

## Appendix. Evidence Tables for Dementia and Co-Occurring Chronic Conditions Systematic Literature Review\*

\*for Categories with Sufficient Evidence only

### Mortality

|   |    |
|---|----|
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|--|----|

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|---|----|

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SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author Yr Tracking # Study Design Duration  | # Sample (N) Mean age (SD) % Female Other Demographics Location DB   | Chronic Condition (CC): Definition Assessment Who made assessment   | Cognitive Impairment (CogImp)/Dementia (Dem): Measures, Criteria, % with Dem/CogImp, Mean Scores, Assessment  | Mortality Outcomes   | Quality   |
|---|--|---|---|--|---|
| First author<br><i>Each row/article is numbered consecutively in this table for easier reference.</i> | Sample size<br>Mean age (SD)<br>% Female<br>Other Demographics (e.g. education, race/ethnicity) if reported  | Which chronic conditions (CC) were included (e.g. depression). Overall comorbidity refers to a measure of multiple chronic conditions.<br><br>Assessment<br>Type of assessment (e.g. clinical exam/diagnosis, records review) along with details about the assessment, including who made the assessment. | Measures<br>Objective measures used to screen for or diagnose dementia-severity level cognitive impairment or dementia (e.g. MMSE). Cut off scores are provided where available.<br><br>Criteria<br>Diagnostic criteria used to determine presence of dementia (and dementia types) in the sample (e.g., DSM-III-R) | Outcomes indicating how cognitive impairment/ dementia impacts mortality in samples with multiple chronic conditions or one target condition<br><br>e.g., Hazard Ratios (HR) for mortality, mean survival times for people with and without dementia<br><br>Where provided, adjusted analyses will follow unadjusted analyses. | SAMPLE<br>N > 500<br><br>GENDER<br>>= 30% each male and female<br><br>MCC<br>>= 2 CC's<br><br>DEM<br>Both measures and criteria for measuring cognitive impairment<br><br>CC<br>Assessment details reported for measuring CC<br><br>MULTI<br>Multivariate Analysis<br><br>COV ASE<br>Adjusted for age, sex, education |
| Year of publication   | Study location   |   | % with dementia / cognitive impairment  |  |   |
| Article Tracking #  | Database (if reported)<br><br><i>Note: If demographic data is not available for the whole sample, other data will be provided here (e.g., age range, % female for w/ and w/o dementia; age range).</i> |   | Mean scores for objective measures (where available)  |  |   |
| Study Design (if reported)  |  |   | Details about the assessment of cognitive impairment / dementia (e.g. who conducted the assessment)   |  |   |

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| Author<br>Yr<br>Tracking #                          | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality   |
|---|--|--|---|--|---|
| 29 studies  | N=52,644   | Multiple chronic conditions  | Community-based samples: 17 studies<br><br>Clinic-based samples: 12 studies   |  | SAMPLE: 23<br>GENDER: 27<br>MCC: 29<br>DEM: 12<br>CC: 24<br>MULTI: 26<br>COV ASE: 12      |
| <i>Community-based</i>                              |  |  |   |  |   |
| 1. Gombojav<br>2011<br>#A5                          | N=2496<br>73.6(5.9)<br>57.7% female<br>% no formal education:  | Overall comorbidity, HTN,<br>Smoking (former and current),<br>Drinking, BMI  | Measures: MMSE<24<br>(18-23=mild severity; <18=severe)<br><br>Criteria: None  | % died (unadjusted):<br>CogImp: 54.8% (mild 49.1% and severe 61.2%)<br>No CogImp: 41.9%<br>p = 0.00  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: NA<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No  |
| Longitudinal<br>Mean 11.8 yrs                       | CogImp: 80%<br>No CogImp: 50%<br><br>INTL: South Korea<br><br>Database NR<br>(survey of Kangwa<br>County cohort) | Overall comorbidity: Subjects<br>answered yes or no to the<br>the question ~do you have any<br>chronic disease or past accident<br>or injury due to which you feel<br>uncomfortable in your daily life<br>including work?<br>HTN was measured in two BP<br>readings. HTN=SBP/DBP:<br><= 140/90 mm Hg | 44% with CogImp<br><br>Mean MMSE (SD):<br>mild severity: 17.6 (1.1)<br>severe severity: 11.9 (3.1)<br>no CogImp: 24.1 (2.9)<br><br>MMSE administered by the<br>investigation team | Adjusted HR All-Cause Mortality (95% CI):<br>No CogImp = referrant group<br>Male<br>severe: 1.33 (0.99 - 1.77); p = 0.0577<br>mild: 1.28 (1.01-1.61; p = NR<br>Female<br>severe: 1.59 (1.25 - 2.00); p < 0.001<br>mild: 1.32 (1.04-1.67), p = NR |   |
| 2. Nikolova<br>2011<br>#29                          | N=1164<br>82(7.3)<br>>50% female<br>>50% had >= h.s. educ  | Overall comorbidity<br><br>Standardized Instrument:<br>A 16-item questionnaire on<br>prevalence of common CC.<br>Number of diseases was coded<br>as: 0-1 disease, 2-3, 4-5, and<br>more than 6 diseases.   | Measures: SPMSQ >= 5<br><br>Criteria: None  | Adjusted OR Mortality (95% CI):<br>5.57 (0.84, 36.39)<br>p=NS  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Secondary<br>analysis of<br>SIPA RCT<br><br>3 years | INTL: Montreal, Canada<br><br>SIPA (Research Program<br>on Integrated Services for<br>the Elderly)               | Trained Interviewer  | 30% with cognitive impairment<br><br>Assessment details NR  |  |   |

