

Association between long-term smoking and leisure-time physical inactivity - a cohort study among Finnish twins with a 35-year follow-up

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Authorship statement

All authors meet the ICMJE authorship requirements as follows: 1) substantial contributions to conception and design (MP, JK, KS, PS, TK, AR) OR the acquisition of data (JK), AND the analysis and interpretation of data (all authors: MP, JK, KS, PS, TK, AR); AND 2) the drafting of the article or its critical revision for important intellectual content (all authors: MP, JK, KS, PS, TK, AR); AND 3) final approval of the version to be published (all authors: MP, JK, KS, PS, TK, AR). All authors (MP, JK, KS, PS, TK, AR) also agreed be accountable for all aspects of the work in ensuring that questions related to the accuracy or integrity of any part of the work are appropriately investigated and resolved.

ABSTRACT

Objectives

To investigate longitudinal associations of smoking and a change in smoking status with leisure-time physical inactivity. In addition, to control whether familial confounding (genetics and shared environment) influences the associations.

Methods

Data were based on the population-based Finnish Adult Twin Cohort of 5254 twin individuals born in 1945-1957 (41% men) and who participated in all four surveys over a 35-year follow-up (1975–2011). Logistic and conditional logistic regression models with multiple covariates were used for analyses.

Results

Compared to never-smokers, long-term daily smokers (1975–1990) had the highest likelihood for both long-term inactivity and to change into inactive by 2011. Recurrent smoking was associated with long-term inactivity. Instead, in comparison to persistent daily smokers, quitting smoking decreased the likelihood of becoming physically inactive at leisure-time. The associations remained in the analyses which accounted for multiple covariates and/or familial confounding.

Conclusions

Daily smoking increases the likelihood of remaining or becoming physically inactive over the decades. Our results emphasize the importance to prevent smoking initiation but also to support early smoking cessation in promotion of lifelong physical activity.

Keywords: Adults; Follow-up cohort study; Exercise; Physical inactivity; Smoking; Twins

INTRODUCTION

Physical inactivity and tobacco smoking are major preventable risk factors for non-communicable diseases, including cardiovascular diseases, chronic respiratory diseases, diabetes, and many cancers (Lee et al. 2012; Lim et al. 2012). Both contribute to increasing health care costs (Krueger et al. 2014) and shortened lifespans (Lim et al. 2012). Globally, around one-third of adults are reported as being physically inactive during their leisure time (World Health Organization 2011), whereas more men (31%) than women (6%) are daily smokers (Ng et al. 2014). Even though age-adjusted prevalence rates of daily smoking (Ng et al. 2014) and leisure-time physical inactivity (Borodulin et al. 2016) have shown decreasing trends over recent decades in many high-income countries, the prevalence of smoking and inactivity is still high or is even increasing in other countries (Dumith et al. 2011; Ng et al. 2014).

Smoking often coexists with other unhealthy lifestyle habits (Auer et al. 2014; Morris et al. 2016). Accumulating evidence from longitudinal studies indicates that adults who are daily smokers are less physically active in their leisure time than non-smokers (Auer et al. 2014; Azagba and Asbridge 2013; Paavola et al. 2004). However, the association between smoking and leisure time physical inactivity has not been consistent (Bauman et al. 2012; Kaczynski et al. 2008). More longitudinal studies with a follow-up over several decades and several time points to track changes in smoking and leisure-time physical activity (LTPA) are needed to establish whether a change in smoking behavior would affect physical activity behavior.

Both smoking (Li et al. 2003; McGue et al. 2014) and LTPA (de Geus et al. 2014; de Vilhena e Santos et al. 2012) are known to have a moderate-to-strong genetic component. Studies about the familial confounding underlying smoking and physical inactivity are, however, rare. Twin studies

provide possibility to a co-twin control design which applies determine whether the association between the factors is influenced by familial confounding (McGue et al. 2010). All twin pairs raised together have the same family background and thus share many environmental exposures in childhood. Dizygotic (DZ) twin pairs also share, on average, 50% of their segregating genes, whereas monozygotic (MZ) twin pairs are virtually identical on the gene sequence level (Boomsma et al. 2002). Twin pairs discordant for leisure-time physical inactivity (one twin within a pair is active and the co-twin inactive) would provide the possibility to determine whether the association between smoking and inactivity is influenced by familial confounding. If familial factors play a role, the association between smoking status and inactivity should exist in the analyses of the whole cohort (i.e. twins treated as individuals) but not among pairs discordant for inactivity. If genetics plays a role, then the association should be present within DZ twin pairs, but not within MZ twin pairs. Further, if the association were found both within MZ and DZ pairs, the finding would suggest independence from familial factors and support a direct association between smoking and physical inactivity. Due to the lack of longitudinal studies that control for familial confounding, we investigated the longitudinal association between smoking status and leisure-time physical inactivity in the Finnish adult twin cohort with a follow-up from 1975 to 2011. Our specific aims were to analyze the effects of smoking behavior change and familial confounding on these associations.

METHODS

Sample

Four waves of postal surveys (participation rates 72%-89%) from the population-based Finnish Older Twin Cohort of same-sex adult twins were gathered in 1975, 1981, 1990, and 2011 (Kaprio 2013). To maintain continuity, the original questions used in 1975 were used wherever possible over time. Our analysis was restricted to twins born between 1945 and 1957 invited to the 2011 survey (Piirtola et al. 2016). Among the 5575 individuals participating in all four surveys, 321 individuals (6% of the total sample) had missing or incomplete LTPA data in at least one survey and were therefore excluded. In 1975, there were no differences in leisure-time physical inactivity, smoking status, or mean alcohol consumption between the 321 excluded and the 5254 included participants. However, the mean age of the 321 excluded individuals (51% men) was 0.7 (SD 0.22) years higher ($P=0.001$), and they had on average 1.18 (SD 0.15) years less education ($P<0.001$) and had a 0.32 kg/m² (SD 0.16) higher BMI ($P=0.042$) compared to the 5254 included participants at baseline. Our final sample consisted of 5254 individuals. Analyses and results related to selection bias and drop-out during the 35-year follow-up (1975 – 2011) are presented in the Electronic Supplementary Material [ESM] 1.

