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Development and validation of a brief dementia screening indicator for primary care

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Abstract

Background—Detection of "any cognitive impairment" is mandated as part of the Medicare annual wellness visit, but screening all patients may result in excessive false positives.

Methods—We developed and validated a brief Dementia Screening Indicator using data from four large, ongoing cohort studies (the Cardiovascular Health Study [CHS]; the Framingham Heart Study [FHS]; the Health and Retirement Study [HRS]; the Sacramento Area Latino Study on Aging [SALSA]) to help clinicians identify a subgroup of high-risk patients to target for cognitive screening.

Results—The final Dementia Screening Indicator included age (1 point/year; ages, 65–79 years), less than 12 years of education (9 points), stroke (6 points), diabetes mellitus (3 points), body mass index less than 18.5 kg/m² (8 points), requiring assistance with money or medications (10 points), and depressive symptoms (6 points). Accuracy was good across the cohorts (Harrell's C statistic: CHS, 0.68; FHS, 0.77; HRS, 0.76; SALSA, 0.78).

Conclusions—The Dementia Screening Indicator is a simple tool that may be useful in primary care settings to identify high-risk patients to target for cognitive screening.

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Kevwords

Dementia; Screening; Risk prediction modeling; Primary care

1. Introduction

There are currently 5.2 million individuals with Alzheimer's disease and other dementias in the United States, including one in eight of those age 65 years or older [1]; however, 50% to 75% of these cases are undiagnosed [2,3,4]. Current U.S. Preventive Service Task Force guidelines, which were published in 2003, recommend screening only when cognitive impairment or deterioration is suspected [5,6]. However, many patients with dementia appear normal during clinical encounters and do not exhibit traditional signs or symptoms until they are screened. The 2010 Patient Protection and Affordable Care Act now requires providers to "detect any cognitive impairment" as part of the newly established annual wellness visit (AWV) for Medicare recipients, but no specific guidelines for achieving this new mandate were provided [7].

Primary care clinicians face numerous barriers to dementia diagnosis, including patient visit time constraints; complexities of diagnosis, reimbursement and follow-up; lack of an effective treatment; and concern about causing distress in patients and their families [4,8]. However, research published during the past decade suggests that early diagnosis of dementia has many potential benefits for clinicians, patients, and family members. These benefits include earlier initiation of dementia treatment, which may delay symptom progression in some patients [9]; better management of care for other medical issues [10], which could potentially reduce avoidable hospitalizations [11]; greater patient involvement with planning for their future care and modification of the home environment for their protection; and caregiver participation in support programs, which have been found to reduce burden and depression [12,13].

To provide guidance regarding detection of cognitive impairment during the AWV, an Alzheimer's Association work group recently recommended that a brief cognitive screening be performed on all Medicare patients who show either signs or symptoms of cognitive impairment or do not have an informant available to confirm a lack of cognitive impairment [14]. A potential consequence of this approach, however, is that many low-risk patients—especially younger Medicare patients—may be screened simply because they do not have an informant with them at the time of the AWV. This could take time away from other important clinical issues and could potentially result in large numbers of false-positive findings, unnecessary referrals for expensive follow-up testing, and needless worry to patients and families.

The objective of the current study was to develop and validate a brief Dementia Screening Indicator for use in primary care settings to help identify a subgroup of patients that does not have overt signs or symptoms of cognitive impairment but, based on other risk factors, has an increased risk of dementia and could be targeted for cognitive screening. Thus, our goal was to develop an easy-to-use clinical tool that uses information already available or easily

obtainable in the primary care setting to stratify older patients into low- and high-risk groups, with the goal of targeting cognitive screening to the latter group.

2. Methods

2.1. Brief overview and study cohorts

This research was performed by four teams of investigators, who coordinated their efforts to develop and validate a common Dementia Screening Indicator. Each team was associated with a well-established, community- or population-based cohort study. The four studies consisted of the Cardiovascular Health Study (CHS; team, KY, DB, and AK), the Framingham Heart Study (FHS; team, SS, AB, and SP], the Health and Retirement Study (HRS; team: DW, KL, and RM], and the Sacramento Area Latino Study on Aging (SALSA; team: MH, AL, and JN). These studies were selected because they included individuals from diverse geographic and race/ethnic backgrounds, ascertained subjects for incident dementia, and evaluated a broad palette of potential risk markers.