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| Author<br>Yr<br>Tracking #            | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality  |
|---------------------------------------|--|--|--|---|--|
| 3. Millan-<br>Calenti<br>2010<br>#A28 | N=579<br>75.1 (7.5)<br>56.3% female<br>No formal education:<br>CogImp: 87.4%<br>No CogImp: 83.2%   | Overall comorbidity, Depression,<br>Visual /hearing impairments<br><br>Self-Report, Standardized Instrument,<br>Clinical Exam/ Diagnosis, Records<br>Review:<br><br>Probable clinical depression:<br>Assessed by a psychologist using<br>GDS-SF>/=6. Medical histories were<br>collected by a physician or a trained<br>nurse in charge of the participant<br>during the research. Participants<br>report was given by the patient or their<br>relatives according to medical records.<br>Comorbidity conditions were defined<br>according to the CCI. | Measures: MMSE<br><br>Criteria: None<br>20.6% with cognitive impairment<br><br>Mean (SD) MMSE: 23.9 (5.6)<br>(all subjects)<br><br>MMSE cut-offs based on pop-based age-<br>education adjusted norms: <20 for 80+<br>w/ 0 -4 yrs of education to <29 for age<br>65-69 college-educated. Majority of<br>subjects (>80%) had no formal<br>education. MMSE administered by a<br>psychologist. | Predicted 10-year survival expectancy<br>Mean (SD) (unadjusted):<br>No CogImp: 3.2 years (3.0)<br>CogImp: 2.3 years (2.9)<br>p = NR | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Design NR<br><br>Duration NR          | Older subjects (85+) were<br>more cognitively impaired<br>(29.4%) and presented more<br>co-occurring CogImp and<br>depression (22.1%) than<br>other ages.<br><br>INTL: Naron Council, A<br>Coruna, Spain<br><br>Database NR (community-<br>dwelling) |  |  |   |  |