Leisure-time physical inactivity

The focus of this study was on a low level of LTPA, that is, *leisure-time physical inactivity* (outcome variable), and especially in long-term inactivity or change into inactive. The LTPA questions were the same in all surveys except in 1990. The LTPA questions and the metabolic equivalent of task (MET) calculations for this cohort have been described in detail earlier (Piirtola et al. 2014; Piirtola et al. 2016). Briefly, LTPA (exercise) was elicited by asking, “Is your leisure-time physical exercise on average as intensive as...”, with four response alternatives: walking,

walking and jogging, jogging, or running. In addition, daily time for commuting by physically active means (walking, jogging, and cycling) to and from work was gathered as a separate question and included in the total MET index of LTPA. The MET index/day was calculated by multiplying the general intensity (4–13 METs) and average duration and frequency of activities at each time point and converted into MET hours/day. A sub-study has shown intraclass correlations of 0.68 for the LTPA (exercise) and 0.93 for commuting activities between the questionnaire-based MET index and a comprehensive structured face-to-face interviewed MET index (Waller et al. 2008). At each survey, the mean daily LTPA energy expenditure ≤ 1.5 MET hours (≤ 10.5 MET hours/week) was applied to define inactive (Dumith et al. 2011; Lee et al. 2012). Those with a MET index > 1.5 were defined as active. Our focus was on long-term LTPA behavior. Thus, a combined LTPA status in 1975 and 1981 was used for a baseline LTPA status; an inactive group was formed from those inactive both in 1975 and 1981 ($n=1728$), and those active in one or both surveys were defined as active ($n=3526$) (Figure 1).

Smoking status

In each survey, smoking status (main independent variable) was elicited the same way and categorized into five classes: never-smoker, former smoker, occasional smoker, daily smoker, and other (i.e. illogical reporting/ missing information) (Kaprio and Koskenvuo 1988). For the statistical analyses, the baseline smoking status (1975/81) was categorized into five classes: never-smokers, quitters before 1975, quitters between 1975 and 1981, daily smokers, and others (EMS 2A).

Correspondingly, the long-term smoking status (1975/90) was categorized into seven major classes: consistently never-smokers, quitters before 1975, quitters during 1975/81, quitters during 1981/90, recurrent smokers (i.e. those unsuccessful in cessation and restarting smoking), persistently daily smokers, and others (ESM 2B). The smoking category “others” was included in the analyses to

maintain all participants in the analyses. No interpretation was made of the results from the “other” group given its heterogeneity.

Covariates

Covariates related to leisure-time physical inactivity and/or smoking were assessed primarily from the 1981 questionnaire, except for *socioeconomic status*, which was elicited only in 1975 and consisted of six main classes: upper and lower non-manual workers, skilled and unskilled manual workers, farmers, and others (including students, conscripts, full-time homemakers, and otherwise not classified). Other covariates were *age* (years), *education* (obtained as nine categories of education attainment, then converted into years of education), *marital status* (dichotomized as single, divorced or widowed, vs married or cohabiting), *body mass index (BMI)* (kg/m², computed from self-reported weight and height), *alcohol consumption* (grams of absolute alcohol per day, based on the average consumption of beer, wines, and spirits), *life satisfaction* (categorized into three classes: satisfied, intermediate, and dissatisfied), and *working status including physical loading* (sedentary work, more active work, not at work for any reason).

Statistical analyses

Stata SE version 13.1 (StataCorp, College Station, Texas, USA) was used for all analyses. Logistic regression models were used to calculate Odds Ratios (OR) with 95% Confidence Intervals (CI) for the whole cohort (twins treated as individuals). To account for the dependency of twin individuals within a pair, a cluster option was used to obtain robust standard errors. All analyses were first adjusted for age and sex, then for multiple covariates. First, the cross-sectional association was analyzed between baseline smoking status (1975/81) and baseline inactivity (1975/81) (Figure 1). Then, the associations were analyzed between baseline smoking status and future inactivity in 1990 or in 2011. Thereafter, we analyzed the associations of smoking status with long-term inactivity

(remaining inactive) and a change in LTPA status (becoming inactive) from the baseline to 2011. Because the effect of LTPA in 1990 on the association between the long-term smoking status and inactivity over 35 years was minimal (data not shown), LTPA in 1990 was not included as a covariate in the final analyses. In all analyses, the group of consistent never-smokers was used as a reference group. In analysis to clarify the effect of quitting smoking on inactivity, the persistent daily smokers were used as reference.

We tested the interaction between sex and smoking status (at baseline or long term) on inactivity (at baseline, in 1990, or in 2011) with a log-likelihood test. A statistically significant interaction between sex and smoking status was found for inactivity ($P=0.039$) at baseline but not for inactivity in 1990 ($P=0.58$). Further, interaction between sex and long-term smoking status (1975/90) was not found on inactivity in 2011 ($P=0.10$). Thus, men and women were combined in all analyses with sex as a covariate. We also tested if the effect of long-term smoking status on inactivity would differ across zygosity in twin individuals. However, no interaction between zygosity and smoking was found ($P=0.69$).

