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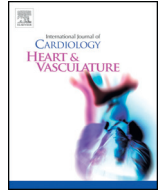
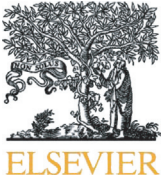
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Geographical variation and predictors of physical activity level in adults with congenital heart disease

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Abbreviations: APPROACH-IS, assessment of patterns of patient-reported outcomes in adults with congenital heart disease – international study; CHD, congenital heart disease; CI, confidence interval; HBS-CHD, health behaviour scale; METS, metabolic equivalents; NYHA, New York Heart Association (class); OR, odds ratio; PAL, physical activity level; PRO, patient-reported outcomes; WHO, World Health Organization.

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ABSTRACT

Background: Physical activity is important to maintain and promote health. This is of particular interest in patients with congenital heart disease (CHD) where acquired heart disease should be prevented. The World Health Organization (WHO) recommends a minimum of 2.5 h/week of physical activity exceeding 3 metabolic equivalents (METS) to achieve positive health effects. It is unknown whether physical activity levels (PAL) in adult CHD patients differ by country of origin.

Methods: 3896 adults with CHD recruited from 15 countries over 5 continents completed self-reported instruments, including the Health Behaviour Scale (HBS-CHD), within the APPROACH-IS project. For each patient, we calculated whether WHO recommendations were achieved or not. Associated factors were investigated using Generalized Linear Mixed Models.

Results: On average, 31% reached the WHO recommendations but with a great variation between geographical areas (India: 10%–Norway: 53%). Predictors for physical activity level in line with the WHO recommendations, with country of residence as random effect, were male sex (OR 1.78, 95%CI 1.52–2.08), NYHA-class I (OR 3.10, 95%CI 1.71–5.62) and less complex disease (OR 1.46, 95%CI 1.16–1.83). In contrast, older age (OR 0.97, 95%CI 0.96–0.98), lower educational level (OR 0.41, 95%CI 0.26–0.64) and being unemployed (OR 0.57, 95%CI 0.42–0.77) were negatively associated with reaching WHO recommendations.

Conclusions: A significant proportion of patients with CHD did not reach the WHO physical activity recommendations. There was a large variation in physical activity level by country of origin. Based on identified predictors, vulnerable patients may be identified and offered specific behavioral interventions.

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1. Introduction

Due to improvements in the treatment and management of congenital heart disease (CHD), most children with congenital heart disease are expected to reach adulthood and the population of adults with CHD continues to grow [1,2]. However, the risk of complications increases as patients grow older [3]. With increasing age, there is also the risk of acquired heart disease, especially in those with traditional risk factors for cardiovascular disease such as hypertension, diabetes, and hyperlipidaemia [4,5]. In an adult CHD population, prevention of acquired heart disease is especially important given the risks associated with re-intervention [6,7] and pre-existing limitations in physical capacity [8].

A physically active lifestyle has the potential to modify cardiovascular risk factors and promote general health [9–11]. Most patients with CHD experience some degree of limitations in aerobic capacity, most pronounced in those with complex heart lesions [8]. This may pose barriers for being physically active. However, studies have suggested that adults as well as children with CHD are physically active on the same level as healthy subjects [12,13]. Nevertheless, approximately one-half to three-quarters of both patients and healthy subjects do not reach the World Health Organization (WHO) recommendations of 2.5 h per week of physical activity of 3 metabolic equivalents (METS) or more [12,14].

Several patient-related factors are potentially associated with low physical activity level (PAL) in patients with CHD, such as reduced aerobic capacity [8,15], impaired muscle function [16–18], self-concept [19], self-efficacy [20], parental overprotection [21], and restriction recommendations by their cardiologists [22]. In general, physical activity level may also be affected by external factors such as seasonal variation [23–25], socio-economic and local environmental factors [26–29], and by country of origin [30]. These findings raise the question whether the degree of physical activity level also varies in different countries in patients with CHD. In the present study, physical activity level was analyzed in a large international cohort of adults with CHD, including geographical variation in physical activity level and general predictors of physical activity level in this particular population.

