (HDL)-cholesterol and PA level ($\chi^2_1 = 3.62$; $P = 0.057$; $\phi = 0.24$).

Conclusions: Our results suggest that patients considered active showed lower total comorbidity than inactive patients and higher protection levels against developing cardiovascular comorbidity.

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INTRODUCTION

The normal ageing process combined with the long-term deterioration produced by spinal cord injury (SCI) compromises the patient's general health status and increases the risk of acquiring further disabilities. The decline in cardiovascular function as a result of ageing is more pronounced in people with SCI than in the general population. Moreover, the pathological conditions associated with SCI, such as hyperinsulinemia, obesity and high cholesterol, could aggravate cardiovascular diseases,^{1–3} which are now the first cause of death in the SCI population. In addition, those with complete SCI have abnormal heart rate responses and blood pressure as compared with their healthy peers, which indicates alterations of the autonomous system⁴ and reduced lung capacity, also characteristic of SCI.⁵

As regards the metabolic system, the frequency of glucose intolerance is higher in people with SCI and could be the cause of the higher risk of premature diabetes mellitus,⁶ possibly aggravated by their greater body mass index and central obesity characteristics.

As physical activity (PA) can modulate some of these SCI alterations, it is a relevant factor in primary and secondary prevention of several metabolic and cardiovascular diseases associated with SCI. Indeed, the few epidemiological studies carried out on the SCI population have shown encouraging results in terms of PA.^{7–10} The recommendations relating to exercises designed to improve stamina and maximal oxygen consumption (VO₂max) in people with SCI are not very different from those that have been established for the general

population.^{11–13} These recommendations advise from three to five 20–60-min sessions per week at an intensity of 50–80% of VO₂max.¹⁴

Few studies have evaluated the daily PA in this population and even fewer have attempted to quantify the PA and energy expenditure measured by an accelerometer during a 1-week period. Most of these studies used a questionnaire to ascertain the level of activity,^{15,16} and some monitored the subjects' heart rate and indirect calorimetry during activities performed in a laboratory.^{17,18}

As we considered it important to explore the relationship between PA and health-related factors in SCI subjects, the aim of this study was to quantify the comorbidities in two groups of SCI subjects who either did or did not perform regular PA.

MATERIALS AND METHODS

Participants

A convenience sample of 67 SCI patients was recruited from the database of the Hospital de la Vall d'Hebrón and the Hospital La Fe de Valencia for a cross-sectional study that lasted for a period of 3 months. All the participants had complete traumatic motor SCI with injury levels between T2 and T12 (American Spinal Injury Association Impairment Scale A) and all were full-time manual wheelchair users.

The exclusion criteria were as follows: psychiatric disorder that could interfere with participation, siringomyelia, nervous impairment in upper extremities, ischemic cardiac pathology, recent osteoporotic fracture, mechanical ventilator dependent, acute cancer process or recent ischial, sacral or trochanteric pressure ulcers.

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The epidemiological characteristics of the population are given in Table 1.

All the participants provided a written informed consent, all the procedures were conducted in accordance with the principles of the World Medical Association's Declaration of Helsinki and the protocols were approved by the ethical committee of the Vall d'Hebrón Hospital.

Experimental procedure

All the subjects were clinically examined by a physician and SCI rehabilitation specialist. The body mass index and blood pressure were obtained, and the subjects were asked whether they suffered urinary infections, constipation, dysreflexia symptoms and whether they were in need of bladder voiding. The activity monitor was then attached to their non-dominant wrist¹⁷ and instructions were given regarding its use. All the patients' clinical data were obtained from the database of each hospital.

PA measurement

An Actigraph Model GT3X (Manufacturing Technology Inc., Fort Walton Beach, FL, USA) was used to measure PA by means of the accelerometer worn by the patients. The Actigraph was initialized using 30 Hz frequency, and the data were transformed into counts·seg⁻¹. After 1 week, the participants returned the accelerometer to the hospital. Matlab R2010a (Mathworks Inc., Natick, MA, USA) software was used for preprocessing, segmentation and feature extraction from the signals. All the data over 5000 counts·seg⁻¹ were considered not to have been generated by the subjects and were removed, and any periods of 20 consecutive minutes without registrations were not considered valid. The activity records were considered valid if at least 10 h per day had been obtained in 4 days, of which 3 were during the week and 1 at the weekend.

