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Depressive Symptoms and Gait Dysfunction in the Elderly

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Abstract

Objective—Assess the association between depressive symptoms (not meeting the criteria for major depression) and gait dysfunction in older adults.

Design—Cross-sectional study.

Setting—Einstein Aging Study, a community-based longitudinal aging study.

Participants—Six hundred ten nondemented and nondepressed community-residing adults age 70 and older.

Measurements—Depressive symptoms measured using the 15-item Geriatric Depression Scale. To obtain a comprehensive assessment of gait, eight individual quantitative gait parameters were assessed: velocity (cm/s), stride length (cm), cadence (steps/min), swing phase (seconds), stance phase (seconds), double support phase (seconds), stride length variability (SD of stride length), and swing time variability (SD of swing time). Multiple linear regression analysis was applied to study the association of depressive symptoms with gait, adjusting for potential confounders including demographic variables, medical illnesses, and clinical gait abnormalities.

Results—Increased level of depressive symptoms was associated with worse velocity, stride, and swing time variability. The relationship of the remaining five gait variables with depressive symptoms was not significant in the fully adjusted models.

Conclusions—Higher levels of depressive symptoms are associated with worse performance in specific quantitative gait variables in community-residing older adults.

Keywords

depressive symptoms; epidemiology; elderly; gait

It is estimated that about 14% of the U.S. population of age 65 and older suffer from depressive symptoms that do not meet criteria for major depressive disorder (MDD), which is approximately 2.5 times higher than the prevalence of MDD.1 In addition to the associated risk of converting to MDD,2 depressive symptoms in older adults have been

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reported to be associated with increased risk of disability,3 decreased quality of life, and result in increased societal costs due to higher health care needs.4

Gait performance is recommended as a clinical marker of health and function in older adults, 5,6 and predicts multiple adverse health outcomes including disability, falls, dementia, and death in our and other studies.5^{,7},8 Like depressive symptoms, the prevalence of gait disorders is high in older populations.6 Although slowing of gait in clinical depression is well established,9-12 little is known about the nature of gait dysfunction in elderly without MDD. It is unclear from studies that have included both depressed and nondepressed elderly4,13 whether the observed associations between gait and depressive symptoms are explained by the presence of subjects with MDD in these samples. Thus, these studies do not establish whether gait dysfunction occurs in the earliest or mildest stages of depressive symptoms. On the contrary, previous studies of gait in adults with depressive symptoms but who did not meet criteria for MDD have been mostly limited to older adults in hospital or nursing homes9^{,10,12} or younger adults.9⁻¹² Older individuals with depressive symptoms have been reported to have slow gait velocity.13 However, gait is a complex motor behavior with many other measurable facets besides velocity. For instance, gait variability was a stronger predictor of falls than velocity and was the only predictor of injurious falls in our cohort.7

A more comprehensive examination of gait performance in older adults may not only increase our understanding of brain substrates and pathologic processes that are common to gait and depressive symptoms but may also provide insights into new interventions. To address limitations of previous studies, we conducted a cross-sectional study of the relationship between depressive symptoms and gait function, in an ambulatory community-residing sample of 610 older adults (age 70 and older) who were free of dementia and MDD.

METHODS

Sample

Subjects were participants in the Einstein Aging Study.14 The primary aim of the Einstein Aging study is to identify risk factors for cognitive decline. Study design and methods have been previously reported.6 In brief, potential subjects (age 70 and older) identified from population lists of Bronx County were contacted by a letter that explained the purpose and nature of the study and then by telephone. The telephone interview included verbal consent, medical history questionnaire, and cognitive screening tests.7 Exclusion criteria included severe audiovisual loss, bed bound because of illness, and institutionalization. After the interview, an age-stratified sample of subjects who matched on a computerized randomization procedure was invited for further evaluation at our clinical research center at the Albert Einstein College of Medicine. Informed consent was obtained at enrollment according to protocols approved by the local institutional review board. Subjects returned for yearly follow-up assessments.

