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Decline in Fast Gait Speed As a Predictor of Disability in Older Adults

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Running head: Decline in gait speed predicts disability

ABSTRACT

Objectives: To determine whether fast gait speed at study baseline and change in gate speed had independent associations with disability onset.

Design: Cohort study with 11-year follow-up (1999–2010).

Setting: Three-City Study, Dijon center, France.

Participants: Community-dwelling individuals aged 65 to 85 (N=3,814, 61% female).

Measurements: Fast gait speed (6 m) was assessed up to five times and disability (mobility (Rosow-Breslau scale), instrumental activities of daily living (IADLs; Lawton-Brody scale), basic activities of daily living (ADLs; Katz scale)) six times. A hierarchical disability indicator was constructed; participants were considered disabled if they reported difficulties with mobility and IADLs or with mobility, IADLs, and ADLs. The association between baseline fast gait speed and its slope of change and disability incidence was examined using joint models for longitudinal and time-to-event data.

Results: Over follow-up, 628 participants (68% women) developed disability. Mean fast gait speed at baseline was 1.54 m/s, and annual decline was approximately 0.017 m/s. The hazard ratio of disability per standard deviation (SD) (–0.22 m/s) slower baseline fast gait speed was 1.77 (95% confidence interval (CI)=1.60–1.94) and for one-SD (–0.013 m/s) faster annual decline was 1.38 (95% CI=1.10–1.73) when both parameters were included in a sex- and age-adjusted model. All associations remained statistically significant in multivariable models, except for slope of change when chronic conditions were added to the model; of chronic conditions, dyspnea was the main contributor.

Conclusion: Accelerated decline in fast gait speed was associated with disability independent of baseline fast gait speed. These results underline the interest of measuring gait speed repeatedly and the need to identify determinants of decline, because they are likely to be targets for prevention and treatment to reduce disability risk.

Key words: aged, cohort study, disability, epidemiology, motor decline.

Given increasing life expectancy, the burden of disability is expected to rise. Disability is associated with risk of hospitalization, institutionalization, and death.¹⁻³ Identifying persons at risk of disability before they become disabled may allow high-risk populations to be identified and preventive strategies to be implemented.⁴

Previous research has shown that slow gait speed is a robust predictor of onset of disability,⁵⁻⁸ but the majority of these studies used a single measure of gait speed to predict subsequent disability. There are two shortcomings with this approach. One, a single assessment might simply reflect low vitality and poor general health. If so, attempts to improve gait speed are unlikely to reduce risk of disability. Two, measurement error is more likely to bias findings from studies that use single assessments. Decline in gait speed has been shown to be associated with mortality,⁹⁻¹¹ but few studies have examined its association with disability, with inconsistent findings.¹²⁻¹⁵

Whether baseline fast gait speed and change in fast gait speed over follow-up were independently associated with incidence of disability over 11 years of follow-up was examined in a cohort of French community-dwelling older adults from the Dijon center of the Three-City (3C) Study.

METHODS

Study population

The 3C Study recruited community-dwelling individuals aged 65 and older from electoral rolls in three French cities in 1999–2001 (Bordeaux, Dijon, Montpellier).¹⁶

The present study is based on data from Dijon (n=4,931), where a specific study on physical performance was undertaken on adults aged 65 to 85. Participants younger than 85 were invited to the study center to be interviewed in person and for additional investigations at baseline and after 2 (Wave 1, 2001–02), 4 (Wave 2, 2003–04), 7 (Wave 4, 2006–07), 9 (Wave 5, 2008–09), and 11 (Wave 6, 2010–12) years. Older participants were seen at home; from

Wave 2 onward, all participants were offered the opportunity of being seen at home. Wave 3 (2005–06) consisted only of a self-administered questionnaire.

The ethical committee of the Kremlin-Bicêtre University-Hospital (France) approved the study protocol, and all participants gave written informed consent.

Gait speed

Gait speed was measured in participants younger than 85 who visited the study center at baseline and Waves 2, 4, 5, and 6 using two photoelectric cells (6 m apart) connected to a chronometer. Participants were asked to walk at usual and fast (without running) speeds. They started walking 3 m before the start line and stopped walking after the end line so that the measure included neither acceleration nor deceleration time. At Wave 6, for participants younger than 85 who were seen at home, portable photoelectric cells (Racetime2 kit light radio, MicroGate) were used using the same protocol as at the study center. Gait speed was therefore measured at home over 6 m in most instances (85%); shorter distances (3.5–5.9 m) were used if there was not enough space at home.

Gait speed (m/s) was computed as 6 m divided by the time taken to walk this distance (seconds). Short-term reproducibility was assessed using two measures 5 minutes apart in a random sample of 51 participants (mean age 80.1 ± 3.4). Intraclass correlation coefficients (standard errors (SEs)) were 0.84 (0.02) for usual speed and 0.92 (0.02) for fast speed.¹⁷

Given greater annual decline (-0.017 vs -0.005 m/s), in agreement with other reports,^{18,19} and greater interindividual variability (random variability of the slope (SE) 0.0002 (0.00002) vs 0.0001 (0.00001)) in fast speed than usual speed, fast speed was used in the analyses.

Disability

Three disability domains (supplementary methods) were assessed six times (baseline, Waves 1, 2, 4, 5, 6). Mobility was assessed using the French translation of the Rosow and Breslau scale, which evaluates the ability to perform heavy work around the house, walk half a mile, and climb stairs.²⁰ The French version of the Lawton-Brody instrumental activity of daily

living (IADL) scale evaluates the ability to use a telephone, manage drugs and money, use public or private transportation, and shop and, additionally for women, prepare meals and do housework and laundry.²¹ Basic activities of daily living (ADLs) were assessed using the French version of the Katz scale, which evaluates whether participants need help with bathing, dressing, toileting, transferring from bed to chair, and eating; incontinence was excluded because it reflects organ impairment rather than functional limitation.²² For each domain, participants were considered disabled if they could not perform at least one activity without a given level of help, as defined according to the respective instrument.