Finally, we repeated the longitudinal analyses for associations of baseline and long-term smoking status with inactivity using a co-twin control design (McGue et al. 2010). For the cross-sectional pair-wise analyses, each twin individual was classified as either inactive or active, as was done in analyzing the whole cohort (Figure 1). For the longitudinal discordant analyses, twin individuals were dichotomized either into predominantly inactive ($n=1337$) or predominantly active ($n=2955$) according to each individual's 35-year LTPA behavior (Figure 1) (Piirtola et al. 2016). The dichotomization was done to retain enough power in the analyses. Individuals and twin pairs with mixed LTPA behavior ($n=962$) were excluded from these longitudinal discordant analyses. The data of 5254 twin individuals included altogether 1604 full twin pairs: 588 MZ and 944 DZ twin pairs. A

further 72 pairs were of uncertain zygosity based on the questionnaire-based classification algorithm, showed a high degree of agreement with multiple polymorphic genetic markers (Sarna et al. 1978). At baseline, we identified 588 twin pairs (374 DZ; 190 MZ pairs, 24 uncertain zygosity) discordant for their LTPA. The number of long-term LTPA discordant twin pairs was 370 (238 DZ; 114 MZ pairs, 18 uncertain zygosity). The discordant analyses were performed using conditional (fixed-effects) logistic regression.

RESULTS

The mean age of the participants (41% men) was 24 years (range 18–31) in 1975 and 60 years (range 53–67) in 2011. At baseline, 33% of the participants were inactive (31% of men; 34% of women), 45% were never-smokers, 12% had quit smoking before 1975 and 13% between 1975 and 1981, whereas 22% were daily smokers. Table 1 describes the smoking status and the covariates by sex and the LTPA level at baseline. In 1990, 34% were inactive (39% of men; 31% of women), and 36% in 2011 (43% of men; 33% of women). The long-term smoking behavior (1975/90) is shown in the EMS 3.

At baseline, both daily smokers and quitters between 1975 and 1981 had a higher likelihood of being inactive (Table 2), and the association remained after accounting for multiple covariates. In the longitudinal analyses, being a daily smoker at baseline (1975/81) was associated with inactivity nine years later in 1990 both in the age- and sex-adjusted and in the multiple-adjusted models (Table 2). Being a daily smoker or quitter between 1975 and 1981 was also associated with being inactive 30 years later in 2011 also in the multiple-adjusted model.

In the age- and sex-adjusted analyses of the long-term smoking behavior (1975/1990), persistent daily smokers showed a higher likelihood for both remaining inactive and becoming inactive in 2011 (Table 3). Additionally, recurrent smokers were more likely to remain inactive. Any history of smoking showed a trend for remaining or becoming inactive in 2011, while accounting for multiple covariates did not change the associations.

An additional analysis was run for the effect of quitting smoking on remaining or becoming inactive between the baseline and 2011. Compared to persistently daily smokers, quitters before 1975

(OR=0.41, 95% CI: 0.25, 0.67), quitters during 1975/81 (OR=0.53, 95% CI: 0.32, 0.89), and quitters during 1981/90 (OR=0.42, 95% CI: 0.25, 0.71) were less likely to become inactive in the multiple-adjusted analyses. The association between quitting smoking and remaining inactive was significant only in quitters during 1975/81 (OR=0.57, 95% CI: 0.38, 0.85).

Furthermore, we tested the effect of familial confounding on the associations (Table 4). Daily smoking was associated with inactivity both in the cross-sectional and in the longitudinal discordant pair analyses. The association was strongest for persistent daily smokers, but evident also for any kind of smoking history in the longitudinal analyses. In the analyses for MZ and DZ twins separately, the association remained significant for the DZ pairs, whereas the estimates for MZ pairs attenuated.

DISCUSSION

Principal findings

Our study within a cohort of 5254 twin individuals with responses to four comprehensive surveys over a 35-year follow-up showed that daily smoking, especially being a long-term daily smoker or a recurrent smoker, was associated with long-term leisure-time physical inactivity. In comparison to persistent daily smokers, quitters had a lower likelihood of inactivity. Accounting for familial confounding indicated that any kind of smoking history was associated with inactivity over 35 years. Our study adds to the previous findings suggesting a direct effect of smoking on physical inactivity.

Comparison with other studies

In particular, our finding of the association of long-term daily smoking with long-term physical inactivity merits attention. Despite some previously reported evidence about a lesser amount of LTPA among persistent daily smokers compared to never-smokers in a long-term continuum (Auer et al. 2014; Azagba and Asbridge 2013; Nagaya et al. 2007; Paavola et al. 2004; Perkins et al. 1993), our study is among the first with a follow-up over three decades (Auer et al. 2014; Borodulin et al. 2012) and more than two time points in the analyses (Auer et al. 2014; Azagba and Asbridge 2013; Nagaya et al. 2007; Paavola et al. 2004). In a 25-year follow-up with eight time points, increasing years of smoking were associated with a decrease in the amount of LTPA, whereas an increase in LTPA was associated with successful smoking cessation (Auer et al. 2014), which is in line with our results. However, indications exist that not all daily smoking is associated with less LTPA (Azagba and Asbridge 2013). For example, a 6-year follow-up study with several survey points showed that only daily smokers with nicotine dependence, that is, heavy smokers, who usually have a longer smoking history and who are on average less successful in smoking cessation,

had a higher likelihood for inactivity compared with less dependent smokers (Azagba and Asbridge 2013). We did not have measures of nicotine dependence, a limitation that would call for further studies. It is also relevant that there is no global definition for physical inactivity (Dumith et al. 2011; Lee et al. 2012; Sedentary Behaviour Research Network 2012). Thus, the effect of smoking status on leisure-time physical inactivity might differ across studies depending on the definition of physical inactivity used.

Another important finding was that in comparison to persistent daily smokers, quitting smoking was related to a lower likelihood of inactivity. Earlier results suggest that each additional year of successful cessation increases the positive net effect of smoking cessation on an increase in LTPA compared to those who have continued smoking (Auer et al. 2014). We were only interested in those with the lowest amount of LTPA, not in an increase in LTPA. However, an increase in habitual exercise after smoking cessation was indicated in a study with annual time points over seven years among healthy men, whereas a smoking relapse also decreased the amount of LTPA (Nagaya et al. 2007).

