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Q2

Abstract

**Q1** 

Alzheimer's

# Functional limitations and health care resource utilization for individuals with cognitive impairment without dementia: Findings from a US population-based survey

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Introduction: Little is known about functional limitations and health care resource utilization of people with cognitive impairment with no dementia (CIND).
Methods: Respondents with stable or progressive cognitive impairment (CI) after the first (index) indication of CIND in 2000–2010 were identified from the Health and Retirement Study (HRS). Re-

spondents never exhibiting CI were identified as potential controls. Propensity score–based optimal matching was used to adjust for differences in demographics and history of stroke. Differences between cohorts were assessed accounting for HRS survey design. **Results:** After matching, CIND respondents had more functional limitations (difficulty with  $\geq 1$  activities of daily living: 24% vs. 15%;  $\geq 1$  instrumental activities of daily living: 20% vs. 11%) and hospital stays (37% vs. 27%) than respondents with no CI (all P < .001). Seventy five percent of CIND respondents developed dementia in the observable follow-up (median time: ~6 years).

**Discussion:** Even before dementia onset, CI is associated with increased likelihood of functional limitations and greater health care resource use. © 2016 Published by Elsevier Inc. on behalf of the Alzheimer's Association. This is an open access

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Keywords: CIND; Functional limitations; Health care resource use; Burden; Dementia

## 1. Introduction

In 2007, the Aging, Demographics, and Memory Study estimated that the prevalence of dementia in the United States among individuals aged 71 years and older was 13.9% [1]. Alzheimer disease (AD) is the most common cause of dementia and accounts for 60%–80% of all dementias in the United States, followed by vascular dementias that account for up to 20% of all dementia patients [2,3].

Several studies have documented the functional and economic burden associated with AD and related dementias. For example, in 2009, more than a third of people with AD

\*Corresponding author. Tel.: 513-907-6945; Fax: ■■■. E-mail address: andrews\_jeffrey\_scott@lilly.com required some assistance with activities of daily living (ADL), such as dressing, bathing, and getting in and out of the bed [2]. The direct costs associated with AD and related dementias in the United States were estimated to be \$226 billion in 2015. In addition, nearly \$18 billion were attributable to costs associated with informal caregiving for people with AD and related dementias in 2014 [2].

However, little is known about the implications of cognitive impairment without dementia—a cognitive status known to develop as many as 18 years before clinical AD diagnosis [4] and affect approximately 10%–20% of Americans aged 65 years and older [5–7]. Recent studies have found that the incidence and prevalence of cognitive impairment without dementia are higher than those for dementia [7,8]. Prior research has also found that cognitive impairment with no dementia (CIND) is associated with

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110 substantial comorbidities and limitations in ADL and 111 instrumental activities of daily living (IADL). For 112 example, in the Cache County Study, Lyketsos et al. found 113 that participants with CIND had substantially higher rates 114 of comorbid conditions than those with normal cognition 115 116 [9]. Using data from a nationally representative sample of 117 participants aged 71 years and older, Gure et al. found that 118 45% of subjects with CIND had difficulty with >1 IADL 119 compared to 13% of subjects with normal cognition [10]. 120 In a similar study, Fisher et al. found that although respon-121 122 dents with CIND generally maintained their functional inde-123 pendence, their caregivers spent approximately 4 hours/day 124 to help them with IADLs [11]. 125

These studies, however, are limited to respondents aged 126 65 years and older and may not represent the broader popu-127 128 lation with cognitive impairment, many of whom may be 129 younger [2,6]. In addition, to the best of our knowledge, 130 no study to date has evaluated the health care resource use 131 among respondents with CIND, as compared to similar 132 respondents with normal cognition. Understanding the 133 134 functional and economic implications of CIND, including 135 in a younger population, is especially important given the 136 refinement of diagnostic criteria for CI and earlier stage 137 Alzheimer disease, and the emergence of new technologies 138 that may facilitate earlier diagnosis of cognitive 139 140 impairment and its causes [12]. In addition, new treatments 141 in development are likely to target patients at earlier stages 142 of disease. The objective of the present study was to compare 143 differences in patient characteristics, functional limitations, 144 and health care resource use between people with CIND and 145 146 those with no cognitive impairment (no CI) using a nation-147 ally representative sample of the US population enrolled in 148 the Health and Retirement Study (HRS). In addition, the 149 study assessed the rates of progression to dementia among 150 respondents with CIND, the time to progression, and the 151 152 burden associated with development of dementia in the sub-153 group of CIND respondents who progressed within 2 years 154 after incident CIND indication. 155

#### 2. Methods

### 2.1. Data

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161 The study used data from the RAND version M of the 162 publicly available HRS survey data sets for respondents 163 164 enrolled in the study. The survey design and questionnaires 165 have been described previously [13]. Briefly, the HRS is a 166 longitudinal household survey data set facilitating study of 167 retirement and health among the noninstitutionalized popu-168 lation over age 50 years in the United States. The HRS 169 170 includes rich demographic, clinical, economic, and health-171 related data. Of particular interest, the survey includes a 172 detailed cognitive assessment, which has been used to study 173 cognitive functioning among older Americans [5]. Certain 174 HRS data elements which are not part of the integrated 175 176 RAND HRS database (e.g., caregiver assistance) were

accessed directly from the core HRS files and merged at the respondent level. In addition, respondent level weights, strata, and cluster information (provided by HRS) were used in all analyses described in the following to account for the complex survey design.

#### 2.2. Measures of cognitive assessment

For each survey wave with valid cognitive assessment data, a respondent's cognitive status was determined following the approach used by Langa et al. [14,15]. Different stages of respondents' cognitive health were defined using the 27-point TICS scale for all selfrespondents (this scale includes: 10-word immediate and delayed recall tests of memory, serial 7s subtraction test, and the backwards counting test) and the 11-point composite scale for respondents requiring proxy informants (the composite scale includes: proxy's assessment of respondent's memory and limitations in five IADLs, and interviewer assessment of respondent's cognitive ability). Using the composite scores, respondent's cognitive status was classified as

- No CI
  - Self-respondent—score of 12 or higher
  - Proxy respondent—score of 0 to 2
- CIND
  - Self-respondent—score of 7 to 11
  - Proxy respondent—score of 3 to 5
- Dementia
  - Self-respondent—score 0 to 6
  - Proxy respondent—score of 6 or higher

#### 2.3. Study sample and time periods

Following the classification of cognitive functioning, the data were examined to identify respondents with earliest indications of CIND in 2000 or later. The first wave indicating CIND was considered as the index wave. To increase the likelihood of including respondents whose cognitive impairment is consistent with a progressive pattern due to an underlying neurodegenerative process such as AD, those with waves indicating an improvement in cognitive status (i.e., CIND followed by no impairment or dementia followed by CIND/no impairment) were excluded from the analyses. Respondents were required to have complete information regarding demographics and comorbidity profile, cognitive assessment, metrics of physical functioning, and resource use in the index wave as well as the waves immediately preceding and following the index wave.

Respondents with no evidence of cognitive impairment during the observable years of data were considered as potential controls. The index wave for the no CI cohort was selected at random, and respondents were required to have similar information as the CIND cohort in the index wave as well as the waves before and after the index wave. 177

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