The mathematical model developed by García-Massó *et al.* was used to analyze the PA.¹⁷ This model, using indirect calorimetry as gold standard, showed an *r*-value of 0.86 and a root mean square error of 2.23 ml kg⁻¹ min⁻¹. The PA levels were divided into sedentary (that is, <1.5 Metabolic equivalents (METS)), low (that is, 1.5–2.99 METS), moderate (that is, 3–5.99 METS) and vigorous (that is, >6 METS).

In accordance with the literature,^{15,19} the sample was divided into two groups: the subjects who performed at least 60 min of PA per week at the moderate–vigorous level were considered active (*n* = 37) and those who did less than 60 min were considered inactive (*n* = 30).

Table 1 Participants' clinical profile

	Inactive (<i>n</i> = 30)	Active (<i>n</i> = 37)	Mean between-group difference with 95% CI	P-value
<i>Gender</i>				
Male	20 (66.7)	31 (83.8)	—	—
Female	10 (33.3)	6 (16.2)	—	—
<i>Neurological level</i>				
≤T6	14 (46.7)	21 (56.8)	—	—
>T6	16 (53.3)	16 (43.2)	—	—
Age (years)	50.63 (14.12)	43.3 (12.16)	7.34 (0.92 to 13.75)	0.02
Weight (kg)	73.85 (12.78)	70.81 (13.0)	3.04 (−3.29 to 9.37)	0.34
Height (m)	1.7 (0.08)	1.73 (0.1)	−3.37 (−7.89 to 1.16)	0.14
BMI (kg m ⁻²)	25.67(4.48)	23.57 (3.51)	2.10 (0.15 to 4.05)	0.03
Time since injury (years)	15.77 (11.71)	17.76 (11.6)	−1.99 (−7.71 to 3.73)	0.48

Abbreviations: BMI, body mass index; CI, confidence interval; T6, 6th thoracic level. Data are expressed as mean (s.d.) or mean (95% CI). For gender and neurological level, the number of participants is shown *N* (%).

Medical data

A panel of experts selected the medical variables following the European Cooperation Office (EuropeAid) recommendations, after a review of the main scientific databases (for example, WOS, Pubmed, Cochrane and so on).

The variables selected were as follows: neurological level, time from injury, hypertension, dyslipidemia, diabetes mellitus, cardiological or pulmonary diseases, metabolic diseases, vascular problems, psychological disorders, fractures and other musculoskeletal pathologies, allergies, pressure ulcers and urinary infections that required admission to hospital. The subject's lipid profile was measured to assess the risk of cardiovascular disease. Total cholesterol (TC) and triglycerides (TGs) were determined using standardized enzymatic procedures. High-density lipoprotein (HDL) and low-density lipoprotein (LDL) were selectively precipitated and the TC/HDL ratio was calculated. Total proteins and albumin were studied.

The metabolic syndrome was then assessed following the criteria of the American Association of Clinical Endocrinologists.²⁰

Statistical analysis

The statistical analyses were performed on SPSS 20.0 software (IBM Corporation, Armonk, NY, USA). The mean and s.d. were used to describe data. Although the main goal of this study was descriptive, we also explored the possible differences between physically active and non-active people. To this purpose, the mean differences between groups and the 95% confidence interval were computed. Further, the Mann–Whitney *U*-test was conducted. Categorical variables were tested using the χ^2 -test and effect size ($\phi = \sqrt{\chi^2/N}$).²¹ Besides, the odds ratios (ORs) were reported.

RESULTS

General results of the participants

The clinical profiles of the participants are shown in Table 1. The active group was slightly younger and taller and showed a lower BMI than the inactive group.