Interpretation of the discordant analyses

A unique aspect of this study was the twin study design, which permits controlling for familial confounding. In twin pairs discordant for LTPA, the twin who was a daily smoker had a higher likelihood of being inactive in both the cross-sectional and longitudinal analyses compared to his/her non-smoking twin brother or sister. The longitudinal associations between smoking status over 1975–1990 and inactivity pointed to the same direction regardless of quitting, in other words, any kind of smoking history increased the likelihood for inactivity. Hence, our results indicate the importance of not only support for quitting smoking, but also the prevention of smoking initiation in the first place. Notably, in the long-term discordant analyses, the point estimates attenuated to

statistical non-significance in the MZ pairs for all smoking categories, suggesting limited power in the separate analyses of MZ and DZ twins.

Both LTPA and smoking have a strong genetic component (de Geus et al. 2014; de Vilhena e Santos et al. 2012; Li et al. 2003; McGue et al. 2014), whereas the heritability of physical inactivity has been reported to be only modest (de Geus et al. 2014). Since smoking usually starts already during the teenage years (Auer et al. 2014), the influence of parental practices and living conditions in childhood is obvious. To the best of our knowledge, prospective twin studies analyzing the association of smoking with LTPA while controlling for familial confounding are rare if any. However, we acknowledge an earlier study conducted among the younger Finnish twin cohort, where the longitudinal association in the reverse direction, that is, between physical inactivity in adolescence and smoking in adulthood, was not confounded by familial factors (Kujala et al. 2007).

Pathways between smoking and inactivity

The mechanisms are not fully known underlying the association between smoking and physical inactivity. One potential explanation is the association between respiratory capacity and LTPA, since smoking reduces pulmonary function and therefore compromises LTPA performance (Higgins et al. 1991). However, smoking is related to inactivity already in adolescence (Aarnio et al. 2002) when the negative effects of smoking on pulmonary functions are evident but still modest (Gold et al. 1996). Long-term smoking is, however, associated with many chronic diseases (Lim et al. 2012), which might reduce physical activity. Other factors (social behavior, life circumstances, or early experiences with smoking experimentation) may also explain the association of smoking with inactivity. Notably, smoking has also been used as a weight control mechanism (Potter et al. 2004). Another pathway might be related to psychosocial factors such as mood, stress, and depression, which may affect smoking and consequently inactivity (Kaczynski et al. 2008). Long-term smoking

has been shown to be associated with depression (Korhonen et al. 2011), which might modify LTPA behavior (Kaczynski et al. 2008). Instead, smoking cessation can reduce depression, anxiety, and stress and improve positive mood (Taylor et al. 2014b), whereas LTPA is known to reduce depressive symptoms (De Moor et al. 2008). However, in our study life dissatisfaction, used as a proxy for depression (Koivumaa-Honkanen et al. 2004), did not weaken the association between smoking and physical inactivity. Despite the fact that successful smoking cessation has been reported to improve mental health in a meta-analysis (Taylor, McNeill, et al. 2014), a clear causal association between smoking and depression was not supported by another Mendelian randomization meta-analysis (Taylor, Fluharty, et al. 2014).

The pathway between smoking and inactivity may also be affected by the complex associations between several other lifestyle choices, because health-related lifestyle choices usually occur in combinations (Auer et al. 2014; Morris et al. 2016). For example, changing one lifestyle behavior could act as a catalyst to change another (Kaczynski et al. 2008), and in influential lifestyle changes, smoking seems to play a central role (Auer et al. 2014; Morris et al. 2016; Paavola et al. 2004). In our study, quitters still had a higher likelihood for inactivity, even when controlling for alcohol consumption and BMI as reflectors of other lifestyle factors. Hence, interventions aimed to influence lifestyles should include simultaneous support for many behavioral changes, such as smoking cessation.

Study strengths and limitations

This longitudinal cohort study with twin pairs has several strengths. The comprehensive surveys repeated four times provided a unique opportunity not only to follow the same individuals over 35 years but also to investigate the long-term development of both smoking and LTPA behaviors. The Finnish Twin Cohort is representative of the Finnish adult population (Kaprio and Koskenvuo

1988). The proportion of daily smokers decreased during the 35-year follow-up as a result of increasing cessation with age and the drop-out rate of smokers due to death and non-participation. Overall, this decrease is also in line with the decrease in smoking in Finnish adults (Midlöv et al. 2014; Morris et al. 2016; Ng et al. 2014). The dataset also provided a powerful case-control setting to control for familial confounding on the associations of interest. In addition to familial confounding, the study also controlled for several covariates with a known influence on smoking and LTPA.

Regarding study limitations, more current daily smokers and occasional smokers in 1975 had died by 2011, and fewer smokers participated in the last survey in 2011. Thus, this drop-out most likely dilutes the associations in our analyses. Since we had time points situated 6, 9, and 21 years apart, additional changes regarding both smoking and physical inactivity might have occurred during the 35-year follow-up. Any unrecorded changes in these factors of interest could have had an impact on the observed associations and would dilute the effect. However, we believe that four time points over 35-year of follow-up has rarely been done before. Another limitation in this study is that the measure of LTPA was not based on objective but on self-reported assessments without information about domestic (everyday) activities. However, objective measurements were not available for large cohort studies in the 1970s. Our analyzing methods did not allow taking into account time-dependent changes in co-variables, for example in marital status. This may have affected our results. However, the main focus of this study was on the association between smoking status and long-term leisure-time physical inactivity or change into inactive during the follow-up. The analyzing methods were selected to capture whether stability or change in smoking status would affect stability or change in leisure-time physical inactivity over time. However, further studies using alternate methods, such as bivariate cross-lagged path model, are needed to investigate causal

relationship between varying smoking status and full spectrum of leisure-time physical activity as well as the influence of co-variates on the associations.