For each study, a special data set was created for the secondary data analyses performed in this investigation. Data were not pooled across studies; rather, each team conducted its own analyses. Table 1 provides an overview of basic characteristics of the cohorts associated with the four studies. Although the studies have been active for many years, the information about data collection, eligibility, and sample size contained in the table is specific to the current investigation. Detailed descriptions of the design and methods of each study are available in the scientific literature for CHS [15], FHS [16,17], HRS [18], and SALSA [19]. Additional information about the studies is available online at study-specific websites (CHS [20], FHS [21], HRS [22], and SALSA [23]. All studies obtained written, informed consent from study participants. The CHS analyses also were approved by the Committee on Human Research at the University of California, San Francisco, and the Research & Development Committee at the San Francisco Veterans Affairs Medical Center.

2.2. Ascertainment of dementia

Each study used different procedures to ascertain dementia prevalence and incidence, and the details are available elsewhere (CHS [24–26], FHS [27,28], HRS [29,30], SALSA [19]). However, the minimum requirements for dementia diagnosis across the four studies were (*i*) cognitive impairment in at least two domains that reflected a decline from prior levels and (*ii*) sufficient severity to affect daily function. In CHS, FHS, and SALSA, diagnoses were based on consensus committee review. In HRS, diagnoses were based on validated cut points on a brief cognitive battery [29,30].

2.3. Predictive factors

We first developed a list of potential predictive factors (or predictors) that were available in all or most of the cohorts and that would be readily available or easily assessed in most primary care settings. Demographic factors included age, sex, and years of education. Medical history included stroke, diabetes, hypertension, and coronary heart disease. Depressive symptoms were assessed on the basis of individual items from the Center for Epidemiologic Studies Depression Scale [31] and current use of antidepressant medication.

The Center for Epidemiologic Studies Depression Scale items that were administered in all four cohorts were: I felt that everything I did was an effort, My sleep was restless, I could not "get going." Function was determined on the basis of self-report of difficulty with individual activities of daily living, instrumental activities of daily living, and mobility. Items common across the four cohorts included self-reported difficulty with eating, dressing, bathing, using the toilet, managing either money or managing medications, using the telephone, getting out of a bed or a chair, walking, climbing stairs, and performing heavy housework. Body mass index (BMI) was measured on the basis of weight in kilograms divided by height in square meters and was classified as underweight (BMI, <18.5), normal (BMI, 18.5–24.9), overweight (BMI, 25.0–29.9), or obese (BMI, 30 or greater).

2.4. Analyses

Knowing that the risk of dementia in the elderly increases with advancing age, we first used an empirical process to select an age at which the risk of dementia was considered high enough that routine screening for cognitive impairment should be considered on the basis of age alone. To accomplish this goal, we calculated 6-year Kaplan–Meier dementia incidence rates as a function of 5-year baseline age categories for each cohort (Table E1). Although dementia incidence rates varied widely across the studies, we observed a sharp increase in 6-year risk at age 80 years, with an approximate doubling of risk in persons age 80 to 84 years compared with those age 75 to 79 years, and absolute risks among persons 80 years or older that was more than one in four in all studies. Hence, we agreed collectively that risk was adequately high in all the studies that, at an age of 80 years or older, cognitive screening could be considered on the basis of age alone.