The mean weekly time for each PA level and sedentary activity level of all the participants can be seen in Table 2. As we had expected, the active patients performed more minutes of light, moderate, vigorous, moderate-to-vigorous activity and had higher average METs than the inactive patients. In fact, this difference is greater in METs and moderate-to-vigorous weekly PA variables.

The most frequent comorbidities associated with SCI were as follows: urinary tract infections requiring hospital admission (52%), pressure ulcer (39%) and dyslipidemia (22%; see Figure 1).

PA and comorbidity

We found differences between the active and inactive groups in the total number of pathologies, with fewer comorbidities in the active group. Each pathology was analyzed separately and a statistical association was found between PA group and diabetes mellitus ($\chi^2_1 = 3.96$; $P = 0.047$; $\phi = 0.25$). On the basis of the OR, the active subjects were 7.2 times less likely to suffer non-insulin-dependent diabetes mellitus than inactive subjects.

The metabolic syndrome was assessed through American Association of Clinical Endocrinologists criteria. Sixty-three percent of the inactive patients presented two or more risk factors for metabolic syndrome, in contrast to the 59% of the active patients who presented none or only one risk factor.

Blood variable values were similar in both groups (Table 3). The HDL-cholesterol >45 mg dl⁻¹ was associated with a higher PA level ($\chi^2_1 = 3.62$; $P = 0.057$; $\phi = 0.24$). On the basis of the OR, the odds of a subject having more than 45 mg dl⁻¹ of HDL were 2.86 times higher if they were physically active than inactive.

Table 2 Daily physical activity of participants and sedentary behavior

	Inactive (n = 30)	Active (n = 37)	Mean between-group difference with 95% CI	P-value
METS	1.44 (0.82)	1.57 (0.1)	-0.13 (-0.18 to -0.09)	<0.001
Sedentary (min per day)	631.05 (184.25)	653.21 (192.6)	-22.16 (-114.85 to 70.54)	0.64
Light (min per day)	239.69 (82.86)	335.43 (104.62)	-95.73 (-142.60 to -48.86)	<0.001
Moderate (min per day)	3.86 (3.01)	26.75 (19.64)	-22.89 (-29.52 to -16.26)	<0.001
Vigorous (min per day)	0.01 (0.04)	0.22 (0.54)	-0.21 (-0.39 to -0.03)	0.04
Moderate-to-vigorous (min per day)	3.87 (3.0)	26.97 (19.96)	-23.10 (-29.83 to -16.37)	<0.001

Abbreviation: CI, confidence interval.
Data are expressed as mean (s.d.) or mean (95% CI).

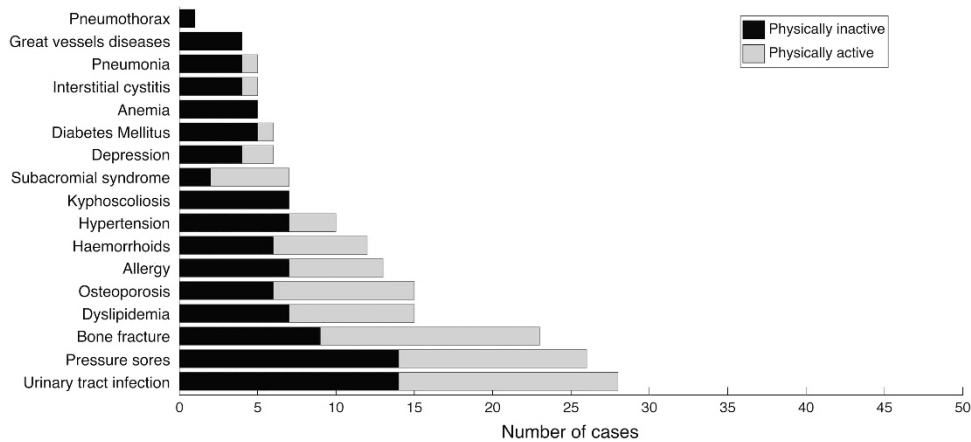


Figure 1 Number of cases of each comorbidity for active and inactive subjects.

