

Original Contribution

Prediction of Severe, Persistent Activity-of-Daily-Living Disability in Older Adults

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In a prospective cohort of nondisabled adults aged 65 years or more in the Established Populations for Epidemiologic Studies of the Elderly (1981–1987 and 1985–1992), we used a competing risk approach to predict the 5-year risk of severe, persistent activities-of-daily-living (ADLs) disability, defined as dependence in ≥3 ADLs for 2 consecutive annual interviews or for 1 interview followed by death in the subsequent year. During 5 years, 6.8% developed severe, persistent ADL dependence, and 14.6% died without severe, persistent ADL dependence in the derivation cohort ($n = 8,301$); the corresponding percentages were 6.8% and 15.8% in the validation cohort ($n = 4,177$). A model based on age, current employment, visual impairment, self-rated health, diabetes mellitus, history of stroke or brain hemorrhage, cognitive function, and self-reported physical function showed good calibration. Discrimination, assessed by C statistics, for <70, 70–74, 75–79, and \geq 80 years, was 0.75, 0.74, 0.65, and 0.66 in the derivation cohort and 0.70, 0.72, 0.70, and 0.65 in the validation cohort, respectively. In conclusion, a simple risk score based on routinely available clinical information can predict severe, persistent disability in 5 years. Future studies should examine whether physical performance measures can further improve prediction in the oldest old.

activities of daily living; aged; prognosis

Abbreviations: ADL, activity of daily living; EPESE, Established Populations for Epidemiologic Studies of the Elderly.

Despite primary prevention efforts and therapeutic advancements, recent data in the United States suggest an expansion of life with disability, rather than compression of morbidity [\(1](#page-7-0)). Over 20% of older Americans reported at least 1 activityof-daily-living (ADL) disability, and long-term-care expenditure for disabled older adults reached \$135 billion in 2004 ([2](#page-7-0)). With the growing interest in quality of life and health-care cost reduction, ADL dependence is a meaningful, patient-oriented outcome because individual disease–oriented outcomes often fail to capture the overall impact of multiple comorbidities on a person's function. It also predicts institutionalization [\(3](#page-7-0)), home service use [\(4\)](#page-7-0), hospitalization ([5\)](#page-7-0), and mortality ([6](#page-7-0)).

Extensive research has identified a wide range of risk factors for ADL disability, including low socioeconomic status, sensory impairment, poor self-reported health, comorbidities, cognitive and physical functional impairment, and subclinical disease. However, the translation of research into clinical prediction has been slow. Although several prognostic tools are available $(7-12)$ $(7-12)$ $(7-12)$, they are often resource intensive or require trained assessors $(7-11)$ $(7-11)$ $(7-11)$ $(7-11)$. Some focused on mortality (12) (12) . Moreover, previous research did not consider the dynamic nature of disability or competing events that alter the risk of disability. In the era of health-care reform, identifying individuals who are more likely to lose their ability to live independently and utilize more resources is crucial to achieve more appropriate care, cost control, and meaningful use of resources.

This study aimed to develop and validate a practical tool that predicts 5-year risk of severe, persistent loss of independence in performing ADLs in a large representative cohort of community-dwelling older adults, using information that can be easily obtained in a general practice setting.

MATERIALS AND METHODS

Study population

This study used data from the Established Populations for Epidemiologic Studies of the Elderly (EPESE), a prospective cohort study of noninstitutionalized adults, aged 65 years or more, in 4 US communities [\(13](#page-7-0)). The details of study design and conduct were described elsewhere ([14](#page-8-0)–[16\)](#page-8-0). Participants underwent baseline interviews in 1981–1982, except in the North Carolina cohort (in 1985–1986), and annual followup interviews for 6 years. Because additional interviews beyond 6 years were conducted in selected cohorts with varying intervals, we used data up to the sixth annual follow-up. After excluding 248 participants with missing data on more than 50% of potential predictors and 1,665 participants with ADL dependence at baseline, 12,478 participants were included in this analysis. The institutional review boards at Beth Israel Deaconess Medical Center and Hebrew SeniorLife exempted this study.