A hierarchical disability indicator was constructed²³ that defines four levels of increasing disability by summing up responses to the three dichotomized disability items in a hierarchy (0=fully independent, 1=dependent only in relation to the Rosow scale, 2=dependent on the Rosow and IADL scales but not the ADL scale, 3=dependent in all domains). This approach has the advantage of taking three disability domains into account simultaneously; it has a reproducibility coefficient of 0.99 and a scalability coefficient of 0.98.^{23,24} Few people were disabled in all three domains, leading to the creation of two groups: 2/3 (moderate/severe disability) and 0/1 (no/light disability).

Covariates

Sociodemographic measures drawn from study baseline included age, sex, education (no education to primary school, secondary school, high school to university), and marital status (married; divorced, separated, widowed; single). An unhealthy behaviors score was established as the number of unhealthy behaviors (range 0–4) using measures from study baseline: low to intermediate physical activity (walking <1 h/d and/or exercising <1 time/wk); consumption of fruits and vegetables less than once per day; current or recent exsmoker (quit smoking <15 years before baseline); never, former, or heavy drinker (>21 alcoholic drinks/wk for men, >14 for women).²⁵

Anthropometric measures included body mass index (BMI) and height. Weight was measured during clinical examinations or self-reported during interviews at all waves. Height was measured or self-reported at baseline and Wave 4. When both measures were available, clinical assessments were used. BMI (kg/m^2) was calculated as weight divided by height squared; baseline height was used to compute BMI at baseline and Waves 1 and 2, and height from Wave 4 was used to compute BMI at Waves 4 to 6.

The following covariates were assessed at study baseline and at each wave of data collection. Cognition was assessed using the Mini-Mental State Examination (MMSE), with higher scores corresponding to better function. Depressive symptomatology were measured using the Center for Epidemiologic Studies Depression Scale (CES-D), with scores of 16 or greater corresponding to high depressive symptoms;²⁶ use of psychotropic drugs (antidepressants, anxiolytics, benzodiazepines, hypnotics) was also recorded. Trauma included history of bone fracture or recurrent falls (≥ 2 falls) over the 2 years preceding each visit. Chronic conditions included self-reported diabetes mellitus, Parkinson's disease, vision difficulties (difficulty recognizing familiar faces at a distance of ≤ 4 m, with or without glasses), deafness, dyspnea (classes II, III, or IV of the New York Heart Association classification: breathless with minor effort, during ordinary activity, or at rest), regular use of nonsteroidal antiinflammatory drugs for joint pain, and knee or hip replacement for osteoarthritis. Cardiovascular disease and risk factors included stroke, coronary heart disease, lower-limb arteritis, hypertension (systolic blood pressure ≥ 140 mmHg, diastolic blood pressure ≥ 90 mmHg, or antihypertensive medication), and lipid-lowering drugs as a surrogate for hypercholesterolemia. Expert committees validated incident stroke and coronary heart disease events using hospital and other medical records.²⁷

Statistical analysis

Individuals who were disabled at baseline or had missing data for any of the covariates, for whom disability status was unknown at all waves, or for whom fast gait speed was never

measured during follow-up were excluded. Because fast gait speed was measured between age 65 and 85 only, follow-up for disability was restricted to that range. Disability was assessed at each wave, so its precise date of onset was unknown; the midpoint between the data collection wave at which the participant was deemed to have disability and the previous one was therefore used as the date of disability onset. Follow-up was until Wave 6 for participants who remained disability free and for whom fast gait speed was measured at that wave, date of onset of disability for participants in whom fast gait speed was assessed at the previous wave, and date of last assessment at which fast gait speed was measured for all other participants censored as nondisabled at that date.

Participant characteristics were described as a function of disability status at the end of follow-up. Change in fast gait speed was first examined as a function of disability status using a linear mixed model, including the intercept and slope as random effects and a backward scale for time (in years). In addition to an indicator of disability status at the end of follow-up and its interaction with time, the model was adjusted for age at the end of follow-up, sex, and the interaction between sex and time.

To examine the association between baseline fast gait speed and change in fast gait speed over time and the incidence of disability, a joint modeling approach was used that jointly estimates the parameters of longitudinal and survival submodels (supplementary methods). The intercept and slope (change over time) of the longitudinal submodel were treated as random effects, and sex and baseline age (centered at 65) and their interactions with time were treated as fixed effects. A Weibull function was used for the survival submodel, adjusted for sex and baseline age (centered at 65). Because the independent effects of baseline fast gait speed and change therein on the incidence of disability was of interest, the random intercept (corresponding to individual differences in baseline fast gait speed) and slope (corresponding to individual differences in the slope of change in fast gait speed) were included in the survival submodel to estimate hazard ratios (HRs).^{28,29}

The role of covariates in explaining the association between fast gait speed and disability was assessed by including them in the survival submodel from the baseline assessment (sociodemographic measures, unhealthy behaviors, height) or as time-dependent variables (BMI, MMSE tertiles, depressive symptomatology, trauma, chronic conditions, cardiovascular disease, and risk factors). Models with covariates were compared with one without them; their contribution was estimated as percentage change in HRs as $100 \times (\log HR_{\text{Model1}} - \log HR_{\text{Modeli}}) / \log HR_{\text{Model1}}$.

To exclude reverse causation bias, participants who developed disability in the first 4 (Waves 1–2) and 7 years of follow-up (Waves 1–4) were excluded from sensitivity analyses. The main analyses did not take into account the possibility that some participants may recover from disability; the analysis were repeated by excluding participants who recovered from disability and remained disability free throughout the remaining follow-up.

Analyses were performed using SAS 9.3 (SAS Institute, Inc., Cary, NC) and Stata 12.1 (stjm command; StataCorp LP, College Station, TX); P-values were two-sided and those $\leq .05$ were considered statistically significant.