We then developed the Dementia Screening Indicator in 65- to 79-year-olds to determine which individuals in this younger age group had a risk of dementia similar to that observed in the older participants. We examined bivariate associations between predictor variables and 6-year incident dementia using χ^2 tests for categorical variables and t tests for continuous variables. Cox proportional hazards regression was then used to develop multivariable models of time to dementia onset, with subjects censored at death or the last evaluation. Each group determined independently which variables were most predictive of dementia incidence within their cohort. We then compared across the four cohorts to identify a subset of variables that were predictive consistently. When studies identified different variables that appeared to reflect similar domains, or asked similar questions in slightly different ways, we used an iterative group consensus process to harmonize variables across the studies. For example, some studies asked about difficulty managing money whereas others asked about difficulty managing medications, and all studies found that this "domain" was highly predictive of dementia risk. Therefore, our final model included the variable "difficulty managing money or medications." Each group then performed a final Cox regression analysis using the harmonized predictors, and a fixed-effects meta-analysis was performed to calculate the pooled hazard ratio estimates for each predictor across the four studies. Fixed-effects and random-effects approaches typically yield similar point estimates and differ mainly in their estimates of precision [32], which were not needed for the current study. A point system was then created by dividing the β coefficients for each variable in the final model by the β coefficient for age and rounding to the nearest integer.

Discrimination of the final Dementia Screening Indicator using full model coefficients and the point system was assessed using Harrell's C statistic. In addition, sensitivity analyses were performed for CHS and HRS to assess discrimination within white, black, and Latino groups. Calibration of the final model was assessed by plotting actual risk as a function of decile of predicted risk for the four studies. Kaplan–Meier survival curves were calculated for each cohort using different risk score cut points to compare dementia incidence in low-and high-risk younger participants (65–79 years) with older participants (80–84 years). Several different cut points were examined, and the final cut point was chosen to provide the closest match between high-risk 65- to 79-year-olds and typical 80- to 84-year-olds across the four cohorts. Sensitivity and specificity values are not included because our tool is designed to identify a high-risk subgroup to consider for cognitive screening, rather than to detect dementia per se.

3. Results

Potential predictors and corresponding baseline cohort characteristics are presented in Table 2. The mean age ranged from 71 to 73 years, and the proportion of women was slightly greater than 50%. Education levels varied widely, with less than 12 years of education reported in 13% of FHS participants vs. 71% of SALSA participants. Participants in SALSA also had greater percentages of obesity, based on BMI, and more comorbid medical conditions, including stroke, hypertension, and diabetes.

Although a slightly different constellation of factors was identified in each study as being predictive of dementia, the general domains identified were similar. The variables that were predictive consistently across the four studies and were retained in the final model included older age, fewer years of education, history of stroke, presence of diabetes mellitus, low BMI, assistance needed with money or medications, and a composite depressive symptom variable (current usage of antidepressant medication or reporting that "everything was an effort" for 3 or more days per week during the past week; Table 3).

Discrimination of the final model based on Harrell's C statistic (95% confidence interval) was good for each study (Table 3): CHS, 0.68 (0.65–0.72); FHS, 0.77 (0.73–0.82); HRS, 0.76 (0.74–0.77); SALSA, 0.78 (0.72–0.83). Sensitivity analyses within CHS and HRS also suggested that discrimination was good within different race/ethnic groups—CHS: whites, 0.70 (0.66–0.73); blacks, 0.65 (0.58–0.72); HRS: whites, 0.75 (0.73–0.77); blacks, 0.70 (0.68–0.72); Latinos, 0.71 (0.67–0.75). Calibration of the final model suggested good correspondence between predicted and actual risk over deciles of predicted risk, particularly for the greatest deciles (Fig. E1).

Table 3 also shows the hazard ratios for each final predictor within the four studies as well as the combined hazard ratio and number of points assigned for each factor. Age was assigned 0 point at age 65 years and 1 point per year for ages 66 to 79 years. Therefore, the point values for the other predictors can be interpreted as the increased risk associated with being older by that number of years. For example, a person with a history of stroke (6 points) would have an average dementia risk equivalent to someone without a history of stroke who was 6 years older. The point values for the other factors were as follows: fewer

than 12 years of education, 9 points; BMI less than 18.5, 8 points; diabetes, 3 points; needing help with money or medications, 10 points; and depressive symptoms, 6 points. Therefore, the Dementia Screening Indicator score may range from 0 points (age 65 years, no other predictors present) to 56 points (age 79 years, all other predictors present).