Definition of ADL dependence and other outcomes

Disability in 7 ADLs was self-reported at baseline and subsequent annual interviews, using the following question: "Other than when you have been in a hospital, was there any time in the past 12 months when you needed help from some person or equipment to do . . . (each of the following activities)?" for walking across a small room, bathing or shower, personal grooming, dressing, eating, getting from a bed to a chair, or using the toilet. Whether the help was from another person, equipment, or both was also asked: We considered requiring personal assistance to define ADL dependence. The event of interest was severe, persistent ADL dependence, defined as requiring personal help in \geq 3 ADLs for at least 2 consecutive annual interviews or for 1 interview followed by death in the subsequent year to capture prolonged, severe disability status that is comparable to requiring a significant amount of personal assistance or nursing home care. Those who died during the follow-up without severe, persistent ADL dependence were considered as having experienced the competing event (refer to "Statistical analysis" below). As our outcome was defined on the basis of ADL status in 2 consecutive annual interviews, the fifth year was the last possible time in which the outcome could occur. Our definition of disability reduces misclassification from self-report and is more strongly associated with adverse outcomes and resource utilization than other definitions (e.g., self-reported difficulty, impairment in any ADLs, or disability at a single assessment) [\(17](#page-8-0)–[21\)](#page-8-0).

Vital status was obtained through obituaries, hospitalization records, proxy interviews, and death certificates. Hospitalizations during the past 12 months were self-reported (yes/no) for selected conditions (myocardial infarction, stroke, cancer, and fracture). Assuming that each condition was associated with a distinct hospitalization, we summed the number of hospitalizations for 5 years.

Potential predictors

We chose 30 potential predictors of ADL disability that were identified from literature and collected via questionnaire and in-person assessment in the EPESE (Web Table 1 available at <http://aje.oxfordjournals.org/>). Multiple response categories for certain predictors were collapsed to ensure a sufficient number of events in each category, to improve interpretability, and to reduce model complexity without compromising model fit. Cognitive function was measured by using the Short Portable Mental Status Questionnaire [\(22](#page-8-0)), a validated instrument that comprises 10 questions on memory, orientation, current events, and a mathematical task. Cognitive function was classified as "normal" if the number of errors was \leq 2; "mild" if 3 or 4; and "moderate to severe" if \geq 5 [\(22](#page-8-0)). A cutpoint of 3 errors corresponds to the Mini-Mental Status Examination score of 23 ([23](#page-8-0)). Because physical performance was measured at the sixth annual examination in the EPESE cohort, we used self-reported physical function obtained from questionnaires. These measures have been shown to be reproducible and validated against performancebased measures ([24,](#page-8-0) [25\)](#page-8-0).

Statistical analysis

All analyses were performed in Stata/SE 11.2 software (StataCorp LP, College Station, Texas). Two-sided P < 0.05 was considered statistically significant. The data set was randomly divided into a derivation cohort $(2/3; n = 8,301)$ and a validation cohort $(1/3; n = 4, 177)$. Characteristics were compared between the 2 cohorts by using the *t* test or Wilcoxon rank-sum test for continuous variables and the Pearson χ^2 test for categorical variables.

Missing data imputation. For those who had any interval missing on the number of ADL dependencies (7.4%), we imputed the missing values with the last nonmissing values for that individual carried forward. At baseline, 2.7% had missing data on more than 5 predictors. We implemented multivariable imputation that used all baseline variables, the time-to-event variable, and outcome status.

Model development. We used cause-specific proportional hazards regression to model the time to first occurrence of severe, persistent ADL dependence as a function of predictors, while accounting for death without severe, persistent ADL dependence as a competing event that prevents the occurrence of our outcome of interest [\(26](#page-8-0)–[28](#page-8-0)). Because the exact event dates were not available, we assumed that they occurred at the midpoint between the 2 interviews.

In selecting predictors, we initially fitted a model with all 30 predictors in the derivation cohort, using severe, persistent ADL dependence as the outcome, and carried out backward elimination to find a model with the minimal Schwarz Bayesian Information Criterion [\(29](#page-8-0)). Because the proportional hazards assumption was violated for age categories, we fitted age-stratified Cox models. Categorical variables were modeled ordinally, if monotonic relationships existed and likelihood ratio tests favored a more parsimonious model over the model with indicator terms. Any significant interaction terms among age, sex, and other predictors that resulted in a lower Bayesian Information Criterion were retained.

Model performance. In the derivation and validation cohorts, calibration was assessed by calibration plots and the goodness-of-fit χ^2 statistic ([30\)](#page-8-0). Discrimination was evaluated by C statistics that were modified to the context of competing risk such that individuals who failed from competing events remained in the risk set at all times [\(28](#page-8-0)).