Mobility Assessment

Quantitative Gait Assessment—Subjects received quantitative gait assessments at baseline and at each annual follow-up visit using a computerized walkway ($180.0 \times 35.5 \times 0.25$ inches) with embedded pressure sensors (GAITRite; CIR Systems. Havertown, PA). Research assistants asked subjects to walk on the instrumented mat at their normal pace for two trials in a quiet well-lit hallway wearing comfortable footwear and without any attached monitoring devices. White lines on the floor marked start and stop points, which included three feet from the walkway edge for initial acceleration and terminal deceleration. On the basis of footfalls recorded on the walkway, the software automatically computes gait

parameters as the mean of two trials. The following eight gait variables that have been reported to have associations with cognitive function and dementia as well as falls in our previous studies were selected for analysis:7¹⁵ velocity (cm/s), stride length (cm), cadence (steps/min), swing phase (seconds), stance phase (seconds), double support phase (seconds), stride length variability (standard deviation; SD of stride length), and swing time variability (SD of swing time). The GAITRite system is widely used in clinical research settings, and excellent reliability has been reported in our and other studies.7¹⁶

Clinical Gait Assessment—Study clinicians clinically evaluated gait at each visit as part of the standard neurologic examination, blinded to the quantitative assessment. Gait was determined to be normal or abnormal due to neurologic (such as hemiparesis or Parkinson disease) or nonneurologic (such as arthritis or joint deformities) causes after visual inspection of walking patterns. More detailed descriptions and Web links to videos of individual gait subtypes are available in our previous publications.6^{,14} The clinical gait assessment has established test–retest reliability and predictive validity for outcomes such as dementia and falls.6^{,14}

Depressive Symptoms

Participants were screened for depressive symptoms with the 15-item Geriatric Depression Scale (GDS).17¹8 The GDS has been reported to have Cronbach's α coefficient of 0.94 and a split-half reliability coefficient of 0.9419 in elderly populations. The GDS has been utilized to describe the relationship of depressive symptoms with other health outcomes in older adults including mortality20 as well as risk of developing major depression21 and falls.22

Study Sample

Between June 2004 and February 2010, 723 Einstein Aging Study participants received detailed clinical and quantitative motoric assessments as part of a gait and mobility substudy.6·7 Reasons for not obtaining assessments included clinician or tester unavailability for conduction motoric assessments, subject illness, or subject refusal. Subjects who were not included were similar in terms of age, sex, and educational status at enrollment.

Of the 723 subjects, 41 with prevalent dementia and 70 with MDD (defined as self-reported history of MDD and previously validated GDS cutscore of 10 or higher to capture undiagnosed cases of MDD23) were excluded. Two additional subjects with missing GDS scores were excluded. Thus, 610 older adults were eligible for this analysis. The subjects who were excluded had significantly different GDS scores but were not significantly different in age, sex, race, and education compared to eligible subjects.

Covariates

Study subjects were interviewed using structured questionnaires at baseline and annual follow-up study visits about sociodemographic variables, medical illnesses, medications, and activities of daily living. Details were verified with family members and caregivers when available. History of diabetes mellitus, heart failure, hypertension, angina pectoris, myocardial infarction, strokes, Parkinson disease, chronic obstructive lung disease, and arthritis reported by subjects at baseline gait visit or at any previous Einstein Aging Study visit was used to calculate a summary illness index (range: 0–9) as previously reported.24 Medical records and primary care providers were also consulted to verify or obtain additional information.6 Any use of antidepressant, antipsychotic, antianxiety, or antimanic drugs by subjects was summarized as "psychotropic use." Research assistants recorded blood pressure, height, and weight using standard protocols. Mean arterial pressure was

calculated as the sum of diastolic blood pressure and one-third pulse pressure (systolic– diastolic blood pressure). General cognitive status was assessed with the Blessed-Information-Memory-Concentration test.25 Participation in 10 common physical leisure activities (tennis, golf, swimming, bicycling, dancing, brisk walking, yoga, aerobics, weight training, and bowling) was assessed using our validated and reliable Physical Activity Scale (PAS).26 For each activity, subjects received two points for participating two or more days per week; one point for weekly participation; and zero points for participating occasionally or never. We summed individual activity points for each individual activity to generate a summary PAS score.26

Data Analysis

Multiple linear regression analysis was used to investigate the association between depressive symptoms and each of the eight gait parameters (variability measures were log transformed to account for skewed distributions), adjusting for age, sex, race, and education. Associations are reported as parameter estimates with 95% confidence intervals (CIs). The Benjamini and Hochberg's approach27 was used to control the false discovery rate and p-values for individual gait variables that remained significant (Table 1) were indicated.