RESULTS

At study baseline, 379 (8.1%) of 4,692 individuals aged 65 to 85 were disabled and excluded from the analyses. After excluding individuals with missing data for covariates (n=66), disability status (n=25), or fast gait speed (394 participants with missing measures at all waves and 60 with missing measures before they became disabled), the analyses were based on 3,814 participants. Excluded participants (n=499) were older (76.2 vs 73.2, $P < .001$) and less educated (no education to primary school, 38.5% vs 33.8%, age-adjusted $P = .02$) than those included in the analyses.

During a total follow-up of 11 years (mean 5.1 ± 4.2 years), 3,814 participants contributed 20,338 person-years; 628 (68.0% women) developed disability (incidence, 30.9/1,000 person-years). Table 1 shows that participants who developed disability were more likely to be

female, to be married, to walk more slowly, and to be in worse health. In particular, there was a strong association between dyspnea and disability over follow-up (odds ratio (OR)=2.23, $P<.001$). Mean fast gait speed at baseline was 1.54 ± 0.30 m/s. Older and less-educated participants, women, and those in worse health walked more slowly; dyspnea had the strongest effect (beta=-11.65, $P<.001$) (Supplementary Table 1).

In an unadjusted linear mixed model, mean annual decline in fast gait speed was -0.017 m/s. Those who walked faster at baseline experienced somewhat greater decline over follow-up than those who walked more slowly (covariance of random intercept and slope=-8.26, SE=1.93, $P<.001$), although this effect was not strong, as evidenced in the spread of participants in the cross-tabulation of quartiles of the individual predictions (best linear unbiased prediction) of intercepts and slopes (Supplementary Table 2).

Figure 1 presents predicted trajectories of fast gait speed according to disability status at the end of follow-up for 75-year-old women (estimates in Supplementary Table 3). Those who developed disability during follow-up had a 20% ($p<.001$) faster annual decline in fast gait speed (-0.024 m/s, 95% confidence interval (CI)=-0.028, -0.021) than those who remained disability free (-0.020 m/s, 95% CI=-0.022, -0.019).

Table 2 shows the association between fast gait speed at baseline and change in fast gait speed and incidence of disability. In the simplest model (Model 0), the HR of disability per SD (-0.22 m/s) slower fast gait speed at baseline (random intercept) was 1.73 (95% CI=1.57-1.90). In a model with the random intercept and slope (Model 1), the HR of disability per SD was 1.77 (95% CI=1.60-1.94) for slower fast gait speed at baseline and 1.38 (95% CI=1.10-1.73) for 1-SD (-0.013 m/s per year) decrease in the random slope (representing faster decline in fast gait speed). For baseline fast gait speed, the association remained significant in the fully adjusted model (HR=1.54, 95% CI=1.39-1.72); covariates explained 23.6% of the association. The highest percentage reduction was for chronic conditions (10.4%). For change in fast gait speed, the association became nonsignificant in the fully adjusted model

(HR=1.05, 95% CI=0.83–1.32); covariates explained 86.3% of the association. The highest percentage reduction was for chronic conditions (58.9%). Dyspnea was the chronic condition that played the strongest role, for baseline speed (6.6%) and change (42.1%).

Participants who developed disability during the first 4 to 7 years of follow-up were excluded from sensitivity analyses (Table 3). Baseline fast gait speed remained associated with disability, with HRs in the same range as in main analyses. The HR for change in fast gait speed was similar to that of main analyses when excluding the first 4 years and tended to be somewhat stronger when excluding the first 7 years. During follow-up, 178 participants recovered from disability and remained disability free throughout follow-up. Analyses excluding these participants showed associations that were even stronger than those from the main analyses (Supplementary Table 4).

DISCUSSION

Based on a large cohort of French community-dwelling older adults followed over 11 years, slower fast gait speed at baseline and faster decline in fast gait speed thereafter were independent predictors of incident disability. The strength of the association was stronger for fast gait speed at baseline, but HRs were similar to that associated with decline (1.4 to 1.8). Chronic conditions, especially dyspnea, explained the largest part of the associations. Finally, fast gait speed at baseline was associated with incident disability 7 years after, independent of change in fast gait speed over this period.

These findings extend those of previous studies showing that poor physical performance is associated with disability,^{5–8} although few studies have examined the role of change in physical performance. One study in Americans aged 65 and older showed that poor grip strength and slow gait speed 11 years after baseline and faster decline in grip strength and stride length 5 to 11 years after baseline independently predicted three domains of disability (mobility, upper-extremity function, ADLs) 18 years after baseline.¹⁴ Another study in Americans aged 72 and older examined the association between physical performance (gait

speed, timed chair stands, speed of turning) and their change between baseline and Year 1 and ADL disability;¹² decline in physical performance was associated with disability at Year 1 after adjusting for baseline performance but was not associated with disability at Year 3 after adjusting for physical performance at Year 1. These contradictory results may result from the small interval used to determine change in performance and from the long interval between the last gait speed measure and the disability assessment for the second analysis. In addition, change was assessed based on two measures only, which may be subject to regression to the mean. Another study of 93 disabled women aged 65 and older assessed weekly over 24 weeks examined the short-term effect of change in physical function (summary score including gait speed, standing balance, rising from a chair) on ADLs and mobility.¹³ At each time point, baseline and change in physical performance were independently associated with disability. Finally, a study in Italians aged 65 and older reported that the latest assessed lower extremity performance is a strong predictor of incident mobility and ADL disability independent of previous performance.³⁰ That study measured physical performance twice (3 years apart) and examined incident disability 6 and 9 years later, whereas the current study measured fast gait speed five times and disability six times over 11 years. The small interval between the two assessments of physical performance and the long interval between the first measure and the disability assessment may account for inconsistent findings.

Using up to five assessments to estimate change in fast gait speed over 11 years, it was shown that it predicted disability independent of fast gait speed at the start of the study. Therefore, assessment of change in fast gait speed, in addition to single assessments, allows the identification of persons whose fast gait speed is initially fine but experience faster decline over follow-up as being at risk of disability. Although those who walked faster tended to have a more-pronounced decline in fast gait speed over follow-up, the correlation was not strong, and participants were relatively evenly distributed across quartiles of initial fast gait speed and change in fast gait speed.