A cut point of 22 points or more on the Dementia Screening Indicator enabled stratification of 65- to 79-year-olds into high-risk and low-risk groups (Fig 1). Dementia incidence in high-risk 65- to 79-year-olds was comparable with dementia incidence in the general population of 80- to 84-year-olds in all four cohorts. The percentages of 65- to 79-year-olds whose scores were 22 points or more and who would, therefore, be recommended for cognitive screening were 8.2% in CHS, 6.0% in FHS, 11.2% in HRS, and 26.7% in SALSA.

A Web-based version of the Dementia Screening Indicator is available [33].

4. Discussion

We developed and validated a simple Dementia Screening Indicator that was designed specifically for primary care settings to help clinicians identify a subgroup of older patients with an increased risk of dementia who could be targeted for cognitive screening. The Dementia Screening Indicator includes seven simple items—age, educational attainment, history of stroke, presence of diabetes mellitus, difficulty managing money or medications, low BMI, and depressive symptoms—that are combined to create a point score, with greater scores indicating greater risk of developing dementia. Dementia Screening Indicator scores or 22 points or more identified 65- to 79-year-olds whose 6-year dementia risk was comparable with 80- to 84-year-olds in all four cohorts. Furthermore, the percentage of subjects identified as high risk based on this score was relatively low, ranging from 6% in FHS to 27% in SALSA, suggesting that usage of this tool in clinical settings could potentially improve efficiency by targeting cognitive screening to a small, high-risk subgroup.

The accuracy of the Dementia Screening Indicator was consistently good across four large community- or population-based cohort studies with wide-ranging demographic and race/ethnic characteristics, suggesting broad generalizability. C statistics ranged from 0.68 to 0.78, which is consistent with other widely used clinical risk indices such as the Framingham Heart Index [34] and the Breast Cancer Risk Assessment Tool [35].

Several other dementia risk prediction tools have been developed during the past few years [36–44]. However, none are ideal for primary care or the AWV because they consider either a restricted range of risk factors (e.g., midlife only [41], vascular only [42,43], general health questions only [44]) or include measures of cognitive function [37–40], or other tests that are expensive or require special equipment [37]. The goal of the current study was to provide a tool that can be used easily in primary care settings to identify a subset of patients with an increased dementia risk who could then be targeted for screening with a standard cognitive instrument.

Although the Patient Protection and Affordable Care Act now requires detection of "any cognitive impairment" during the AWV [7], screening all older adults for cognitive

impairment is not currently recommended by the U.S. Preventive Services Task Force [6]. This conclusion was based on lack of evidence for clinical benefit as well as concern regarding potential harms associated with false-positive findings that could result in costly follow-up testing and unnecessary patient worry. However, these guidelines were published a decade ago, and there is now a growing body of research suggesting that timely diagnosis of dementia has many potential benefits for clinicians, patients, and family members. Earlier diagnosis enables earlier initiation of treatment, which delays worsening of symptoms in some patients [9]. In addition, knowledge that a patient has dementia may enable clinicians to manage care more effectively for other medical issues by simplifying medication regimens, providing written directions, and increasing involvement of family members and other mechanisms of social support [10]. This, in turn, may lead to the prevention of avoidable hospitalizations [11]. Early detection of cognitive impairment may also benefit patients by enabling them to be involved with decision making related to their future care, by prompting alterations in the home environment to increase safety, and by enabling them to participate in research studies if they wish. Last, early detection of cognitive impairment can benefit caregivers by providing them with an opportunity to participate in support programs, which have been found to reduce caregiver burden and depression [12,13].

On the basis of empirical considerations, we determined that routine cognitive screening should be considered for all older adults beginning at age 80 years. We recognize that not all will agreewith this conclusion. However, inour view, the potential benefits of routine screening likely outweigh the potential harms, given the high incidence of dementia in this age group.