In additional analyses (Model 2), adjustments for the following potential confounders were applied: medical illness index, presence of clinical gait abnormalities, PAS score, Blessed test score, mean arterial pressure, body mass index (BMI), and psychotropic medication use. Missing values appeared random and showed no evidence of being related to gait using multivariate logistic regression. Thus, the main analysis for Model 2 is based on the 503 subjects without missing covariate data. Regression diagnostics for all models were examined analytically and graphically and were adequately met.

All statistical analyses were performed in STATA 10.1 (StataCorp LP, College Station, TX, 2009).

RESULTS

Baseline characteristics of study subjects are presented in Table 2. The sample was predominantly female (61%), well-educated (mean [SD]: 14.0 [3.4] years), mostly Caucasian (67%), with mean age 80.3 (SD: 5.5) years and a mean GDS score of 2.0 (SD: 1.9).

Results of linear regression analysis of the association between depressive symptoms and individual gait parameters are presented in Table 1. In the basic model adjusting for age, sex, race, and years of education, higher GDS scores was associated with worse performance on all gait measures except stride length variability. A one-point increase in the GDS corresponded to decreases of 2.98 cm/s in velocity, 2.39 cm in stride length, and 1.17 steps/min in cadence; and increases of 0.01 seconds in stance time, 0.003 seconds in swing time, 0.01 seconds in double support time, and 0.06 in the log transformed swing time SD (seconds).

Upon further adjustment for additional covariates including in Model 2 the relationship with depressive symptoms remained significant for the velocity, stride length, and swing time variability variables only (Table 1 and Figure 1).

CONCLUSIONS

The findings of this study show a significant association between depressive symptoms (in the absence of MDD) and quantitative gait dysfunction in a large, well characterized cohort

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of community-dwelling nondemented older adults. Increasing numbers of depressive symptoms in this sample was associated with worse performance on seven out of the selected eight gait measures in our study after adjusting for demographic variables. The association with depressive symptoms remained significant for velocity, stride length and swing time variability even after adjusting for several additional confounders such as vascular risk factors, overall disease burden, physical activity levels, and cognitive status. These results suggest that depressive symptoms have specific rather than global effects on gait performance in older adults.

To our knowledge, this is the first study to systematically examine the association of depressive symptoms with comprehensive gait assessments in community residing older adults. Our results are supported by previous studies that have examined the relationship between depressive symptoms and other aspects of mobility.4,12,13,28 Increasing levels of depressive symptoms, not meeting the criteria for MDD, were predictive of greater decline in standing balance, walking speed, and chair rise performance in a cohort of noninstituionalized older adults, age 71 and older.4 Low performance on tests of executive function, slower gait speed, and depressive symptoms has been identified to coexist in community-dwelling nondemented elderly.13 Differences in double support, stride length,10 gait velocity, 10, 12 arm swing, lateral body sway, posture of upper body, and vertical up and down movements of the upper body12 have been reported between patients with clinical depression and healthy elderly controls, with worse performance in the former group. These studies, however, have been mostly limited to older adults in hospital or nursing homes, 10,12 patients with major depression,9⁻¹² measures of mobility other than gait,4,13,22 gait velocity alone,4,13 or younger adults.9-12 Our study extends these findings to a broader assessment of gait performance in nondemented community-dwelling elderly without MDD.