Conclusions

Being a long-term persistent daily smoker increases the likelihood of remaining or becoming physically inactive whereas quitting smoking decreases the likelihood of inactivity. However, since any kind of smoking history associates with physical inactivity in the long-term discordant analyses, efforts to prevent smoking initiation and to support smoking cessation during early adulthood along with the promotion of a healthy lifestyle might be important for promoting physical activity. The association between smoking and inactivity seems to be independent of other factors. Extensive health behavior counselling and effective tobacco control policies are needed in promoting lifelong physical activity.

COMPLIANCE WITH ETHICAL STANDARDS

Ethical approval

All procedures performed in studies involving human participants were in accordance with the ethical standards of the institutional and/or national research committee and with the 1964 Helsinki declaration and its later amendments or comparable ethical standards. The study protocol was approved by the Ethical Committee of the University of Helsinki, Department of Public Health. This present analysis did not involve any contact with the study participants and only used existing epidemiological data.

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Conflicts of interest

JK reports personal fees from Pfizer unrelated to the submitted work. TK reports personal fees from Pfizer for consulting on nicotine dependence but unrelated to the submitted work. None of the other authors have anything to declare.

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FIGURE LEGENDS

Figure 1. Categorization of leisure-time physical activity by surveys (1975, 1981, 1990, 2011), at combined baseline (1975/81), and in analyzing remaining (long-term) inactive or becoming inactive both in the whole cohort (n=5254) and in the leisure-time physical activity discordant twin pairs among the Finnish Older Twin Cohort, Finland, 1975-2012.

Abbreviations: LTPA= leisure-time physical activity; MET= metabolic equivalent; A=active (>1.5 MET h/day); I=inactive (\leq 1.5 MET h/day); II= inactive in 1975 and 1981; AA= active in 1975 and 1981; AI= active in 1975, inactive in 1981; IA= inactive in 1975, active in 1981; III = inactive in all four time points (1975, 1981, 1990 and 2011); AIII= active in 1975, inactive in 1981, 1990 and 2011; AAI= active in 1975, 1981 and 1990, inactive in 2011; AII= active in 1975 and 1981, inactive in 1990 and 2011; AAI= active in 1975, 1981 and 1990, inactive in 2011; IAI= inactive in 1975 and 1981, active in 1990, inactive in 2011; IAI= inactive in 1975, active in 1981, inactive in 1990 and 2011; AAAA: active in all four time points (1975, 1981, 1990 and 2011); IAAA= inactive in 1975, active in 1981, 1990 and 2011; IIAA= inactive in 1975 and 1981, active in 1990 and 2011; IIIA= inactive in 1975, 1981 and 1990, active in 2011; AAIA= active in 1975 and 1981, inactive in 1990, active in 2011; AIAA= active in 1975, inactive in 1981, active in 1991 and 2011.

Table 1. Sex-Specific Characteristics [% (N)] of Smoking Status and Confounding Factors by Leisure-Time Physical Activity at Baseline (1975/81) Among 5254 Twin Individuals From the Finnish Older Twin Cohort, Finland, 1975-2012.

Characteristics	Men (n=2143)			Women (n=3111)		
	Inactive in both 1975 and 1981 (n = 674) % (n)	Change in activity between 1975 and 1981 (n = 741) % (n)	Active in both 1975 and 1981 (n = 728) % (n)	Inactive in both 1975 and 1981 (n = 1054) % (n)	Change in activity between 1975 and 1981 (n=1240) % (n)	Active in both 1975 and 1981 (n = 817) % (n)
Smoking status 1975/81						
Never-smokers	29 (196)	35 (262)	42 (308)	47 (498)	51 (628)	56 (454)
Quitters before 1975 ^a	12 (78)	12 (89)	17 (121)	11 (113)	9 (117)	11 (88)
Quitters 1975-1981	13 (84)	13 (98)	12 (88)	14 (148)	13 (164)	10 (85)
Daily smokers	34 (227)	28 (209)	20 (144)	21 (212)	18 (226)	15 (125)
Others ^b	13 (87)	11 (83)	9 (67)	8 (83)	8 (105)	8 (65)
Social class 1975						
Upper white collar	6 (41)	8 (60)	8 (61)	2 (21)	3 (40)	5 (40)
Lower white collar	14 (94)	16 (117)	20 (147)	31 (329)	34 (420)	43 (353)
Skilled worker	52 (348)	48 (355)	45 (326)	33 (347)	29 (365)	28 (230)
Unskilled worker	7 (48)	9 (70)	7 (50)	11 (114)	11 (136)	10 (80)
Farmer	6 (43)	6 (42)	4 (28)	4 (42)	1 (18)	1 (12)
Other (including students)	14 (97)	13 (95)	16 (113)	19 (200)	21 (259)	12 (101)
Life satisfaction 1981						
Satisfied	17 (112)	20 (148)	23 (170)	21 (223)	24 (297)	25 (204)
Intermediate	68 (457)	65 (481)	65 (473)	65 (684)	65 (801)	63 (512)
Dissatisfied	16 (105)	15 (112)	12 (85)	14 (147)	11 (142)	12 (101)
Marital status 1981						
Married or living together	67 (450)	68 (503)	68 (495)	77 (816)	70 (867)	66 (538)
Single, divorced, or widowed	33 (224)	32 (238)	32 (232)	23 (238)	30 (373)	34 (279)
Work physical loading 1981						
Sedentary work	30 (203)	27 (197)	34 (248)	27 (280)	35 (433)	43 (352)
More active work	58 (391)	65 (482)	60 (435)	35 (371)	42 (524)	43 (350)
Not at work	12 (80)	8 (62)	6 (45)	38 (403)	23 (283)	14 (115)
	Mean (SD)	Mean (SD)	Mean (SD)	Mean (SD)	Mean (SD)	Mean (SD)
Education 1981 (years)	8.4 (2.8)	8.8 (3.1)	9.3 (3.3)	8.8 (2.9)	9.2 (3.1)	9.1 (3.0)
Age 1981 (years)	30.3 (3.8)	30.3 (3.7)	30.5 (3.7)	29.9 (3.8)	29.8 (3.8)	30.2 (3.7)

BMI 1981 (kg/m ²)	23.6 (2.9)	23.6 (2.8)	23.4 (2.5)	22.0 (3.2)	21.6 (2.9)	21.4 (2.6)
Alcohol consumption 1981 (g/day)	12.5 (16.0)	10.7 (12.5)	10.1 (11.4)	4.1 (5.5)	4.3 (6.7)	4.2 (5.6)

a) Quitters who stopped smoking before 1975 and remained non-smokers until 1981.

b) The group of “others” includes occasional smokers, initiators, recurrent smokers, and other kinds of non-persistent smoking behavior.