Calculation of risk score. To facilitate clinical application, we developed a scoring system by assigning a score to

Table 1. Selected Characteristics in the Derivation and Validation Cohorts, the Established Populations for Epidemiologic Studies of the Elderly, United States, 1981–1987 and 1985–1992

Abbreviation: SD, standard deviation.

 a One pound = 0.45 kg.

b Cognitive function was classified as "normal" if the number of errors on the Short Portable Mental Status Questionnaire was \leq 2; "mild" if 3 or 4; and "moderate to severe" if \geq 5 (of 9) [\(22\)](#page-8-0).
^c Half a mile or about 8 ordinary blocks.

 d Body mass index: weight (kg)/height (m)².

each predictor in proportion to its regression coefficient in our final model. We assessed a potential loss in discrimination by comparing the C statistic from the risk score with that from the model. According to the distribution of the risk score in the derivation cohort, participants were classified into tertiles of risk score: low (from 0 to 7); moderate (from 8 to 15); and high (≥16). For each category, we calculated the observed absolute risk of severe, persistent ADL dependence (event of interest) and death without severe, persistent ADL dependence (competing event), as well as the risk of being free of both events in the derivation and validation cohorts. In secondary analyses, we estimated the 5-year risk of any or recurrent hospitalizations for selected conditions (myocardial infarction, stroke, cancer, and fracture), using Poisson regression with log follow-up time as the offset variable.

RESULTS

At the end of 5 years, 6.8% developed severe, persistent ADL dependence, and 14.6% died without severe, persistent ADL dependence in the derivation cohort ($n = 8,301$). The corresponding percentages were 6.8% and 15.8% in the validation cohort ($n = 4,177$). Loss to follow-up was 2.9% and 2.5%, respectively. Distributions of demographic characteristics and other predictors were similar in both cohorts (Table [1\)](#page-2-0).

Model development

In the derivation cohort, most predictors were statistically significantly associated with severe, persistent ADL dependence, as expected from our large sample size. However, the model that included 8 predictors achieved the smallest Bayesian Information Criterion (Table 2). Because the model was stratified by age because of violation of the proportional hazards assumption, the association with age was not directly estimable from the model. We found that self-rated health was a stronger predictor of severe, persistent ADL dependence in younger adults, but less so in older adults ($P_{\text{interaction}}$ < 0.001).

Table 2. Cox Proportional Hazards Model for Severe, Persistent Activity-of-Daily-Living Dependence in the Derivation Cohort, the Established Populations for Epidemiologic Studies of the Elderly, United States, 1981–1987 and 1985–1992

Predictors	Response Categories	HR ^a	95% CI
Currently working at a paying job	No vs. yes	2.08	1.33, 3.23
Able to read ordinary newspaper print	No vs. yes	1.44	1.17, 1.77
Self-rated health	Per each category increase (excellent/ good, fair, or poor)	1.22	1.07, 1.39
Diabetes mellitus	Yes vs. no	1.38	1.12, 1.71
Ever had stroke or brain hemorrhage	Yes vs. no	1.74	1.32, 2.29
Cognitive function ^b	Per each category increase (normal, mild, or moderate/ severe)	1.85	1.64, 2.08
Able to walk half a mile ^c	No vs. yes	1.61	1.33, 1.95
Able to do heavy housework	No vs. yes	1.51	1.24, 1.83

Abbreviations: CI, confidence interval; HR, hazard ratio.

a Age-stratified Cox model that included all predictors; indicators for study sites were fitted for severe, persistent activity-of-daily-living dependence. Interaction terms were not included in this model to better demonstrate the associations with the main-effect terms. Refer to Web Table 2 available at <http://aje.oxfordjournals.org/> for the final model with interaction terms.

^b Cognitive function was classified as "normal" if the number of errors on the Short Portable Mental Status Questionnaire was ≤2; "mild" if 3 or 4; and "moderate to severe" if \geq 5 (of 9) ([22\)](#page-8-0).
^c Half a mile or about 8 ordinary blocks.

Because this interaction term improved the overall model fit by reducing the Bayesian Information Criterion, it was included in our final model (Web Table 2). There was no statistically significant interaction between sex and other predictors ($P_{\text{interaction}} > 0.05$).

Model performance

To estimate the absolute risk of severe, persistent ADL dependence, we fitted separate cause-specific Cox models (Web Table 2) and computed cumulative incidence function. There was a good agreement between the predicted risk estimated from the cause-specific hazards model and the observed risk (Figure 1 and Web Figure 1); the goodness-offit χ^2 statistic = 5.50 (P = 0.79) in the derivation cohort, and the goodness-of-fit χ^2 statistic = 3.56 (P = 0.94) in the validation cohort. However, it was evident that a standard Cox model that did not consider competing risk overestimated the risk (Figure 1).