Our study focused on older adults with depressive symptoms but not meeting criteria for MDD. Although MDD is the most frequently studied and well-characterized depressive syndrome,29 the incidence of clinically significant nonmajor forms of depression also steadily increases with age.1 We chose to exclude subjects with MDD, to address the nature of gait dysfunction in the earliest or mildest stages of depression. Moreover, it would be difficult to make inferences about the association between gait and depressive symptoms in the larger non-MDD population if subjects with MDD were included. When we repeated our analyses including subjects with clinical depression in the study sample, the association between depressive symptoms with velocity, stride length, and swing time remained significant. However, as may be expected, the strength of the association was increased. For instance, addition of clinically depressed subjects to the study sample strengthens the association of GDS scores with gait velocity (B -1.31, 95% CI -2.07 to -0.55) compared to the model that excluded depressed subjects (B -1.17, 95% CI -2.05 to -0.30).

There are various possible mechanisms by which depressive symptoms may be related to quantitative gait dysfunction. Vascular mechanisms may be an important link, as it has been reported that there are associations between late-onset depression and gait abnormalities with vascular diseases (such as hypertension, diabetes, or coronary artery disease), frontal–subcortical dysfunction, and white matter lesions.30⁻³⁴ Slower velocity, shorter stride length, and longer double support time have been shown to be associated with subclinical strokes and high white matter ischemic disease on imaging studies.35 Subjects with extensive white matter lesions were three to five times more likely to have depressive symptoms as compared with individuals with mild or no white matter lesions in a population-based cohort study of adults age 60–90.36 Motor (i.e., gait), cognitive, and neuropsychiatric disorders such as depression, may concurrently develop due to disruption of neural circuits originating in the prefrontal cortex37 as the result of neurodegenerative disease, vascular disease, or other diseases.31 Neuroimaging data that could confirm this

vascular hypothesis was not routinely obtained in all subjects in our cohort and needs to be done in follow-up studies.

Gait was a predictor of cognitive decline and incident dementia in our cohort.7 In addition, depression has been demonstrated to be a risk factor for Alzheimer disease.38 To account for cognitive impairment as a possible mechanism linking depressive symptoms and gait dysfunction we adjusted for the Blessed test,25 which correlates well with the pathology of Alzheimer disease.38 However, the significant association remained suggesting that cognitive dysfunction was not the only possible link between gait and depressive symptoms. Depressive symptoms are common in older adults with Mild Cognitive Impairment (MCI) syndrome.39 We have reported that gait dysfunction is common in MCI patients in our cohort.15 Despite excluding subjects with dementia and adjusting for the cognitive status using the Blessed test, the association of gait and depressive symptoms could in part be due to presence of subjects with MCI in this sample,40 and this possibility needs to be explored in future studies.41

Clinical gait abnormalities were included in our fully adjusted models to partially control for nonneurologic limitations of gait such as pain and arthritis as well as neurologic diseases. However, the significant relationships for the three gait variables remained in these additional analyses. We have reported a very low prevalence of neurologic gait subtypes such as parkinsonian, frontal, or hemiparetic that are also associated with mood disorders in the same cohort.6·42 Arthritis has been reported to be associated with depressive symptoms in adults43 and may be another link between depressive symptoms and gait.

The strengths of this study include the large sample size with systematic gait assessments7 and validated clinical and diagnostic procedures for motor and depressive symptoms and signs.7,19 Limitations of the study include the cross-sectional design, which limits inferences about temporality or causality. Although the presumption is that depressive symptoms lead to gait dysfunction, we cannot discount the possibility that gait dysfunction may limit mobility and social interactions increasing risk for developing depressive symptoms in older adults. In support of the reverse hypothesis, a recent study reported that lower walking capacity predicted increased risk of depressive symptoms in older men during an 8-year follow-up period.44 Our findings should be followed up in longitudinal observational studies or clinical trials. Our study excluded subjects with dementia or inability to ambulate at baseline. Therefore, our findings may not generalize to nursing home-based samples, though the observed associations might be stronger in less healthier populations. We limited our examination of gait to the eight variables that have been extensively studied in this cohort.7,15 It is likely that other gait variables may also show independent associations with depressive symptoms. However, most of these other gait variables are likely to be highly correlated with our selected gait variables. Although we examined some possible mechanisms such as vascular disease in our analyses, future studies should examine these observed relationships in more depth using more direct measures of pathology.