Inactive= LTPA energy cost ≤1.5 MET hours per day; Active= LTPA energy cost >1.5 MET hours per day.

Abbreviations: BMI= Body Mass Index; LTPA= leisure-time physical activity; MET= metabolic equivalent.

Table 2. Associations Between Baseline (1975/81) Smoking Status and Baseline or Future Leisure-Time Physical Inactivity Among 5254 Twin individuals From the Finnish Older Twin Cohort, Finland, 1975-2012; Logistic Regression Analyses (Odds Ratios With 95% Confidence Intervals).

Smoking status at baseline (1975/81)	Inactive*					
	at baseline (1975/81)		in 1990		in 2011	
	Age- and sex adjusted model	Multiple-adjusted model **	Age- and sex adjusted model	Multiple-adjusted model **	Age- and sex adjusted model	Multiple-adjusted model **
	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)
Never-smokers	Reference	Reference	Reference	Reference	Reference	Reference
Quitters before 1975 ^a	1.13 (0.93, 1.38)	1.12 (0.91, 1.37)	1.07 (0.88, 1.30)	1.07 (0.88, 1.31)	1.02 (0.85, 1.24)	1.02 (0.83, 1.24)
Quitters 1975-1981	1.28 (1.07, 1.55)	1.25 (1.03, 1.52)	0.99 (0.82, 1.20)	1.01 (0.83, 1.22)	1.21 (1.00, 1.45)	1.18 (0.98, 1.43)
Daily smokers	1.52 (1.31, 1.78)	1.53 (1.30, 1.82)	1.38 (1.19, 1.61)	1.35 (1.15, 1.59)	1.88 (1.62, 2.18)	1.75 (1.49, 2.06)
Others ^b	1.30 (1.05, 1.60)	1.31 (1.04, 1.63)	1.22 (0.99, 1.50)	1.27 (1.03, 1.57)	1.34 (1.08, 1.65)	1.32 (1.06, 1.64)

* The inactive group were compared to those more active. Inactive (LTPA energy cost \leq 1.5 MET hours per day); active (LTPA energy cost $>$ 1.5 MET hours per day). In 1975/81 (baseline), the inactive group was defined as being inactive in both 1975 and 1981, and those defined as “more active” were either active in 1975 and 1981 or changed their LTPA behavior between the surveys.

** Covariates included sex, age at the survey, Body Mass Index in 1981, education years in 1981, use of alcohol (grams) in 1981, working status in 1981, marital status in 1981, life satisfaction in 1981, and socioeconomic status in 1975.

a) Quitters who stopped smoking before 1975 and remained non-smokers until 1981.

b) The group of “others” includes occasional smokers (n=56), initiators (n=139), recurrent smokers (n=193), and other kinds of non-persistent smoking behavior (n=95).

Abbreviations: LTPA= leisure-time physical activity; MET= metabolic equivalent; OR= odds ratio; 95% CI= 95% Confidence Intervals.

Table 3. Prospective Associations of Long-Term Smoking Behavior (1975/1990) and Remaining or Becoming Physically Inactive in Leisure Time Over a 30-year (2011) Follow-up by the Leisure-Time Physical Activity Status at Baseline (1975/81) Among 5254 Twin Individuals From the Finnish Older Twin Cohort, Finland, 1975-2012; Logistic Regression Analyses (Odds Ratios with 95% Confidence Intervals).

Smoking behavior 1975/1990	Inactive in 1975/81 and odds of remaining inactive in 2011		Active in 1975/81 and odds of becoming inactive in 2011	
	Age- and sex-adjusted model	Multiple-adjusted model*	Age- and sex-adjusted model	Multiple-adjusted model*
	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)
Consistent never-smokers	Reference	Reference	Reference	Reference
Quitters before 1975 ^a	1.22 (0.84, 1.77)	1.22 (0.84, 1.78)	1.11 (0.73, 1.67)	1.15 (0.74, 1.77)
Quitters between 1975 and 1981 ^b	1.02 (0.72, 1.46)	1.02 (0.70, 1.47)	1.56 (0.99, 2.45)	1.50 (0.95, 2.37)
Quitters between 1981 and 1990 ^c	1.19 (0.81, 1.76)	1.17 (0.78, 1.75)	1.14 (0.72, 1.82)	1.19 (0.74, 1.93)
Recurrent smokers ^d	1.48 (1.01, 2.16)	1.65 (1.12, 2.44)	1.38 (0.89, 2.14)	1.37 (0.88, 2.15)
Persistent daily smokers	1.91 (1.45, 2.51)	1.80 (1.34, 2.42)	2.07 (1.93, 3.81)	2.81 (1.95, 4.05)
Others ^e	1.50 (1.03, 2.22)	1.49 (0.99, 2.23)	0.97 (0.54, 1.74)	0.99 (0.54, 1.82)

In each survey: inactive (LTPA energy cost \leq 1.5 MET hours per day); active (LTPA energy cost $>$ 1.5 MET hours per day).

- a) Quitters who stopped smoking before 1975 and remained non-smokers until 1990.
- b) Quitters who stopped smoking between 1975 and 1981 and remained non-smokers until 1990.
- c) Quitters who had mixed smoking status in 1975, at least occasional smoking in 1981, and reported quitting smoking before 1990.
- d) Recurrent smokers who restarted their smoking either between 1975 and 1981 (n=102) or between 1981 and 1990 (n=288).
- e) The group of “others” includes initiators (n=140), consistently occasional smokers (n=19), and other kinds of non-persistent smoking behavior (n=181).