Disability is a long-term process that results from the accumulation of deficits over time.

Thus, fast gait speed at study baseline may be a stronger predictor of disability occurring early during follow-up. This hypothesis was tested by excluding cases of disability arising in the 4 to 7 years after the start of the study, and it was found that baseline fast gait speed continued to predict disability at least 7 years later, independent of change in fast gait speed. Therefore, slower fast gait speed is an early marker of future disability and is present several years before disability onset.

The covariates played a stronger role in explaining the association between disability and change in fast gait speed than with baseline fast gait speed. Baseline covariates do not capture all of the lifecourse exposure history of participants before the baseline assessment, whereas changes in covariates over the study period are likely to better capture change in important confounders after the baseline assessment; therefore, it is likely that residual confounding is more important for the baseline association than for the association with change. Multiple factors are likely to be involved in the association between fast gait speed and disability.

Dyspnea had the strongest association with fast gait speed and disability and the strongest role in explaining their association. Dyspnea may result from heart failure, chronic obstructive pulmonary disease, and obesity, all of which are risk factors for slow gait speed and disability.^{24,31} The results of the current study confirm that better prevention or management of chronic diseases may reduce the risk of disability.

These findings need to be considered in light of some limitations. First, joint models cannot accommodate the interval-censored nature of these data, requiring the use of the midpoint between the wave at which disability was reported and the previous wave to define the date of disability onset. Nevertheless, this approach is reasonable when intervals between assessments are not too large.³² Second, the possibility that some people may recover from disability was not taken into account, although analyses excluding these people did not change the conclusions. Third, fast rather than usual gait speed was used for the analyses,

mainly because the former has a steeper decline and larger interindividual heterogeneity than the latter; these results therefore apply to this measure only. Although clinicians may use it less often than usual gait speed, fast gait speed has been implemented in many studies and settings and is simple to measure; the findings are in favor of its clinical value. Fourth, participants excluded from the analyses were older at baseline and less educated but not significantly different from other participants in terms of sex and marital status. Although the absolute risk of disability was probably underestimated, this would be unlikely to bias the estimate of the associations. Fifth, because fast gait speed was measured only between the ages of 65 and 85, the findings cannot be generalized beyond this age range. This study's main strengths include its large size and long follow-up with up to six disability and five fast gait speed measures, allowing a precise estimate of fast gait speed decline to be obtained. Physical performance was assessed using an objective and highly reproducible measure. Joint models have several advantages over time-dependent Cox proportional hazard models. By modeling the longitudinal variable as a random process (fast gait speed), they allow more-precise examination of the association between the longitudinal process and outcome by taking measurement error into account.³³ These models are remarkably flexible, whereas time-dependent Cox proportional hazards models allow only the association between the current value of the longitudinal process and the incidence of the outcome to be examined, not allowing the analysis of change in exposure.

In conclusion, in French older adults, slower fast gait speed at baseline was a robust predictor of disability, even assessed 7 years before disability. Moreover, change in fast gait speed was associated with disability onset over 11 years of follow-up, independent of initial fast gait speed. These results show the importance of repeated physical and disability assessments in older adults to better identify those at higher risk of becoming disabled.

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Author Contributions: Artaud, Elbaz: study design and research. Artaud: data analysis. Dugravot: statistical analysis. Artaud, Singh-Manoux, Elbaz: writing of the paper. Tzourio, Elbaz: data acquisition. Artaud, Elbaz: primary responsibility for final content. All authors read and approved the final manuscript.

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Table 1. Participant Characteristics, Overall and According to Disability Status at the End of Follow-Up

Characteristic	Overall, N=3,814	Not Disabled, n=3,186 (83.5%)	Disabled, n=628 (16.5%) ^a	Odds Ratio (95% Confidence Interval) ^b
Covariates at study baseline				
Age, mean±SD	73.2±4.6	73.3±4.7	73.0±4.0	0.94 (0.86–1.02)
Men, n (%)	1,493 (39.1)	1,292 (40.6)	201 (32.0)	0.69 (0.58–0.83)
Marital status, n (%)				
Married	2,287 (60.0)	1,897 (59.5)	390 (62.1)	1.87 (1.29–2.70)
Divorced, separated, or widowed	1,211 (31.8)	1,009 (31.7)	202 (32.2)	1.55 (1.06–2.26)
Single	316 (8.3)	280 (8.8)	36 (5.7)	Ref.
Education, n (%)				
No education or primary school	1,289 (33.8)	1,072 (33.6)	217 (34.6)	1.04 (0.84–1.28)
Secondary school	1,235 (32.4)	1,026 (32.2)	209 (33.3)	1.04 (0.84–1.29)
High-school or university degree	1,290 (33.8)	1,088 (34.1)	202 (32.2)	Ref.
Number of unhealthy behaviors, n (%) ^c				
0	233 (6.1)	203 (6.4)	30 (4.8)	Ref.
1	958 (25.1)	818 (25.7)	140 (22.3)	1.17 (0.76–1.78)
2	1,563 (41.0)	1,301 (40.8)	262 (41.7)	1.40 (0.93–2.11)
3 or 4	916 (24.0)	750 (23.5)	166 (26.4)	1.57 (1.03–2.39)
Height, cm, mean±SD	161.9±8.7	162.1±8.7	160.8±8.7	0.95 (0.84–1.07)
Fast gait speed, m/s, mean±SD	1.54±0.30	1.55±0.30	1.49±0.30	0.84 (0.76–0.93)
Time-dependent covariates				
Body mass index, kg/m ² , mean±SD ^d	25.8±3.9	25.7±3.9	26.2±4.1	1.17 (1.08–1.27)
Mini-Mental State Examination score, mean±SD ^d	27.4±1.7	27.4±1.7	27.4±1.8	0.96 (0.88–1.04)
Depressive symptoms, n (%) ^e	1,227 (32.2)	968 (30.4)	259 (41.2)	1.52 (1.27–1.82)
Psychotropic drug use, n (%) ^e	1,457 (38.2)	1,177 (36.9)	280 (44.6)	1.30 (1.09–1.54)
Bone fracture, n (%) ^e	530 (13.9)	424 (13.3)	106 (16.9)	1.24 (0.98–1.57)
Falls, n (%) ^e	1,009 (26.5)	799 (25.1)	210 (33.4)	1.44 (1.19–1.73)
Diabetes mellitus, n (%) ^e	462 (12.1)	369 (11.6)	93 (14.8)	1.41 (1.10–1.81)
Parkinson's disease, n (%) ^e	70 (1.8)	50 (1.6)	20 (3.2)	2.15 (1.27–3.64)
Vision difficulties, n (%) ^e	583 (15.3)	457 (14.3)	126 (20.1)	1.45 (1.16–1.81)
Deafness, n (%) ^e	631 (16.5)	508 (15.9)	123 (19.6)	1.39 (1.11–1.74)
Dyspnea, n (%) ^e	743 (19.5)	543 (17.0)	200 (31.8)	2.23 (1.84–2.70)
Nonsteroidal antiinflammatory drugs for joint pain, n (%) ^e	826 (21.7)	663 (20.8)	163 (26.0)	1.28 (1.05–1.56)
Knee or hip replacement for osteoarthritis, n (%) ^e	337 (8.8)	250 (7.8)	87 (13.9)	1.89 (1.46–2.46)
Cardiovascular disease, n (%) ^{e,1}	834 (21.9)	664 (20.8)	170 (27.1)	1.55 (1.27–1.89)
Hypertension, n (%) ^e	3,301 (86.5)	2,741 (86.0)	560 (89.2)	1.41 (1.08–1.86)
Hypercholesterolemia, n (%) ^e	1,716 (45.0)	1,411 (44.3)	305 (48.6)	1.18 (0.99–1.40)