Fig. 2 provides a decisional flow chart to guide clinicians regarding when to screen for cognitive impairment in the primary care setting. Screening is *always* indicated if cognitive impairment is suspected on the basis of clinical observation, patient concern, or family concern. In addition, as noted earlier, we recommend screening for all patients age 80 years or older. In patients age 65 to 79 years, the Dementia Screening Indicator provides a simple tool to help clinicians discriminate between those patients with a high risk vs. a low risk of developing dementia within 6 years, so that screening can be targeted toward those with higher risk. Our findings suggest that patients who receive a score of 22 points or greater on the Dementia Screening Indicator have an elevated risk of developing dementia and should be considered for screening, although individual clinicians may choose to be more or less aggressive.

Our approach builds on recommendations for assessment of cognitive impairment during the AWV developed recently by an Alzheimer's Association work group [14]. These recommendations call for brief cognitive screening in all older patients who (*i*) show signs or symptoms of cognitive impairment on the basis of clinician observation, patient self-report, or informant report; or (*ii*) do not have an informant available to provide information on signs and symptoms. These guidelines also include a list of cognitive screening tools that are well suited for the primary care setting. Patients who screen positive are then referred for a full dementia evaluation. The Dementia Screening Indicator provides a complementary approach in which patients, with or without an informant, would be screened only if they

had signs or symptoms or were identified as high risk based on either age of 80 years or more, or scoring 22 points or more on the Dementia Screening Indicator.

The predictors included in the Dementia Screening Indicator have all been identified previously as being associated with increased dementia risk in prior studies [45–50]. However, these predictors are not necessarily associated causally with increased dementia risk. In fact, several predictors may be markers of preclinical or early-stage disease in some individuals, rather than true risk factors. Requiring assistance with money or medications [51], low BMI [52], and depression [53] all may occur during the preclinical period. In addition, low educational attainment is likely to be a proxy for a wide range of factors that could potentially increase dementia risk by lowering cognitive reserve [54]. Therefore, changes in these predictors will not necessarily lead to changes in risk. Nonetheless, the combination of the seven items included in the Dementia Screening Indicator were predictive across four large, diverse cohorts, suggesting they are consistent markers of increased dementia risk, even if the etiology of the association is not clear.

4.1. Strengths and limitations

A key strength of this study is that the Dementia Screening Indicator was developed and validated using data from four well-characterized, community- or population-based cohort studies that are generally reflective of the U.S. population. Despite a diversity of approaches to measuring the identified risk factors, accuracy remained consistent across the four studies, suggesting the tool would retain its validity regardless of the method used to measure the risk factors. Accuracy was somewhat lower in CHS than the other cohorts, which may reflect the older age of the data (follow-up ended in 1999) or the retrospective nature of the dementia adjudication process [25]. In addition, a greater percentage of participants in SALSAwere classified as high risk. This may reflect the lower educational attainment in this group.

Our study also has limitations. First, we were unable to consider some potentially important predictors because they were not collected routinely in all the studies. One example is physical inactivity, which may contribute to as many as one in five dementia cases in the United States [55]. In addition, our tool was not validated externally, and several race/ethnic groups were not represented in the current study, including Asians, Native Americans, and Pacific Islanders. It would be useful to validate the Dementia Screening Indicator in other study populations in future studies. Additional studies also are needed to assess the feasibility of using the Dementia Screening Indicator in clinical settings and to determine whether it increases the efficiency of dementia detection or improves patient outcomes.

5. Conclusion

The Dementia Screening Indicator is a simple tool designed for use by primary care clinicians to identify high-risk older patients who could be targeted for cognitive screening. Restricting cognitive screening to patients identified as high risk by the Dementia Screening Indicator, as opposed to screening all patients, may enable clinicians to maximize the potential benefits of screening while minimizing potential harms.

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RESEARCH IN CONTEXT

1. Systematic review: We searched PUBMED using *dementia* as a major subject heading term and the words *risk* and *prediction*, *risk index*, *risk score*, and *prognostic index*, with results restricted to English-language studies performed in humans age 65 years or older. A total of 181 references were retrieved, and the titles and abstracts were reviewed. None of the articles identified provided a simple clinical tool that did not include cognitive items and could be used in primary care settings to identify older adults with an increased risk of dementia.