Table 3 Blood analytical variables for lipid profile, total proteins and albumin, stratifying between active and inactive group

	Inactive (n = 30)	Active (n = 37)	Mean between-group difference with 95% CI	P-value
Total cholesterol (mg dl ⁻¹)	182.04 (30.22)	185.96 (36.22)	-3.92 (-20.52 to 12.68)	0.53
HDL-C (mg dl ⁻¹)	49.36 (7.7)	50.64 (9.58)	-1.28 (-5.59 to 3.03)	0.33
LDL-C (mg dl ⁻¹)	117.68 (26.63)	121.03 (30.04)	-3.35 (-17.37 to 10.66)	0.48
Triglycerides (mg dl ⁻¹)	133.12 (50.27)	120.86 (61.46)	12.26 (-15.58 to 40.10)	0.23
Proteins (g dl ⁻¹)	7.14 (0.37)	10.72 (15.72)	-3.58 (-8.82 to 1.66)	0.83
Albumin (g dl ⁻¹)	4.33 (0.27)	4.31 (0.37)	0.01 (-0.15 to 0.17)	0.70
TC/HDL-C	3.79 (0.94)	3.84 (1.25)	-0.05 (-0.60 to 0.50)	0.74

Abbreviations: CI, confidence interval; HDL-C, high-density lipoprotein cholesterol; LDL-C, low-density lipoprotein cholesterol; TC, total cholesterol.
Data are expressed as mean (s.d.) or mean (95% CI).

DISCUSSION

The results obtained suggest that the subjects with an inactive life-style suffered a higher number of SCI-associated comorbidities than active SCI subjects. Although the study design used could not establish causality, an association was discovered between performing PA and suffering from diabetes mellitus. In addition, there were more patients with metabolic syndrome in the inactive group. A level above 45 mg dl⁻¹ HDL, which is considered cardioprotective, could be associated with PA.

We found that a high percentage of the paraplegic population suffered from comorbidities associated with SCI. The most frequent of these were as follows: urinary infection requiring hospital admission (52%), pressure ulcers (39%) and dyslipidemia (22%). These results are similar to those obtained for this population in other studies,²²⁻²⁴ for example, Brinkhof *et al.* found a prevalence of the urinary tract infection and pressure ulcers of 59% and 34%, respectively.²⁵

The average PA value was 1.51 METs, which is a relatively low level of energy expenditure. The patients were involved in low- or very-low-

intensity activities most of the time, although 55% of the subjects fulfilled the PA recommendations to be considered physically active (at least 60 min activity per week of moderate-to-vigorous intensity).

Diabetes and the metabolic syndrome have been strongly associated with cardiovascular diseases.^{26,27} In 1994, Bauman and Spungen²⁸ identified higher glucose and insulin blood levels in an SCI population than in a control group. Insulin sensitivity has been observed in two-thirds of the tetraplegic population and in 50% of paraplegics. Similarly, Lee *et al.* detected the metabolic syndrome in 23% of people with SCI, approximately double the incidence in the control group stratified by gender and age,²⁹ in agreement with the work done by Jones *et al.*³⁰ This suggests that there is a relationship between metabolic syndrome and SCI.

PA has been found to act as a protecting factor against the metabolic syndrome.³¹⁻³⁴ Manns *et al.* found that paraplegic patients with a low level of cardiorespiratory fitness who did not exercise had a higher fasting glucose blood level, lower HDL-cholesterol levels and a larger waist circumference.³² The authors estimated cardiorespiratory

fitness by measuring VO₂max during cardiopulmonary exercise testing with arm cycling and a self-assessment questionnaire. One possible mechanism may be due to premature tissue ageing associated with low levels of the insulin-like growth factor 1 (IGF-1) hormone and other growth factors.²⁸ Inactivity involves a higher physiopathological risk of suffering sarcopenia, hyperglycemia, diabetes and cardiovascular diseases. Although we could not find any relationship between PA and the metabolic syndrome, this may be possible in future studies with a higher sample size.