^aDependent according to the Rosow and instrumental activity of daily living scales with or without the activity of daily living scale (dichotomized hierarchical indicator of disability).

^bFrom logistic regression of disability status adjusted for sex; per 1-standard deviation (SD) increase for quantitative covariates.

^cThe score of unhealthy behaviors was missing for 144 (4%) participants. An additional “missing” category was included for this variable rather than deleting these subjects.

^dMean of all measures taken during follow-up.

^eAt least one report over the follow-up.

^fStroke, coronary heart disease, or lower-limb arteritis.

Table 2. Incidence of Disability in Relation to Fast Gait Speed at Study Baseline and Change in Fast Gait Speed Over Follow-Up: Analyses Based on a Joint Model

Model	Fast Gait Speed at Baseline ^a	Slope of Change in Fast Gait Speed ^b
	HR (95% Confidence Interval)	% Change ^c
0 ^{d,e}	1.73 (1.57–1.90)	
1 ^{e,f}	1.77 (1.60–1.94)	1.38 (1.10–1.73)
1 + sociodemographic characteristics ^g	1.81 (1.64–2.00) –4.1	1.43 (1.14–1.79) –10.9
1 + number of unhealthy behaviors	1.72 (1.56–1.90) 4.6	1.36 (1.08–1.71) 4.5
1 + anthropometric characteristics ^h	1.76 (1.59–1.94) 0.9	1.37 (1.08–1.75) 1.6
1 + cognitive function ⁱ	1.81 (1.64–2.00) –4.2	1.45 (1.14–1.85) –14.9
1 + depressive symptoms, psychotropic drugs ^j	1.77 (1.60–1.95) –0.3	1.35 (1.05–1.73) 6.9
1 + trauma ^k	1.83 (1.66–2.02) –6.4	1.40 (1.11–1.77) –4.3
1 + chronic conditions ^l	1.66 (1.51–1.84) 10.4	1.14 (0.90–1.45) 58.9
1 + cardiovascular disease and its risk factors ^m	1.78 (1.62–1.97) –1.7	1.37 (1.08–1.73) 3.5
Fully adjusted ⁿ	1.54 (1.39–1.72) 23.6	1.05 (0.83–1.32) 86.3

^aPer 1-standard deviation (SD) decrease in the random intercept (0.22 m/s).

^bPer 1-SD decrease in the random slope (0.013 m/s per year).

^cPercentage change = $100 \times (\log \text{hazard ratio (HR)}_{\text{Model } i} - \log \text{HR}_{\text{Model } 0}) / \log \text{HR}_{\text{Model } 0}$.

^dModel 0 included only the random intercept in the survival submodel.

^eAdjusted for sex (reference female) and age at baseline (centered at 65).

^fModel 1 included the random intercept and slope in the survival submodel.

^gBaseline marital status and education.

^hBaseline height and time-dependent body mass index.

ⁱMini-Mental State Examination score (time-dependent).

^jCenter for Epidemiologic Studies Depression Scale or use of psychotropic drugs (time-dependent).

^kBone fracture, falls (time-dependent).

^lDiabetes mellitus, Parkinson's disease, vision difficulties, deafness, dyspnea, nonsteroidal antiinflammatory drug use for joint pain, knee or hip replacement for osteoarthritis (time-dependent).

^mStroke, coronary heart disease, lower-limb arteritis, hypertension, hypercholesterolemia (time-dependent).

ⁿAll covariates are included in the model.

Table 3. Incidence of Disability in Relation to Fast Gait Speed at Study Baseline and Change in Fast Gait Speed Over Follow-up: Analyses Based on a Joint Model Excluding the First 4 and 7 Years of Follow-Up

Model	Baseline Fast Gait Speed ^a	Change in Fast Gait Speed ^b
	HR (95% Confidence Interval)	% Change ^c
Exclusion of the first 4 years ^d		
Model 1 ^e	1.74 (1.51–2.01)	1.32 (0.98–1.78)
Fully adjusted model ^f	1.61 (1.37–1.89) 14.3	1.04 (0.76–1.42) 85.1
Exclusion of the first 7 years ^g		
Model 1 ^e	1.65 (1.26–2.17)	1.61 (1.04–2.50)
Fully adjusted model ^f	1.59 (1.17–2.17) 7.3	1.30 (0.79–2.12) 45.4

^aPer 1-standard deviation (SD) decrease in the random intercept (0.22 m/s).