2. Methods

2.1. Patients and procedure

In total, 4028 adults with CHD from 15 countries in 5 continents participated in the cross-sectional study APPROACH-IS (Assessment of

Patterns of Patient-Reported Outcomes in Adults with Congenital Heart disease – International Study) [31,32] on Patient-Reported Outcomes (PRO). Data were collected from April 2013 to March 2015. Informed consent was obtained from all participants. The study was conducted in accordance with the Declaration of Helsinki. The rationale, design and methodology of APPROACH-IS have been published previously [33].

Patients included in the study met the following criteria: (i) diagnosis of CHD, defined as a structural abnormality of the heart or intra-thoracic great vessels, that was present at birth and had actual or potential functional significance [34]; (ii) 18 years of age or older; (iii) diagnosis established before adolescence (i.e. before 10 years of age); (iv) continued follow-up at a CHD center or included in a national/regional register; and (v) physical, cognitive, and language capabilities necessary to complete self-report questionnaires. Exclusion criteria were prior heart transplantation or primary pulmonary hypertension [33]. The complexity of the congenital heart disease was based on the Bethesda classification [35].

2.2. Measurements

Socio-demographic variables were patient-reported. The self-report questionnaires in APPROACH-IS were administered to eligible patients by surface mail or in clinic during an outpatient visit. The questionnaires have been validated and reliability-tested and measure PROs within different PRO domains, including health behaviour. Data regarding the participants' medical background, such as CHD diagnosis and disease complexity [35], were added to the APPROACH-IS database by a member of the local research team and based on chart review [33].

The Health Behaviour Scale (HBS-CHD), including data on alcohol consumption, tobacco use and physical activity [36], was used to measure physical activity level. The instrument was translated into Chinese, French, German, Hindi, Italian, Japanese, Norwegian, Spanish, Swedish, and Tamil. The questionnaire has a good to excellent content validity and responsiveness [36]. The validity across different languages has not been tested. The HBS-CHD included questions regarding extremely and moderately demanding physical activity during a 7-day week, also including sports during school hours (the latter was relevant for a minority of the studied patients). The number of hours per week spent at an activity ≥ 3 METS and ≥ 6 METS was summarized. Based on the current WHO recommendations on physical activity for promoting health in adults aged 18–64 (i.e. 150 min/week spent ≥ 3 METS or 75 min/week spent ≥ 6 METS or an equivalent combination of both), participants

were dichotomized into two categories: high physical activity level (reaching WHO recommendations) and low physical activity level (not reaching WHO recommendations) based on their participation in physical exercise.

2.3. Statistical analyses

All analyses were performed using SPSS 23 (IBM, Armonk, NY, USA). Data were assessed for normality. Differences in means were tested with Student's *t*-test and ratios with χ^2 -test. The null hypothesis was rejected for *p*-values < 0.05.

The association between patient-specific variables, being age, sex, educational level, employment status, marital status, functional class and disease complexity, versus physical activity level was estimated through generalized linear mixed models that is a form of multilevel logistic regression. We applied a two-level structure in which patients were nested within countries. Hence, all patient characteristics available, which have been used in prior APPROACH-IS reports [31] were used as fixed effects. Country was used as random effect. Generalized linear mixed models do not result in a normally interpretable R^2 statistics. Therefore, we computed the pseudo R^2 using the method described by Nakagawa & Schielzeth [37].

For the physical activity level, full data on 3896 patients was available. For the predictors, missing values occurred in 0.0 to 2.2% of the subjects. Altogether, full data on patient characteristics was available for 3727 (95.7%) of the patients. Therefore, multiple imputation was not used and only patients for whom full data was available for the variables under study were included in the generalized linear mixed models.