- 2. Interpretation: The Dementia Screening Indicator fills an important gap in the field by providing primary care clinicians with a simple tool to help identify a subset of older patients without overt signs or symptoms who have an elevated risk of dementia and could be targeted for cognitive screening. Restricting cognitive screening to high-risk patients, as opposed to screening all patients, may enable clinicians to maximize the potential benefits of screening while minimizing potential harms.
- **3.** Future Directions: Additional studies are needed to examine the feasibility of using the Dementia Screening Indicator in clinical settings and to determine whether it increases the efficiency of dementia detection or improves patient outcomes.

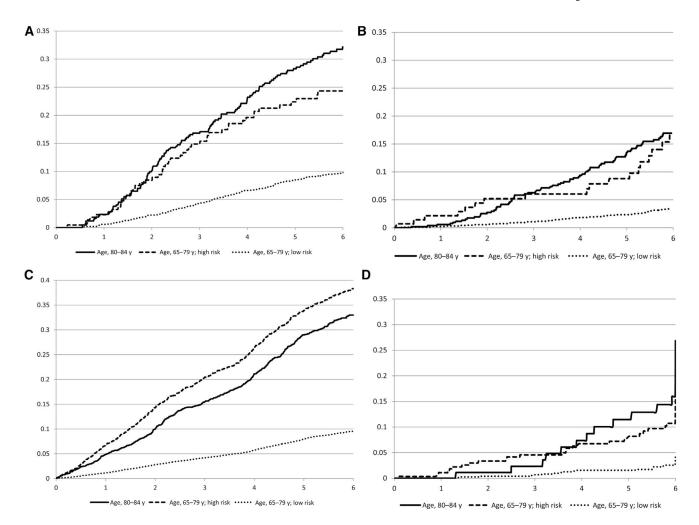


Fig. 1. (A–D) Six-year dementia incidence by age and dementia risk group. The y-axis indicates the proportion of study participants that developed dementia. The x-axis indicates dementia incidence over six years in 65- to 79-year-olds with low dementia risk scores (<22 points, dotted lines), 65- to 79-year-olds with high dementia risk scores (22 points, dashed lines), and normal-risk 80- to 84-year-olds (solid lines) in the Cardiovascular Health Study (A), the Framingham Heart Study (B), the Health and Retirement Study (C), and the Sacramento Area Latino Study on Aging (D). All figures show good separation between the low-risk and high-risk groups, with comparable dementia incidence in high-risk 65- to 79-year-olds and normal-risk 80- to 84-year-olds.

Do you think your patient may have cognitive impairment based on:			
□ your observations □ concerns of patient □ conc	cerns of far	mily or othe	ers
If yes to any: Your patient should be screened for cognitive impairment.			
Is your patient age 80 years or older? □ yes □ no			
If yes: Your patient should be screened for cognitive impairment.			
If no: Administer Dementia Screening Indicator.			
Dementia Screening Indicator			Points
1. How old is your patient? years			
If 65-79 years, assign 1 point per year above age 65. Example: age 65 years	s receives () points;	
age 72 years receives 7 points.			
2. Does your patient have < 12 years of education? ¹	No (0)	Yes (9)	
	10 (0)	168 (9)	
3. Is your patient's BMI < 18.5 kg/m ² ? ²	No (0)	Yes (8)	
	10 (0)	103 (8)	
4. Does your patient have a history of type 2 diabetes?	No (0)	Yes (3)	
	140 (0)	163 (3)	
5. Has your patient ever had a stroke?	No (0)	Yes (6)	
	140 (0)	103 (0)	
6. Does your patient need help from others to manage money or	No (0)	Yes (10)	
medications? ³	140 (0)	103 (10)	
7. Does your patient currently take anti-depressant medications OR report	N- (0)	V (6)	
that "everything was an effort" ≥3 days per week over the past week? ⁴	No (0)	Yes (6)	
Total point score: If \geq 22, your patient should be screened for cognitive impositive	airment.	<u> </u>	

	rarely (<1 day)	□ little (1-2 days)	□ moderate (3-4 days)	□ most (5-7 days)
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If the answer is either "moderate" or "most," then item 7 in the Dementia Screening Indicator receives a score of 6 points.