^bPer 1-SD decrease in the random slope (0.013 m/s per year).

^cPercentage change = $100 \times (\log \text{hazard ratio (HR)}_{\text{Model 1}} - \log \text{HR}_{\text{Model i}}) / \log \text{HR}_{\text{Model 1}}$.

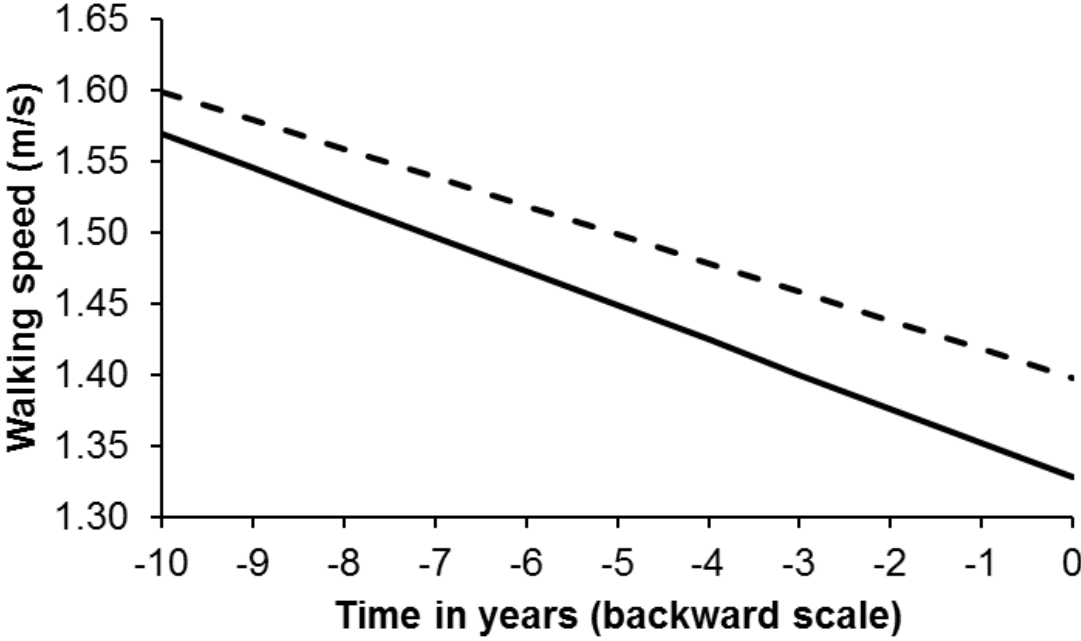
^dBased on 1,849 participants (n=1,112 women, 60.1%), of whom 272 (n=181 women, 66.5%) developed disability.

^eAdjusted for sex (reference female) and age at baseline (centered at 65).

^fAll covariates included in the model. (See Table 2 for a description of the covariates.)

^gBased on 1,211 participants (n=726 women, 60.0%), of whom 77 (n=52 women, 67.5%) developed disability.

Figure 1. Predicted trajectories of fast gait speed in disabled (solid line) and not disabled (dashed line) women aged 75 at the end of follow-up. Fast gait speed was modeled using a linear mixed model including a random intercept and slope with a backward time scale. Estimates from this model are presented in Supplementary Table 3.



Supplementary methods

Disability

Rosow and Breslau scale (mobility)

Questions	Score
A. Able to do heavy work around the house like washing windows, walls, or floors without help	
Yes	0
No	1
B. Able to walk between 500 m and 1 km without help	
Yes	0
No	1
C. Able to walk up and down stairs to the second floor without help	
Yes	0
No	1

Katz ADL scale

Questions	Score
Bathing	
Bathes self completely	0
Needs help in bathing only a single part of the body such as the back, genital area or disabled extremity	0
Needs help with bathing more than one part of the body, or requires total bathing	1
Dressing	
Gets clothes from closets and drawers and puts on clothes and outer garments complete with fasteners	0
May have help tying shoes	0
Needs help with dressing self or needs to be completely dressed	1
Toileting	
Goes to toilet, gets on and off, arranges clothes, cleans genital area without help	0
Needs help transferring to the toilet, cleaning self or uses bedpan or commode	1
Does not go to the toilet	1
Transferring	
Moves in and out of bed or chair unassisted. Mechanical transferring aides are acceptable	0
Needs help in moving from bed to chair or requires a complete transfer	1
Does not leave the bed	1
Feeding	
Gets food from plate into mouth without help	0
Needs help to cut the meal or to butter bread	0
Needs total help with feeding or requires parenteral feeding	1

Lawton-Brody IADL scale

Questions	Score
A. Ability to use telephone	
1. Operates telephone on own initiative; looks up and dials numbers, etc.	0
2. Dials a few well-known numbers	0
3. Answers telephone but does not dial	0
4. Does not use telephone at all	1
B. Shopping	
1. Takes care of all shopping needs independently	0
2. Shops independently for small purchases	1
3. Needs to be accompanied on any shopping trip	1
4. Completely unable to shop	1
C. Mode of Transportation	
1. Travels independently on public transportation or drives own car	0
2. Arranges own travel via taxi, but does not otherwise use public transportation	0
3. Travels on public transportation when accompanied by another	0
4. Travel limited to taxi or automobile with assistance of another	1
5. Does not travel at all	1
D. Responsibility for own medications	
1. Is responsible for taking medication in correct dosages at correct time	0
2. Takes responsibility if medication is prepared in advance in separate dosage	1
3. Is not capable of dispensing own medication	1
E. Ability to Handle Finances	
1. Manages financial matters independently (budgets, writes checks, pays rent, bills goes to bank)	0
2. Manages day-to-day purchases, but needs help with banking, major purchases, etc.	0
3. Incapable of handling money	1
For women only	
F. Food Preparation	
1. Plans, prepares and serves adequate meals independently	0
2. Prepares adequate meals if supplied with ingredients	1
3. Able to heat the dishes already prepared	1
4. Needs to have meals prepared and served	1
G. Housekeeping	
1. Maintains house alone or with occasional assistance (e.g. "heavy work domestic help")	0
2. Performs light daily tasks such as dishwashing, bed making	0
3. Performs light daily tasks but cannot maintain acceptable level of cleanliness	0
4. Needs help with all home maintenance tasks	0
5. Does not participate in any housekeeping tasks	1
H. Laundry	
1. Does personal laundry completely	0
2. Launders small items; rinses stockings, etc.	0
3. All laundry must be done by others	1