3. Results

Out of 4028 participants, 3896 had data on physical activity level. 1217 (31%) reached WHO recommendations on physical activity level. The proportion of patients reaching recommended physical activity level varied among countries ($p < 0.001$), with the lowest proportions in India (10%) and Japan (11%) and highest in Norway (53%), Switzerland (47%), and Sweden (46%). However, the variation was large, also between adjacent countries e.g. France (19%) and Switzerland (Table 1, Fig. 1).

More men than women reached WHO recommendations on physical activity level (37% vs. 26%, $p < 0.001$). Patients with a high physical activity level were younger (32 vs. 36 years, $p < 0.001$), had less complex heart lesions (35% among patients with simple lesions vs. 26% among patients with complex lesions), and higher educational levels (39% of those with a university degree vs. 16% of those who did not finish

high school). Employment status was also associated with physical activity level. Of full-time students, 41% reached WHO recommendations on physical activity level vs. 14% of those who were homemakers or retired. There was an association between self-reported limitations and low physical activity level, with 40% of those who reported no limitations meeting WHO recommendations on physical activity level compared to 10% among those with severe limitations (Table 2).

In a multilevel logistic model with geographical area as a random factor, male sex (OR 1.78, 95% CI 1.52–2.08), NYHA I (OR 3.10, 95% CI 1.71–5.62) and less complex disease (OR 1.46, 95% CI 1.16–1.83) were positively associated with reaching WHO recommendations, whereas higher age (OR 0.97, 95% CI 0.96–0.98), lower educational level (0.41, 95% CI 0.26–0.64) and being unemployed (OR 0.57, 95% CI 0.42–0.77) were negatively associated. Marital status was not associated with physical activity level (Table 3). The pseudo R^2 for this multilevel model was 0.464.

4. Discussion

In the present study, we found that in a globally recruited sample, approximately one third of adults with CHD reached the WHO recommendations on physical activity to maintain or promote health. However, large geographical variations from 10% to slightly above 50% of the population reaching the current recommendations were seen. In a multilevel logistic regression model with geographical area as random effect, sex, age, educational level, employment status, complexity of heart lesions, and self-reported NYHA class were associated with reaching WHO recommendations on physical activity level. This knowledge may help in detecting vulnerable patients and thereby offer specific behavioral interventions.

The reasons for the large variation in physical activity level between different geographical regions are not clear. Factors such as climate, cultural variations, infrastructure and socioeconomic factors may be of importance. With a few exceptions, our studied population reached the recommendations on physical activity level to a similar extent as the reference general population in their respective countries [30,38–40].

In our study we demonstrated that men were more likely to be sufficiently active, which is in line with previous studies on adults with CHD [12,14] as well as the general population [41]. However, there are also conflicting reports on adults with CHD [42] where gender was not associated with physical activity level. The difference in physical activity level between the sexes persisted when adjusted for geographical area. Given this, health care providers should not only ensure activity recommendations are provided for all patients but additional efforts should be made to educate their female CHD patients about the importance of physical activity and the potential long-term benefits.

In this study, the odds for reaching the WHO recommendations on physical activity level decreased by 3% each year of life. It is known that physical activity level decreases with age in the general population [41] and others have reported consistent results in adults with congenital heart disease [42]. A reduction in exercise capacity, which is found in the general population [43] as well as in adults with CHD [8], was a possible explanatory mechanism for lower physical activity level with increasing age. Our data supported that a decreased physical activity level due to increasing age, even at relatively young ages, is a global phenomenon in adults with CHD. These observations underscore the importance of addressing physical activity level in the management of older patients with CHD.

We found that the complexity of CHD was associated with physical activity level. This finding contrasts with previous reports [12,42,44]. In two previous studies, a slightly different definition of complexity was applied with two groups of complexity instead of three as used in this present investigation [35]. In our study, we showed that patients with simple and moderate lesions had higher physical activity level than patients with lesions of severe complexity. However, we noted that differences were modest and the point estimates were fairly similar

Table 1
Distribution of PAL in the different countries.