* Covariates included sex, age at the survey, Body Mass Index in 1981, education years in 1981, use of alcohol (grams) in 1981, working status in 1981, marital status in 1981, life satisfaction in 1981, and socioeconomic status in 1975. Twin-pair identity was used as a cluster in the analyses.

Abbreviations: LTPA= leisure-time physical activity; MET=metabolic equivalent; OR,=odds ratio; 95% CI= 95% confidence intervals.

Table 4. Associations of Baseline (1975/81) and Long-Term (1975/1990) Smoking Behaviors With Being Physically Inactive in the Same-sex Twin Pairs Discordant for Leisure-Time Physical Activity (Inactive vs. Active) at Baseline and Longitudinally in the Finnish Older Twin Cohort during a 35-year follow-up, Finland, 1975-2012; Conditional Logistic Regression Analyses (Odds Ratios with 95% Confidence Intervals).

	LTPA discordant twin pairs at baseline (1975/81)*					
	Basic model **			Multiple-adjusted model***		
	All pairs (n=588)	DZ pairs (n=374)	MZ pairs (n=190)	All pairs (n=558)	DZ pairs (n=355)	MZ pairs (n=179)
Baseline smoking status 1975/81	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)
Never-smokers	Reference	Reference	Reference	Reference	Reference	Reference
Quitters before 1975 ^a	1.20 (0.76, 1.91)	1.19 (0.70, 2.01)	0.98 (0.31, 3.06)	1.30 (0.78, 2.18)	1.30 (0.71, 2.38)	0.89 (0.22, 3.64)
Quitters 1975-1981	1.25 (0.81, 1.94)	1.54 (0.93, 2.58)	0.80 (0.30, 2.16)	1.40 (0.87, 2.28)	1.66 (0.94, 2.92)	1.04 (0.31, 3.53)
Daily smokers	1.55 (1.01, 2.40)	1.49 (0.90, 2.47)	1.73 (0.61, 4.86)	1.80 (1.10, 2.94)	1.64 (0.92, 2.93)	3.22 (0.86, 11.99)
Others ^b	1.43 (0.80, 2.54)	1.35 (0.67, 2.71)	2.31 (0.66, 8.04)	1.47 (0.78, 2.76)	1.26 (0.58, 2.74)	2.97 (0.71, 12.43)
	LTPA discordant twin pairs for 35-year follow-up (1975–2011)****					
	Basic model **			Multiple-adjusted model***		
	All pairs (n=370)	DZ pairs (n=238)	MZ pairs (n=114)	All pairs (n=350)	DZ pairs (n=226)	MZ pairs (n=107)
Longitudinal smoking status 1975/1990	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)	OR (95% CI)
Consistent never-smokers	Reference	Reference	Reference	Reference	Reference	Reference
Quitters before 1975 ^c	2.43 (1.29, 4.57)	2.89 (1.34, 6.22)	1.75 (0.42, 7.30)	3.20 (1.53, 6.69)	5.10 (1.90, 13.74)	2.53 (0.46, 14.02)
Quitters between 1975 and 1981 ^d	2.73 (1.26, 5.92)	4.59 (1.63, 13.00)	1.21 (0.29, 5.13)	3.73 (1.52, 9.15)	7.49 (2.07, 27.14)	3.17 (0.40, 24.95)
Quitters between 1981 and 1990 ^e	2.28 (1.10, 4.74)	2.11 (0.83, 5.37)	3.08 (0.72, 13.07)	2.86 (1.22, 6.74)	2.25 (0.74, 6.86)	7.25 (1.08, 48.71)
Recurrent smokers ^f	2.63 (1.29, 5.36)	4.69 (1.91, 11.49)	0.73 (0.15, 3.56)	3.96 (1.68, 9.32)	14.93 (4.32, 51.61)	0.90 (0.11, 7.04)
Persistent daily smokers	4.70 (2.45, 9.03)	6.55 (2.80, 15.28)	2.54 (0.69, 9.41)	8.07 (3.61, 18.02)	16.51 (5.35, 50.90)	7.07 (1.01, 49.60)

Others ^g	1.78 (0.85, 3.76)	2.03 (1.06, 3.87)	2.14 (0.50, 9.22)	1.99 (0.85, 4.66)	1.68 (0.54, 5.19)	3.91 (0.57, 26.83)
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- a. Quitters who stopped smoking before 1975 and remained non-smokers until 1981.
- b. The group of “others” in 1975/81 includes occasional smokers, initiators, recurrent smokers, and other kinds of non-persistent smoking behavior.
- c. Quitters who stopped smoking before 1975 and remained non-smokers until 1990.
- d. Quitters who stopped smoking between 1975 and 1981 and remained non-smokers until 1990.
- e. Quitters who had mixed smoking status in 1975, at least occasional smoking in 1981, and reported quitting smoking before 1990.
- f. Recurrent smokers who restarted their smoking either between 1975 and 1981 or between 1981 and 1990.
- g. The group of “others” in 1975–1990 includes initiators, consistently occasional smokers, and other kinds of non-persistent smoking behavior.

* One twin within a pair inactive (LTPA energy cost \leq 1.5 MET hours per day) and the other more active (LTPA energy cost $>$ 1.5 MET hours per day) in the survey. In 1975/81 one twin inactive in both surveys (1975 and 1981) and the other twin active in either or both surveys.

** In the same-sex twin pairs, age and sex are automatically adjusted in the basic model.

*** In addition to sex and age, also Body Mass Index in 1981, education years in 1981, use of alcohol (grams) in 1981, working status in 1981, marital status in 1981, life satisfaction in 1981, and socioeconomic status in 1975 are adjusted in the model.