Discrimination was good in younger age and modest in older age: C statistics were 0.65–0.75 in the derivation cohort and 0.65–0.72 in the validation cohort (Table [3\)](#page-4-0). By subgroups, the C statistic was particularly lower in those who were ≥ 80 years.

ADL dependence, death, and hospitalization according to risk score

We generated an algorithm that assigned scores in proportion to regression coefficients, without a substantial loss in discrimination compared with the original model $(C$ statistics were 0.64–0.76 in the derivation cohort and 0.66–0.71

Figure 1. Calibration plot of a cause-specific hazards model and a standard Cox regression model for the prediction of severe, persistent activity-of-daily-living dependence in the validation cohort, the Established Populations for Epidemiologic Studies of the Elderly, United States, 1981–1987 and 1985–1992. The risk was estimated as the 5-year cumulative incidence of severe, persistent activity-ofdaily-living dependence from the cause-specific hazards model that accounted for competing risk.

Table 3. Discrimination in the Derivation and Validation Cohorts, the Established Populations for Epidemiologic Studies of the Elderly, United States, 1981–1987 and 1985–1992

Abbreviation: CI, confidence interval.

a Cardiovascular disease was defined as self-reported history of myocardial infarction, stroke, or diabetes mellitus.

in the validation cohort). The age-specific risk tertiles and corresponding risk of severe, persistent ADL dependence (Figure [2\)](#page-5-0) are useful in assessing an individual's absolute risk as well as relative risk compared with others of similar age. For example, a man aged 73 years whose risk score is over 16 has 10.6% risk of developing severe, persistent ADL dependence in 5 years, which is approximately 3-fold higher than the risk of an average person of his age and is as high as that of an average person who is 10 years older.

In order to provide comprehensive prognostic information, we summarized the 5-year risks of severe, persistent ADL dependence, death without severe persistent ADL dependence, and being alive without both events (Table [4\)](#page-6-0). Those in the upper tertile had higher risks of severe, persistent ADL dependence, as well as death without severe, persistent ADL dependence. The risk tertiles also predicted hospitalizations for myocardial infarction, stroke, cancer, and fracture within 5 years in a dose-dependent manner, particularly among younger subgroups (Web Figure 2).

DISCUSSION

In a representative population of nondisabled, communitydwelling, older adults, we developed and validated a simple risk assessment model that predicted the 5-year risk of severe, persistent ADL dependence using age, 7 self-reported risk factors, and cognitive function that can be easily obtained in a

general practice setting. This model provides the absolute risks of severe, persistent ADL dependence and death without ADL dependence that can be useful in personalized prognostication and care planning. Although we recognize that the age of the data set (ADL data collected 20–25 years ago) is a major limitation for clinical use of our model for contemporary older adults, several features of our approach deserve mention and enhance future research on prediction of disability in older adults.

Comparison with existing models that predict disability

Several frailty indices and validated risk assessment models predict ADL disability in community-dwelling older adults (Web Table 3). The amount and source of information needed for risk calculation vary across the models. The frailty index [\(31](#page-8-0)–[33](#page-8-0)), Sarkisian et al. [\(34](#page-8-0)), and the Vulnerable Elder Survey-13 ([35](#page-8-0), [36](#page-8-0)) screened a wide range of potential predictors, whereas the Cardiovascular Health Study index [\(8](#page-7-0)), the Study of Osteoporotic Fracture index ([37](#page-8-0), [38\)](#page-8-0), and the Short Physical Performance Battery [\(9](#page-7-0), [39\)](#page-8-0) mainly focused on physical performance with or without other predictors. Although variations in outcome definitions and health status of study populations do not allow a direct comparison among these models, C statistics ranged from 0.64 to 0.76.

Compared with existing models, our model has several strengths. We estimated the absolute risk of severe, persistent ADL dependence that is meaningful to clinicians, patients, and policy makers, in the presence of competing event. Although disability is a dynamic status, the severity and duration of ADL disability have rarely been considered in previous research. In a cohort of community-dwelling older adults, approximately 80% of the newly disabled recovered independence within 12 months, and the recovery rate was lower as the severity was higher [\(40\)](#page-8-0). Self-reported dependence was more closely linked to hospitalizations and resource use than self-reported difficulty ([19](#page-8-0)). Dependence in \geq 3 ADLs was associated with an over 3-fold increase in institutionalization within 2–3 years ([18\)](#page-8-0). Thus, our model is likely to capture the most impaired subgroup of this older population with a greater likelihood of adverse health events that can affect quality of life and drive health-care expenditure.