Statistical methods

Joint model

We jointly estimated the parameters of a longitudinal and a survival sub-model:

- *longitudinal sub-model:*

$$WS_{ij} = \beta_0 + \beta_1 t_{ij} + \beta_2 sex_i + \beta_3 age_i + \beta_4 sex_i \times t_{ij} + \beta_5 age_i \times t_{ij} + b_{0i} + b_{1i} t_{ij} + \varepsilon_{ij},$$

where β represents the regression coefficients for fixed effects, b the coefficient for the random effects, ε_{ij} denotes measurement error;

- *survival sub-model:* $h_i(t) = h_0(t) \exp\{\gamma_1 sex_i + \gamma_2 age_i + \alpha_0 b_{0i} + \alpha_1 b_{1i}\}$, where $h_0(t)$ is the baseline hazard function, γ the logarithm of the hazard ratio (HR) of disability associated with covariates, and (α_0, α_1) are the parameters of interest here and correspond to the logarithm of the HR of disability associated with the random intercept and slope respectively (association parameters). We also estimated the HR of disability associated with baseline fast gait speed, through a simpler model that only included the random intercept.

Supplementary Table 1. Participants' Characteristics and their Association with Fast Gait Speed at Study Baseline

Baseline characteristics		Overall ^a	Fast gait speed (m/s) (SD)	Beta (95% CI) ^b	
N (%)		3532	1.54 (0.30)		
Age	<70	799 (22.6)	1.69 (0.30)	Ref.	
	[70 – 74[1005 (28.5)	1.57 (0.29)	-0.10 (-0.12, -0.07)	
	[74 – 78[917 (26.0)	1.51 (0.30)	-0.16 (-0.18, -0.13)	
	≥ 78	811 (23.0)	1.42 (0.29)	-0.24 (-0.27, -0.21)	
Sex	Men	1376 (39.0)	1.69 (0.30)	0.23 (0.22, 0.25)	
	Women	2156 (61.0)	1.45 (0.27)	Ref.	
Marital status	Married	2125 (60.2)	1.59 (0.30)	-0.01 (-0.04, 0.02)	
	Divorced, separated, widowed	1113 (31.5)	1.47 (0.29)	-0.02 (-0.06, 0.01)	
	Single	294 (8.3)	1.50 (0.31)	Ref.	
Education	No education/primary school	1184 (33.5)	1.46 (0.28)	-0.14 (-0.16, -0.12)	
	Secondary school	1159 (32.8)	1.53 (0.30)	-0.08 (-0.10, -0.06)	
	High-school/university	1189 (33.7)	1.64 (0.30)	Ref.	
No of unhealthy behaviours ^c	0	217 (6.4)	1.61 (0.28)	Ref.	
	1	896 (26.2)	1.59 (0.29)	-0.03 (-0.07, 0.01)	
	2	1457 (42.7)	1.52 (0.31)	-0.10 (-0.13, -0.06)	
	3 or 4	845 (24.7)	1.51 (0.30)	-0.12 (-0.16, -0.08)	
BMI (kg/m ²)	Normal	1678 (47.5)	1.57 (0.31)	Ref.	
	Overweight	1383 (39.2)	1.54 (0.30)	-0.08 (-0.09, -0.06)	
	Obese	471 (13.3)	1.43 (0.29)	-0.18 (-0.20, -0.15)	
Height (cm)	Men	Women			
	<165	<153	859 (24.3)	1.49 (0.31)	Ref.
	[165 ; 170[[153 ; 157[842 (23.8)	1.51 (0.29)	0.02 (-0.01, 0.04)
	[170 ; 174[[157 ; 161[894 (25.3)	1.56 (0.30)	0.06 (0.03, 0.08)
	≥174	≥161	937 (26.5)	1.60 (0.30)	0.09 (0.06, 0.11)
MMSE score	< 27		840 (23.8)	1.47 (0.31)	Ref.
	[27 ; 28[613 (17.4)	1.52 (0.27)	0.03 (-0.0004, 0.05)
	≥ 28		2079 (58.9)	1.58 (0.30)	0.08 (0.06, 0.10)
Depressive symptoms	Yes		767 (21.7)	1.43 (0.28)	-0.09 (-0.11, -0.07)
	No		2765 (78.3)	1.57 (0.30)	Ref.

Supplementary Table 1 (continued).