Country	All patients n = 3896	High PAL n = 1217	Low PAL n = 2679	<i>p</i>
Norway	172 (4)	91 (53)	81 (47)	<0.001
Switzerland	270 (7)	128 (47)	142 (53)	
Sweden	463 (12)	213 (46)	250 (54)	
The Netherlands	250 (6)	99 (40)	151 (60)	
Italy	60 (2)	24 (40)	36 (60)	
Australia	130 (3)	51 (39)	79 (61)	
Canada	504 (13)	157 (31)	347 (69)	
Belgium	263 (7)	76 (29)	187 (71)	
Malta	118 (3)	31 (26)	87 (74)	
USA	726 (19)	190 (26)	536 (74)	
Argentina	168 (4)	44 (26)	124 (74)	
Taiwan	248 (6)	51 (21)	197 (79)	
France	88 (2)	17 (19)	71 (81)	
Japan	237 (6)	26 (11)	211 (89)	
India	199 (5)	19 (10)	180 (91)	

Data shown as n (%). *p*-Value represents comparison in physical activity levels between countries.

n = number; PAL = physical activity level.

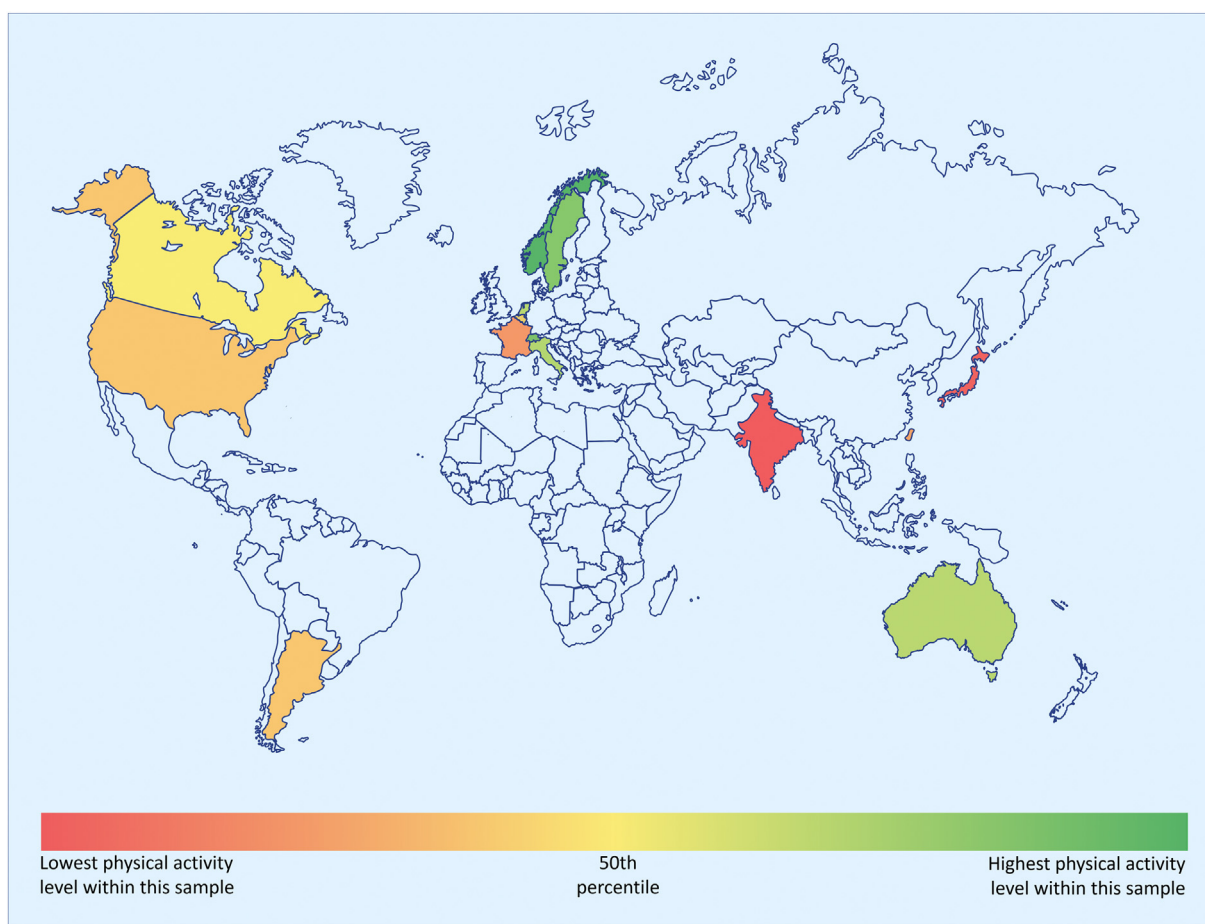


Fig. 1. Comparison of PAL between countries. PAL in adults with CHD was measured using the Health Behaviour Scale (HBS-CHD) and the proportion reaching WHO recommendations on PAL was calculated for each country. There was a great inter-country variation in the proportion of CHD patients reaching recommended PAL. In this heat map, countries included in the study are marked in color. Shades of green denote a higher proportion of CHD patients reaching recommended PAL as compared with shades of red. CHD = congenital heart disease, PAL = physical activity level, WHO = World Health Organization.

for both groups. It may be that patients with lesions of severe complexity in our present study represented more severely limited patients compared with the previous studies. It may also be that the large sample size in this study allowed for detection of smaller differences between groups. Nevertheless, the complexity of the CHD lesion should be considered when giving advice on physical activity to patients with CHD [45,46].

The self-reported NYHA class was strongly associated with physical activity level with almost three times higher odds of reaching recommendations on physical activity level for patients in NYHA class I in contrast to the higher NYHA classes (III and IV). For NYHA class II, the point estimates were in line with NYHA class I but did not reach statistical significance. Our results were in agreement with a previous study that reported higher activity levels