Fig. 2. Decisional flow chart regarding screening for cognitive impairment in the primary care setting.

¹Did not graduate from high school or pass a General Educational Development (GED) Test

²http://nhlbisupport.com/bmi/

³Ask your patient and/or family member if present: *Do you need help from others to manage money or medications?*

⁴Ask your patient: In the past week, how many days did you feel that everything was an effort?

Table 1

Characteristics of the four cohorts used to develop and validate the dementia screening indicator*

Characteristics	Cardiovascular Health Study	Framingham Heart Study	Health and Retirement Study	Sacramento Area Latino Study on Aging
Geographic location	Four local communities (Forsyth County, NC; Washington County, MD; Sacramento County, CA; Pittsburgh, PA)	Local community (Framingham, MA)	Nationally representative probability sample	Local community (6 selected census tracts, Sacramento Valley, CA)
Race/ethnic diversity	85% white, 15% black	100% white	88% white, 7% black, 5% Latino	100% Latino
Data collection period	1989–1999	1990–2010	1998–2010	1998–2008
Data collection cycle, mo	12	12	24	12–15
Data collection mode	Clinic visit, telephone	Study site, home visit	Home visit, telephone	Home visit, telephone
Eligibility at baseline				
Age, y	65+	65+	65+	65+
Dementia free	Yes	Yes	Yes	Yes
Community dwelling	Yes	Yes	Yes	Yes
Sample size at baseline	2794	2411	13,889	1125
Dementia diagnosis requirements				
At least two cognitive domains impaired	Yes	Yes	Yes	Yes
Documented functional disability	Yes	Yes	Yes	Yes

^{*} Information pertains to current investigation. Age range restricted to 65 to 79 years at baseline.

Table 2

Potential dementia predictors, by study cohort

Potential predictor	CHS $(n = 2794)$	FHS $(n = 2411)$	HRS (n = 13,889)	SALSA (n = 1125)
Demographics		_		
Age, y, mean (SD)*	72.9 (3.2)	72.1 (4.4)	71.3 (4.2)	71.3 (4.0)
Sex, female, n (%)	1665 (59.6)	1338 (55.5)	7734 (56.5)	643 (57.2)
Education, <12 y, n (%)	648 (23.2)	322 (13.4)	3844 (25.8)	803 (71.4)
Medical history, n (%)				
Stroke	125 (4.6)	60 (2.5)	946 (6.8)	108 (9.6)
Hypertension	1754 (63.1)	1531 (63.7)	7678 (54.7)	802 (71.3)
Diabetes	289 (10.4)	341 (14.1)	2465 (16.9)	373 (33.2)
Coronary heart disease	492 (18.8)	422 (17.5)	3720 (26.9)	226 (20.1)
Depressive symptoms, n (%)				
Everything an effort $\dot{\tau}$	368 (13.2)	125 (7.3)	3100 (21.3)	195 (17.3)
Sleep was restless †	549 (19.7)	380 (22.2)	4126 (29.9)	271 (24.1)
Could not "get going" †	285 (10.2)	149 (8.7)	2984 (21.6)	110 (9.8)
Current antidepressant use	131 (4.7)	110 (4.6)	874 (6.4)	80 (7.1)
Functional difficulty, n (%)				
Eating	8 (0.3)	8 (0.3)	221 (1.5)	21 (1.9)
Dressing	28 (1.0)	21 (0.9)	981 (7.0)	59 (5.2)
Bathing	36 (1.3)	198 (8.2)	583 (4.3)	78 (6.9)
Using the toilet	11 (0.4)	50 (2.1)	542 (3.9)	33 (2.9)
Managing money or medications	16 (0.6)	41 (1.7)	305 (2.2)	122 (10.8)
Using the telephone	35 (1.3)	18 (0.8)	299 (2.1)	33 (2.9)
Getting out of a bed or chair	84 (3.3)	74 (3.1)	504 (3.5)	47 (4.2)
Walking 0.25 or 0.5 mi	461 (16.7)	244 (10.3) [‡]	3685 (27.0)	223 (19.8)
Walking up stairs	375 (13.6)	525 (22.0)	5367 (43.3)	383 (34.0)
Performing heavy housework	545 (19.7)	355 (14.9)	2773 (21.5)	429 (38.1)
Body mass index, kg/m ² , n (%)				
<18.5	38 (1.4)	22 (0.9)	189 (1.4)	§
18.5–24.9	940 (34.2)	728 (30.2)	4752 (35.3)	226 (20.1)
25.0-29.9	1198 (43.6)	1030 (42.7)	5665 (40.5)	423 (37.6)
30	573 (20.8)	631 (26.22)	3277 (22.7)	476 (42.3)