The relationship between an altered lipid profile and having diabetes or cardiovascular disease has already been well established.³⁵ Some studies have found higher LDL cholesterol and TC levels and lower HDL-cholesterol levels after SCI than in the non-disabled population.^{26,28,32,36,37} In line with the findings of other studies^{31,37} we did not find any relationship between the TC or LDL cholesterol level and physically active subjects. Although inactive paraplegics presented pathological TG levels (10%), we did not find a relationship between these two factors. In the able-bodied population higher cholesterol and TG levels may be modified by PA and dieting.³⁵ There is also a strong relationship between time from SCI and a high lipid profile in the SCI population,³⁸ which suggests that the metabolic changes and physical inactivity associated with SCI may have significant consequences for their lipid profile.

On the other hand, the more active patients had slightly higher HDL-cholesterol levels. The SCI subjects who exercised for 60 min per week (with moderate-to-vigorous intensity) were 2.86 times more likely to present HDL-cholesterol in the cardioprotective range (over 45 mg dl⁻¹). In agreement with our findings, Groot *et al.*³⁷ found a significant relationship between higher PA levels and higher HDL-cholesterol, but not in TC, LDL cholesterol and TG. Brenes *et al.*³⁹ used a questionnaire to ascertain activity level and found significantly lower HDL-cholesterol (34.2 mg dl⁻¹) in 66 sedentary paraplegics as compared with 22 complete SCI subjects with athletic inclinations (47.1 mg dl⁻¹).

Some studies measured daily PA solely by means of questionnaires, or only considered practising athletes as the active SCI population, and thus were not representative of the total SCI population. To our knowledge, only Nooijen *et al.*³⁴ studied lipid profile and PA by accelerometry for two consecutive days. In this study, with a mix of paraplegics and tetraplegics, of which 72% had complete SCI, a significant relationship was found between higher TG levels and physical inactivity.

In the present study, we considered active patients as those who performed at least 60 min of moderate-to-vigorous PA per week, and this population had significantly higher HDL-cholesterol (more than 45 mg dl⁻¹), which suggests that this PA level performed weekly could raise cardioprotective HDL-cholesterol levels.

Regarding diabetes, Cragg *et al.*⁴⁰ found that the OR of Type 2 diabetes was 2.52 times greater in individuals with SCI versus individuals without SCI (95% CI 1.81–3.52). Buchholz *et al.*⁷ in a sample of SCI found that the subjects who performed at least 25 min of physical exercise per day had lower insulin resistance (10%) than inactive subjects (33%). Although we found a lower incidence of Type 2 diabetes than other studies in the literature, there is a relationship between being considered inactive and suffering diabetes after SCI. In our study, the patients considered inactive presented a 7.2 times higher risk of getting Type 2 diabetes than active patients.

One mechanism could be explained by the relationship between sarcopenia, present in the SCI population, and insulin resistance.⁴¹ Moreover, as the SCI population have an altered autonomous nervous

system and vascular system, this could be responsible for the higher metabolism dysfunction.⁴⁰

The primary limitation of the present study is the small number of subjects in the sample, which means a Type II error could have been committed and further associations may not have been detected between PA and the comorbidities studied. Secondly, the age difference between groups (being younger the active people with SCI) could have influenced comorbidity as well as PA. In addition, it should be stressed that the grouping criterion used in this study (that is, 60 min per week of physical exercise) is extensively documented in studies in which the questionnaires are used to assess the PA but not in those whose measurement system is based on accelerometer devices.

At a practical level, further data need to be acquired in observational studies to reach epidemiological conclusions transferrable to the SCI population. Experimental studies with a higher control of confounding variables are also needed in order to establish causal relationships.

CONCLUSIONS

Our results suggest that active people with SCI present lower total comorbidity than inactive people with SCI. In addition, the patients considered active showed improved protection against developing cardiovascular comorbidity due to higher levels of HDL-cholesterol in the cardioprotective range. Our exploratory analysis showed that the active SCI population was also found to have a lower risk of contracting Type 2 diabetes.

CONFLICT OF INTEREST

The authors declare no conflict of interest.

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