In conclusion, increasing depressive symptoms in community residing older adults are associated with quantitative gait dysfunction even in the absence of major depression or dementia. In the primary care setting, elderly patients with early signs of depressive symptoms should be evaluated for therapy or medical treatment and assessed for gait impairment. On the contrary, patients with gait abnormalities should be screened for depression and possible underlying pathologies such as vascular risk factors, which can be targeted for treatment.

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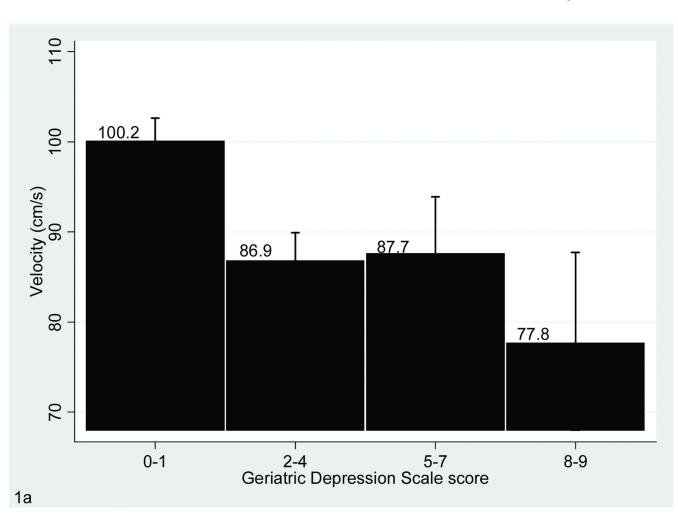
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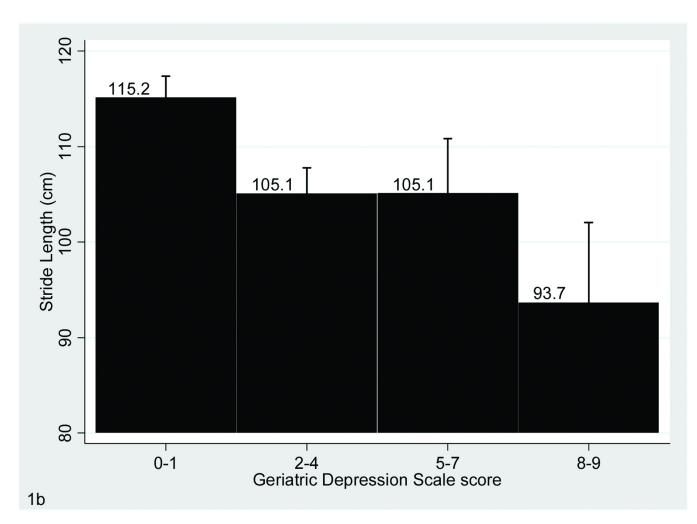
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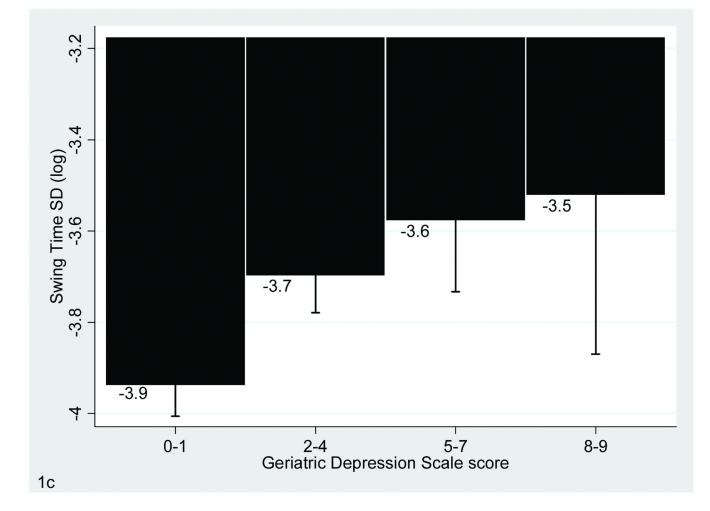


FIGURE 1. Association of Gait With Depressive Symptoms

The bar graphs display the relationship between Geriatric Depression Scale scores and performance on gait velocity (A), stride length (A) and swing time variability (B), which are the three gait parameters that were significantly associated with depressive symptoms in the fully adjusted model. Bars show mean values for the gait variables and error bars indicate standard error of the mean.