for patients in NYHA class I [47]. It was not surprising that patients without limitations are more active than those with limitations for physical activity. For patients describing themselves as physically limited, this is a very strong indicator of having a low physical activity level. These patients are at potential risk of developing complications related to low physical activity and may thus be trapped in a vicious cycle. Patients with higher NYHA classes should be assessed carefully regarding actual physical activity level and offered targeted advice and rehabilitation measures [45].

As in the general population [48–50], higher educational levels were associated with higher physical activity level among patients with CHD. Educational level is likely associated with employment status, which

was also associated with high physical activity level in the current study. Both educational level and employment status are possible to modify and caregivers can support these efforts with discussions beginning in early adolescence on future education, vocation and employment.

Exercise training in adults with congenital heart disease has been proved safe in several trials. There are also recommendations for exercise training that can be applied on an individual basis, taking in account factors such as arrhythmia, arterial saturation and ventricular function [46].

4.1. Study limitations

The present study population only captured patients with CHD and reference data were lacking. While a reference population were not feasible for the current study, the current methods allowed for identification of those with CHD who may be more or less likely to reach WHO recommendations on physical activity level. Only self-reported instruments were used which have inherent limitations regarding under- or overestimation, recall bias, and social desirability bias. On the other hand, only validated instruments were used and our sample was large, which hopefully allowed for a valid output. We did not have access to data on medication. Some common drugs in cardiovascular therapeutics, e.g. beta-blockers, may affect physical performance and thereby also potentially the physical activity level. Our sample was not

Table 2
Overview of patient data.