**** Twin pairs long-term discordant in LTPA, that is, one twin within a pair was inactive (persistently inactive, mainly inactive, or a change from active to inactive during the 35-year follow-up) and the other was more active (persistently active, mainly active, or a change from inactive to active during the 35-year follow-up). Mixed LTPA was excluded from the analyses.

Abbreviations: DZ=dizygotic; LTPA= leisure-time physical activity; MZ= monozygotic; OR= odds ratio; 95% CI= 95% confidence intervals.

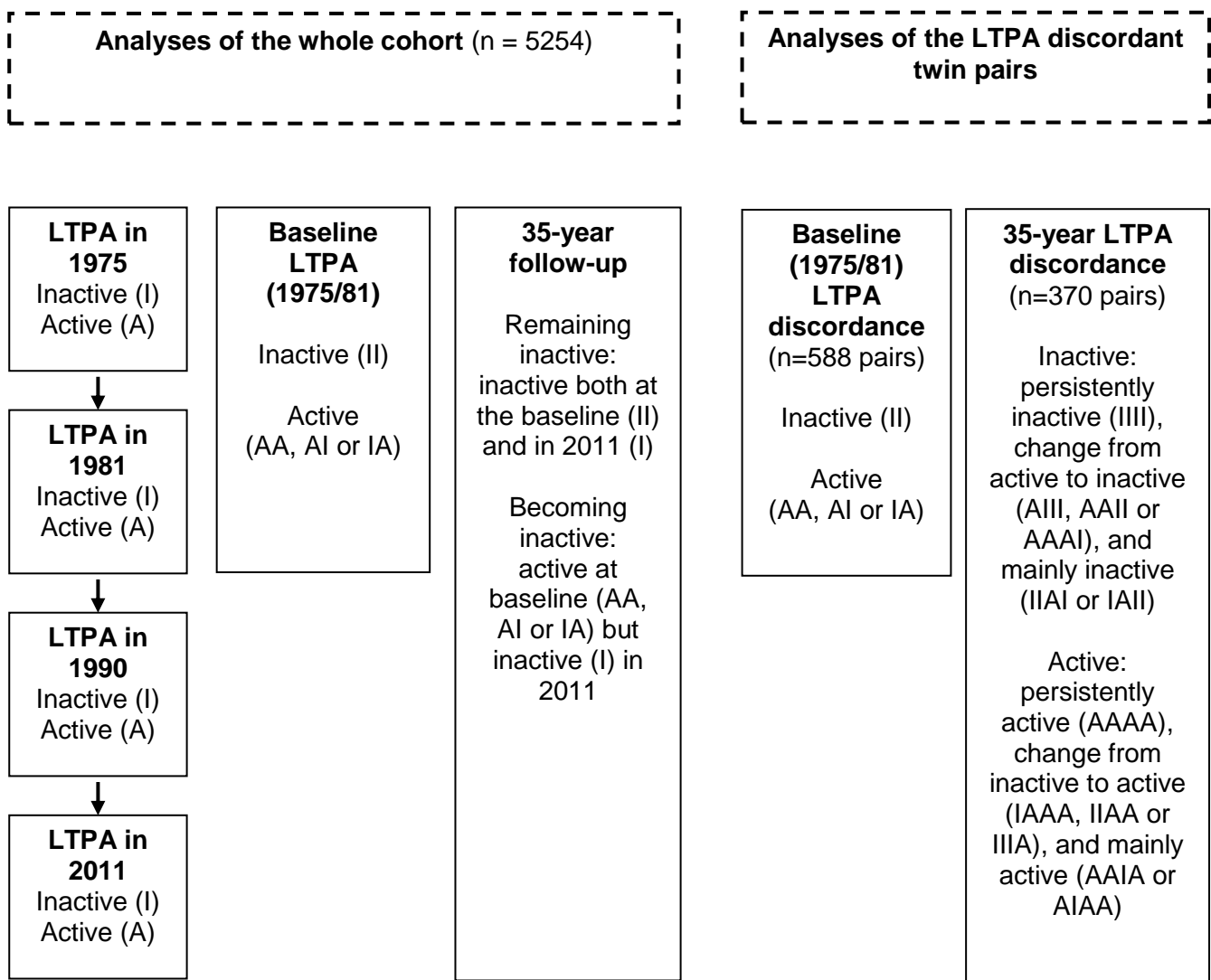


Figure 1. Categorization of leisure-time physical activity by surveys (1975, 1981, 1990, 2011), at combined baseline (1975/81), and in analyzing remaining (long-term) inactive or becoming inactive both in the whole cohort (n=5254) and in the leisure-time physical activity discordant twin pairs among the Finnish Older Twin Cohort, Finland, 1975-2012.

Abbreviations: LTPA= leisure-time physical activity; MET= metabolic equivalent; A=active (>1.5 MET h/day); I=inactive (\leq 1.5 MET h/day); II= inactive in 1975 and 1981; AA= active in 1975 and 1981; AI= active in 1975, inactive in 1981; IA= inactive in 1975, active in 1981; IIII = inactive in all four time points (1975, 1981, 1990 and 2011); AIII= active in 1975, inactive in 1981, 1990 and 2011; AII= active in 1975 and 1981, inactive in 1990 and 2011; AAAI= active in 1975, 1981 and 1990, inactive in 2011; IIAI= inactive in 1975 and 1981, active in 1990, inactive in 2011; IAII= inactive in 1975, active in 1981, inactive in 1990 and 2011; AAAA: active in all four time points (1975, 1981, 1990 and 2011); IAAA= inactive in 1975, active in 1981, 1990 and 2011; IIAA= inactive in 1975 and 1981, active in 1990 and 2011; IIIA= inactive in 1975, 1981 and 1990, active in 2011; AAIA= active in 1975 and 1981, inactive in 1990, active in 2011; AIAA= active in 1975, inactive in 1981, active in 1991 and 2011.