In building our model, we considered multiple biological and psychosocial risk factors of ADL disability to represent heterogeneous disabling processes in community-dwelling older adults. Physical performance (e.g., gait speed) predicts ADL disability, but it was not measured at the EPESE baseline. Biomarkers and imaging studies may provide additional prognostic information with incremental costs. Our objective was to develop a model based on information that can be easily obtained without additional resources and costs in a busy practice setting (e.g., interview and routine physical examination in a clinic room) for a broader application. Our model showed C statistics across subgroups that were similar to other resourceintensive models (Table 3 and Web Table 3).

Potential implications for clinical care

Physicians often feel unsure about when and how to initiate a discussion about advance care planning, particularly

Figure 2. Score sheet to estimate 5-year risk of severe, persistent activity-of-daily-living dependence, the Established Populations for Epidemiologic Studies of the Elderly, United States, 1981–1987 and 1985–1992. The risk was estimated as the 5-year cumulative incidence of severe, persistent activity-of-daily-living dependence from the cause-specific hazards model that accounted for competing risk. Cognitive function was considered "normal" if the number of errors on the Short Portable Mental Status Questionnaire was ≤2; "mildly impaired" if 3 or 4; and "moderately to severely impaired" if ≥5 ([22\)](#page-8-0).

for older adults with multicomorbidities who do not have a terminal diagnosis nor follow a predictable functional decline [\(41](#page-8-0), [42](#page-8-0)). Routine use of a prognostic tool during a primary care visit can offer an opportunity to discuss prognosis and advance care planning, regardless of the levels of predicted risk ([42\)](#page-8-0). Unlike existing prognostic indices based on mortality ([12\)](#page-7-0), our model provides more comprehensive prognostic information on the risks of both severe disability and mortality within 5 years. At the individual level, clinicians can use this information to prioritize competing health issues and reset the goals of preventive care and chronic disease management. For high-risk adults, the focus should be minimizing avoidable harms and maintaining quality of life and function; for low-risk to moderate-risk adults, the

focus should be increasing life expectancy and preventing disability. Importantly, almost one third of adults aged 80 years or more are in the low-risk group. They should not be excluded from preventive care and aggressive chronic disease management, based on age alone. At the health-care system level, practice-based interventions that target these individuals may reduce hospitalizations and health-care costs.

Limitations

Our study did not examine whether using a risk model leads to better clinical outcomes. Because our goal was to predict severe disability that was comparable to nursing

Table 4. Risks of Severe, Persistent Activity-of-Daily-Living Dependence and Death by Age in the Derivation and Validation Cohorts, the Established Populations for Epidemiologic Studies of the Elderly, United States, 1981-1987 and 1985-1992^a

Abbreviation: ADL, activity of daily living.

^a The risk was estimated as the 5-year cumulative incidence of severe, persistent activity-of-daily-living dependence and of death without severe, persistent activity-of-daily-living dependence from the cause-specific hazards model that accounted for competing risk.

home care, we did not examine individual patterns of ADL disability or study causal effects of risk factors. Thus, the effect estimates from our model (Table [2\)](#page-3-0) cannot be interpreted as the expected reduction in disability risk with modification of predictors. Moreover, the discriminatory ability of our model was lower in older age. In older age, C statistics from the full model with all 30 predictors were almost the same as those from our final model, suggesting that additional self-reported information is unlikely to improve discrimination. Further improvement by physical performance or biomarkers remains possible. Future studies should examine an incremental value of physical performance tests or biomarkers, using our model as a referent model.

There is mixed evidence on how the disability patterns have changed in older Americans during the past 2 decades [\(43](#page-8-0), [44\)](#page-8-0). The relations of socioeconomical (e.g., currently working at a paying job) or biological risk factors (e.g., diabetes mellitus or stroke) with disability might have changed over time. Therefore, recalibration of absolute risk, as well as validation in a contemporary cohort, is necessary before widespread application of our model. In addition, hospitalizations were self-reported for selected conditions that accounted for 16% of total hospitalizations among older Americans in 2009 ([45\)](#page-8-0). Finally, the last-observation-carried-forward method for interval missing data on ADL dependence might have underestimated the incidence of ADL dependence and discrimination of our risk model, if those who did not respond to the interview had worsening disability.

Conclusions

A practical risk assessment tool that uses routinely available clinical information can predict severe, persistent disability that has important implications for quality of life, resource utilization, and health-care expenditure. Future research should validate our model in an independent cohort, examine the additional improvement in prediction by measurements of physical performance and biomarkers, and evaluate clinical outcomes of a comprehensive program that incorporates our risk model.

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