Variables		Data points (n)	All patients n = 3896	High PAL n = 1217	Low PAL n = 2679	p
Sex		3880				<0.001
Female	n (%)		2035 (52)	524 (26)	1511 (74)	
Male	n (%)		1845 (48)	685 (37)	1160 (63)	
Age years	Mean ± SD	3890	35 ± 13	32 ± 11	36 ± 13	<0.001
Education		3860				<0.001
Less than high school	n (%)		204 (5)	32 (16)	172 (84)	
High school	n (%)		1663 (43)	483 (29)	1180 (71)	
College degree	n (%)		824 (21)	241 (29)	583 (71)	
University degree	n (%)		1169 (30)	454 (39)	715 (61)	
Work situation		3874				<0.001
Part-time or full-time work	n (%)		2493 (64)	861 (35)	1632 (66)	
Homemaker or retired	n (%)		299 (8)	42 (14)	257 (86)	
Job seeking, unemployed, or disabled	n (%)		489 (13)	79 (16)	410 (84)	
Full-time student	n (%)		319 (8)	130 (41)	189 (59)	
Other	n (%)		274 (7)	99 (36)	175 (64)	
Marital status		3878				0.048
Never married	n (%)		1716 (44)	567 (33)	1149 (67)	
Divorced or widowed	n (%)		192 (5)	46 (24)	146 (76)	
Married or living with partner	n (%)		1964 (51)	598 (30)	1366 (70)	
Other	n (%)		6 (0)	2 (33)	4 (67)	
Limitations in physical functioning		3811				<0.001
Not limited during physical activities	n (%)		2052 (54)	829 (40)	1223 (60)	
Slightly limited during physical activities	n (%)		1334 (35)	323 (24)	1011 (76)	
Considerably limited during physical activities	n (%)		277 (7)	30 (11)	247 (89)	
Unable to be physically active without experiencing discomfort	n (%)		148 (4)	14 (10)	134 (91)	
Complexity of heart defect		3896				<0.001
Simple	n (%)		1003 (26)	354 (35)	649 (65)	
Moderate	n (%)		1891 (49)	598 (32)	1293 (68)	
Complex	n (%)		1002 (26)	265 (26)	737 (74)	

p-Values represent comparisons of physical activity levels between different patient variables.
n = number; PAL = physical activity level.

Table 3
Multivariable generalized linear mixed models with patient characteristics as predictors of PAL (n = 3727).

Variables	PAL	p
Sex		
Female	#	
Male	1.78 (1.52–2.08)	<0.001
Age years	0.97 (0.96–0.98)	<0.001
Education		
Less than high school	0.41 (0.26–0.64)	<0.001
High school	0.59 (0.49–0.71)	<0.001
College degree	0.70 (0.57–0.87)	<0.05
University degree	#	
Work situation		
Part-time or full-time work	#	
Homemaker or retired	0.76 (0.51–1.12)	N.S.
Job seeking, unemployed, or disabled	0.57 (0.42–0.77)	<0.001
Full-time student	1.19 (0.90–1.58)	N.S.
Other	1.13 (0.83–1.53)	N.S.
Marital status		
Never married	1.06 (0.88–1.27)	N.S.
Divorced or widowed	1.01 (0.69–1.49)	N.S.
Married or living with partner	#	
Other	0.55 (0.05–6.19)	N.S.
Limitations in physical functioning		
Not limited during physical activities	3.10 (1.71–5.62)	<0.001
Slightly limited during physical activities	1.76 (0.97–3.21)	N.S.
Considerably limited during physical activities	0.85 (0.42–1.71)	N.S.
Unable to be physically active without experiencing discomfort	#	
Complexity of heart defect		
Simple	1.46 (1.16–1.83)	<0.05
Moderate	1.25 (1.03–1.52)	<0.05
Complex	#	

Data presented as Odds Ratios (Confidence Intervals).

N.S. = non-significant; PAL = physical activity level.

Reference category.

population based and not randomly selected. However, the large sample of adults with CHD and the multicenter recruitment of patients hopefully allowed for valid conclusions based on the present data.

5. Conclusions

Almost 70% of adults with CHD did not reach the WHO recommended physical activity level. There was a large variation between countries in the proportion of patients that reached recommended physical activity level, from 10% to slightly above 50%. Given the proportion of patients not reaching recommended physical activity level, many are at potential risk for developing long-term complications related to a low physical activity level. Therefore, issues regarding physical activity level should be encouraged and discussed in all consultations in adults with congenital heart disease. Furthermore, the identified predictors of physical activity level may help to identify vulnerable patients and thereby allow for targeted interventions.

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Declarations of interest

None.

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