Abbreviations: CHS, Cardiovascular Health Study; FHS, the Framingham Heart Study; HRS, the Health and Retirement Study; SALSA, the Sacramento Area Latino Study on Aging; SD, standard deviation; BMI, body mass index.

NOTE. All values are number (percent) unless otherwise indicated. Numbers and percentages may not match as a result of missing data.

^{*} All subjects are age 65 to 79 years.

 $[\]dot{\tau}$ Individual questions from the Center for Epidemiologic Studies Depression (CES-D) scale that were common to all studies, categorized as experiencing symptoms 3 days or more per week during the past week.

[‡]Needs help walking 50 yards.

 $[\]ensuremath{\Sar}$ Participants included in the 18.5-to-24.9 group as a result of small numbers.

Table 3

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Predictors and corresponding hazard ratios for the final dementia risk calculator

	Hazard ratio (95°	Hazard ratio (95% confidence interval)	(le			
Predictor	CHS	FHS	HRS	SALSA	Combined Points	Points
Age, y	1.16 (1.12–1.21)	1.16 (1.12–1.21) 1.21 (1.15–1.29) 1.09 (1.07–1.11) 1.13 (1.06–1.20) 1.10	1.09 (1.07–1.11)	1.13 (1.06–1.20)	1.10	Variable*
Education, <12 y	1.49 (1.15–1.93)	1.49 (1.15–1.93) 1.62 (0.99–2.64)	2.97 (2.58–3.42)	2.97 (2.58–3.42) 1.29 (0.69–2.42) 2.43	2.43	6
$BMI, <18.5\ kg/m^2$	2.66 (1.37–5.18)	2.66 (1.37–5.18) 6.26 (2.51–15.61) 1.82 (1.25–2.64)	1.82 (1.25–2.64)	1	2.25	∞
Diabetes mellitus	1.27 (0.89–1.80)	1.27 (0.89–1.80) 0.86 (0.44–1.67)	1.27 (1.09–1.47)	1.27 (1.09–1.47) 2.28 (1.37–3.79) 1.29	1.29	ю
Stroke	2.01 (1.34–3.03)	2.01 (1.34–3.03) 1.24 (0.39–3.96)	1.75 (1.45–2.12)	1.75 (1.45–2.12) 2.99 (1.70–5.26) 1.86	1.86	9
Needs help, money/medications	1.90 (0.70–5.12)	1.90 (0.70–5.12) 2.56 (1.01–6.46)	2.76 (2.29–3.33)	2.76 (2.29–3.33) 1.93 (1.07–3.47) 2.65	2.65	10
Depressive symptoms †	1.48 (1.11–1.96)	1.48 (1.11–1.96) 2.15 (1.26–3.69)	1.79 (1.60–2.01)	1.79 (1.60–2.01) 1.92 (1.16–3.20) 1.77	1.77	9
C statistic	0.68 (0.65–0.72)	0.68 (0.65–0.72) 0.77 (0.73–0.82)	0.76 (0.74–0.77) 0.78 (0.72–0.83)	0.78 (0.72–0.83)		

Abbreviations: CHS, Cardiovascular Health Study; FHS, the Framingham Heart Study; HRS, the Health and Retirement Study; SALSA, the Sacramento Area Latino Study on Aging; BMI, body mass

* Points for age are determined as follows: 65 years, 0 point; 66–79 years, 1 point/year. For example, a 69-year-old patient would have 4 points.

 † Current usage of antidepressants or reporting that "everything was an effort" for 3 days or more a week for the past week.

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