Table 1

Association between depressive symptoms measured by the GDS and individual gait variables

	Line	Linear Regression Model 1 \ddagger	n Model 1	**	Line	Linear Regression Model 2 $^{\$}$	n Model 2	s
Outcome Variable	GDS Coefficient Lower CI	Lower CI	Upper CI	P value	GDS Coefficient Lower CI	Lower CI	Upper CI	P value
Velocity, cm/s	-2.98	(-3.85 to -2.10)	-2.10)	$<\!0.001^{\dagger}$	-1.17	(-2.05 to	-0.30)	0.01°
Stride Length, cm	-2.39	(-3.14 to	-1.64)	${<}0.001\dot{\tau}$	-0.95	(-1.69 to	-0.21)	$0.01\dot{\tau}$
Cadence, steps/min	-1.17	(-1.66 to	-0.68)	$< 0.001 \mathring{\tau}$	-0.45	(-0.99 to	(60.0	0.1
Stance Time, seconds	0.01	(0.01 to	0.02)	<0.001	0.004	(-0.003 to	0.01)	0.29
Swing Time, seconds	0.003	(0.001 to	0.005)	0.01^{\dagger}	0.002	(-0.001 to	0.004)	0.17
Double Support Time, seconds	0.01	(0.01 to	0.02)	$<\!0.001^{\ddagger}$	0.003	(-0.004 to	0.01)	0.36
Stride Length Variability, log SD	0.02	(-0.01 to	0.04)	0.15	0.01	(-0.02 to	0.03)	0.51
Swing Time Variability, log SD	0.06	(0.04 to	(60.0	$< 0.001^{+1}$	0.03	(0.002 to	0.06)	0.03

Indicates significant P-value (<0.05) after applying correction for multiple comparisons (see Methods for desc

⁴Linear regression model adjusted for baseline gds, age, sex, race, and education; p-value for variables are derived from t-statistic and has 603 residual degrees of freedom

 $\frac{8}{5}$ Linear regression model additionally adjusted for medical illness index, presence of clinical gait abnormalities, Blessed-Information-Memory-Concentration test score, mean arterial pressure, body mass index, physical activity scale score, and psychotropic use; 489 residual degrees of freedom

Table 2

Baseline Characteristics

	Mean (N=610)	SD
Characteristic		
Age	80.3	5.5
Education, years	14	3.4
	Ν	%
Female	374	61.3%
Race		
White	406	66.6%
Black	171	28.0%
Other	33	5.4%
	Mean	SD
Geriatric Depression Scale score (range 0–15)	2.0	1.9
Blessed-Information-Memory-Concentration test score (range 0–32)	2.0	1.9
Height, cm (n=550)	164.7	10.3
Weight, kg (n=547)	74.3	16.6
Body Mass Index, kg/m2 (n=547)	27.3	5.4
Mean arterial pressure, mmHg (n=536)	97.6	10.2
Illness index (range 0–9)	1.2	1.0
Physical Activity Scale (range 0–20)	1.6	2.2
	N	%
Clinical gait abnormality (n=563)	223	39.6%
Psychotropic use	22	3.6%
	Mean	SD
Gait parameters		
Velocity, cm/s	93.1	23.4
Cadence, steps/min	101.3	12.3
Stride Length, cm	109.7	20.9
Stance Time, seconds	0.8	0.2
Double Support Time, seconds	0.3	0.1
Swing Time, seconds	0.4	0.1
Swing Time Variability, SD	0.0	0.0
Stride Length Variability, SD	4.4	2.5