Baseline characteristics		Overall ^a	Fast gait speed (m/s) (SD)	Beta (95% CI) ^b
Psychotropic drugs	Yes	848 (24.0)	1.44 (0.29)	-0.08 (-0.10, -0.06)
	No	2684 (76.0)	1.57 (0.30)	Ref.
Bone fracture	Yes	232 (6.6)	1.46 (0.30)	-0.03 (-0.06, 0.01)
	No	3300 (93.4)	1.55 (0.30)	Ref.
Falls	Yes	204 (5.8)	1.40 (0.28)	-0.08 (-0.12, -0.05)
	No	3328 (94.2)	1.55 (0.30)	Ref.
Diabetes	Yes	263 (7.4)	1.52 (0.31)	-0.06 (-0.10, -0.03)
	No	3269 (92.6)	1.54 (0.30)	Ref.
Parkinson's disease	Yes	33 (0.9)	1.49 (0.28)	-0.08 (-0.16, 0.01)
	No	3499 (99.1)	1.54 (0.30)	Ref.
Vision difficulties	Yes	180 (5.1)	1.43 (0.30)	-0.04 (-0.08, -0.004)
	No	3352 (94.9)	1.55 (0.30)	Ref.
Deafness	Yes	290 (8.2)	1.51 (0.28)	-0.03 (-0.07, -0.0003)
	No	3242 (91.8)	1.54 (0.31)	Ref.
Dyspnea	Yes	458 (13.0)	1.40 (0.28)	-0.12 (-0.14, -0.09)
	No	3074 (87.0)	1.56 (0.30)	Ref.
NSAIDs for joint pain	Yes	510 (14.4)	1.44 (0.30)	-0.08 (-0.11, -0.06)
	No	3022 (85.6)	1.56 (0.30)	Ref.
Knee/hip replacement for osteoarthritis	Yes	159 (4.5)	1.43 (0.28)	-0.10 (-0.14, -0.06)
	No	3373 (95.5)	1.55 (0.30)	Ref.
Cardiovascular disease ^d	Yes	552 (15.6)	1.50 (0.31)	-0.08 (-0.11, -0.06)
	No	2980 (84.4)	1.55 (0.30)	Ref.
Hypertension	Yes	2796 (79.2)	1.53 (0.31)	-0.06 (-0.08, -0.04)
	No	736 (20.8)	1.59 (0.29)	Ref.
Hypercholesterolemia	Yes	1196 (33.9)	1.52 (0.29)	-0.02 (-0.04, -0.003)
	No	2336 (66.1)	1.55 (0.31)	Ref.
Disability at the end of follow-up	Yes	575 (16.3)	1.49 (0.30)	-0.04 (-0.07, -0.02)
	No	2957 (83.7)	1.55 (0.30)	Ref.

^a Based on participants with a baseline fast gait speed measure.

^b Estimates of the regression coefficient (95% CI) from a linear regression of fast gait speed (dependent variable) on covariates, adjusted for age and sex.

^c The score of unhealthy behaviors was missing for 144 (4%) participants. We included an additional 'missing' category for this variable rather than deleting these subjects.

^d Stroke, coronary heart disease, or lower-limb arteritis.

Supplementary Table 2. Cross-Tabulation of Sex-Specific Quartiles of the Predicted Intercept (Representing Baseline Fast Gait Speed) and Slope (Representing Change in Fast Gait Speed) of a Non-Adjusted Linear Mixed Model of Fast Gait Speed

		Quartiles of the predicted intercept of fast gait speed (m/s)				
Quartiles of the predicted slope of change in fast gait speed (m/s per year)		<i>Slower</i>		<i>Faster</i>		
Men	Women	Men	Women	Men	Women	
<i>Smaller decline</i>	≥ -0.016	≥ -0.014	< 1.51	$[1.51; 1.64]$	$[1.64; 1.83]$	≥ 1.83
	$[-0.018; -0.016[$	$[-0.016; -0.014[$	< 1.32	$[1.32; 1.47]$	$[1.47; 1.89]$	≥ 1.59
	$[-0.021; -0.018[$	$[-0.019; -0.016[$				
<i>Greater decline</i>	< -0.021	< -0.019				
			N=404 (10.6%)	N=149 (3.9%)	N=159 (4.2%)	N=200 (5.2%)
			N=341 (8.9%)	N=422 (11.1%)	N=141 (3.7%)	N=96 (2.5%)
			N=115 (3.0%)	N=198 (5.2%)	N=447 (11.7%)	N=188 (4.9%)
			N=103 (2.7%)	N=175 (4.6%)	N=207 (5.4%)	N=469 (12.3%)

Each individual's baseline fast gait speed was estimated as the sum of the fixed effects for the intercept and of the random intercept using best linear unbiased prediction (BLUPs); individual slopes of change in fast gait speed were estimated in a similar way.

**Supplementary Table 3. Linear Mixed Model Estimates of Change in Fast Gait Speed
According to Disability Status at the End of the Follow-up**

Characteristics	Estimate (95% CI)
Intercept ^a	1.398 (1.385, 1.411)
Age (centered at 75y)	-0.010 (-0.012, -0.009)
Sex (men vs. women)	0.215 (0.198, 0.232)
Disability (disabled vs. not disabled)	-0.070 (-0.093, -0.048)
Time	-0.020 (-0.022, -0.019)
Time × Sex	-0.003 (-0.005, -0.001)
Time × Disability status	-0.004 (-0.007, -0.001)

^aThe intercept corresponds to the mean fast gait speed (m/s) at the end of the follow-up for women not disabled and aged 75y.

Supplementary Table 4. Incidence of Disability in Relation to Fast Gait Speed at Study Baseline and Change in Fast Gait Speed over the Follow-up: Exclusion of 178 Participants who Recovered from Disability and Remained Disability-free during the Follow-up

Model	Baseline fast gait speed ^a		Change in fast gait speed ^b	
	HR (95% CI)	% change ^c	HR (95% CI)	% change ^c
<i>Model 1</i> ^d	1.99 (1.77, 2.24)	--	1.52 (1.16, 2.01)	--
<i>Fully adjusted model</i> ^e	1.72 (1.52, 1.95)	21.1	1.09 (0.82, 1.43)	80.5

Analyses are based on 3636 participants (women, n=2205, 60.6%), of whom 450 (women, n=311, 69.1%) developed disability.

^a Per one-SD decrease in the random intercept (0.22 m/s).

^b Per one-SD decrease in the random slope (0.013 m/s per year).

^c Percentage change = $100 \times (\log HR_{\text{Model } i} - \log HR_{\text{Model } 1}) / \log HR_{\text{Model } 1}$.

^d Adjusted for sex (reference, women) and age at baseline (centered at 65 years).

^e All covariates are included in the model (see table 2 for a description of the covariates).