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| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality   |
|--|--|--|---|---|---|
| 4. Wang<br>2010<br>#A239<br><br>Design NR<br><br>5 years<br>Mean (SD) f/u:<br>Dem: 5.15(0.40)<br>No Dem: 4.92<br>-0.32 | N=2162<br>All 65+<br>NR% female<br>Other demog NR<br><br>INTL: Beijing, China<br><br>10/66 Dementia Research<br>Group's population-based<br>study in China | Overall comorbidity<br><br>Assessment details and who did<br>assessment NR | Measures: CSI-D, CERAD; GMS-<br>AGECAT<br><br>Criteria: 10/66 Dementia Research<br>Group algorithm using: CSI-D<br>CERAD; GMS-AGECAT<br><br>6.3% with dementia<br><br>Cognitive measures were admin-<br>istered by trained research assts.<br><br>Other assessment details NR | Mortality rate (unadjusted):<br>Dem: 66.4%<br>No Dem: 37.2%<br>p<.01<br><br>Kaplan-Meier survival curves: Dem had SS shorter<br>survival time than No Dem (logrank test, p<0.001).<br><br>Five-year survival rates (unadjusted):<br>Dem: 16.1%<br>No Dem: 28.5%<br>p<.001<br><br>Severity of dementia (severe/mild, HR: 8.765, 95%CI: 4.436-<br>17.163) and substantial disability (HR: 5.503, 95%CI: 3.017-<br>8.135) were the most significant predictors of shortened<br>survival time in the multivariate analysis. | SAMPLE: Yes<br>GENDER: NR<br>MCC: Yes<br>DEM: Yes<br>CC: No<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment              | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality  |
|----------------------------|---|---|---|--|--|
| 5. Llinas-Regla, 2008 #500 | N=1153<br>Men: 80.3(5.2)<br>Women: 80.9 (5.1)<br>56.5% female<br><br>Avg 5 years education (all subjects)<br><br>INTL: Gironia, Spain<br><br>6 years (mean f/u 4.3 years) | Cancer, heart disease, stroke<br><br>Assessment details and who made made assessment NR | Measures: MEC<24 (Spanish MMSE)<br><br>Criteria: CAMDEX < 70 (for surviving participants); RCDI and DMS-III-R criteria (deceased)<br><br>10.6% with dementia<br><br>Mean (SD) MEC, all subjects:<br>Men = 27.5(4.7)<br><br>Women = 25.0 (5.7)<br><br>A neurologist and neuropsychologist conducted the CAMDEX interview with subjects to assess dementia diagnosis and severity. For subjects that died before the f/u study, a psychologist administered the RCDI to relatives and neurologists used DSM-III-R to diagnose dementia. Subjects who screened positive or were unable to complete the MEC received a further clinical eval and structure interview at home (inc. informants) by the neurologist and neuropsychologist. Persistent disagreements about diagnosis were coded as "no dementia." CDR assessed severity of dementia. | # / % Deaths (unadjusted):<br>Dem: 49 (40.2%)<br>No Dem: 188 (18.2%)<br>p = NR<br><br>Median survival (95% CI) (unadjusted):<br>Dem: 4.71 years (4.4-5.0)<br>No Dem: 6.2 years (6.0-6.3)<br>p<0.001 (log rank test = 30.59)<br><br>Median survival by dementia severity (unadjusted):<br>Mild dementia: 5.5 years (5.2-5.8)<br>Moderate dementia: 4.7 years (4.1-5.4)<br>Severe dementia: 3.2 years (2.6-3.9).<br>No vs Mild: p = NS<br>Mild vs moderate and moderate vs severe, p < .001<br><br>Mortality rate (95% CI) (unadjusted):<br>Dem: 9.8 per 100 person-years (7.3-12.9)<br>No Dem: 4.2 per 100 person-years (3.6-4.9)<br><br>PAR% of death related to dementia diagnosis = 11.8%<br><br>Unadjusted RR Mortality (95% CI) = 2.3 (1.7-3.2)<br><br>Adjusted HR Mortality (95% CI):<br>Severe Dementia: 5.7 (3.7-8.6)<br>Moderate dementia: 1.9 (1.1-3.5)<br>Mild dementia: 0.8 (0.4-1.5)<br><br>Other significant HRs were cancer, advanced age (90+), age 85-89, heart disease; severe dementia was the strongest predictor of death. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: No<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes   | Quality   |
|--|--|--|--|--|---|
| 6. Rothman<br>2008<br>#356<br><br>Prospective<br>cohort study<br><br>7.5 years | N=754<br>78.4(5.3)<br>64.6% female<br>Mean (SD) yrs education:<br>12.0 (2.9)<br>90.5% non-Hispanic white<br>39.5% live alone<br><br>U.S.: New Haven, CT<br><br>Precipitating Events<br>Project (PEP)<br>(community-dwelling) | Inurious Falls, Overall Comorbidity<br><br>Self-Report, Records Review:<br>Participants were asked about<br>overnight hospital stays during the<br>past month and to provide the<br>primary reason for admission.<br>An independent review of hospital<br>records of 44 participants indicated<br>high reliability of self-reported<br>information (kappa value of 0.89).<br><br>An injurious fall was defined as a<br>fall leading to a hospital admission<br>, head injury, or hematoma or<br>bruise of the face or scalp.<br><br>Overall CC: Mean 1.9(1.3) CC.<br>Self-reported, physician-diagnosed<br>CC's were HTN, MI, CHF, stroke,<br>diabetes mellitus, arthritis, hip fracture,<br>chronic lung disease, and cancer<br>(other than minor skin cancers).<br><br>Trained research staff conducted the<br>assessments | Measures: MMSE<24<br><br>Criteria: None<br><br>11.4% with cognitive impairment<br><br>Mean MMSE (SD):<br>CogImp: 21.7(1.5) (min. = 16)<br>No CogImp: NR<br><br>MMSE administered by trained<br>research staff. Other details NR. | Adjusted HR Mortality (95% CI):<br>Model 1 adjusted for demographics and CC:<br>2.4 (1.8-3.1)<br>Model 2 (~Model 1 + 6 other frailty criteria):<br>1.5 (1.1-2.1)<br><br>Notes:<br>Frailty criteria: low physical activity, cognitive impairment,<br>depressive symptoms, weight loss, gait speed, falls<br><br>In Model 1, lowest PA was the strongest predictor<br>of death (out of the 6 frailty criteria) | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

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| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes   | Quality  |
|-----------------------------|---|--|--|--|--|
| 7. Lyketsos<br>2007<br>#805 | N=198<br>CogImp: 86.1(6.73)<br>No CogImp: 84.9 (10.76)<br>% Female<br>CogImp: 80.6%<br>No CogImp: 75%<br>% Caucasian<br>CogImp: 79.9%<br>No CogImp: 89.1%<br>p=NS<br><br>% in large facility<br>CogImp: 70.9%<br>No CogImp: 85.9%<br>p=.02<br><br>U.S.: Central Maryland<br><br>The Maryland Assisted<br>Living Study (MD-AL) | Overall comorbidity, Depression,<br>Polypharmacy/High risk medications<br><br>Self-Report, Standardized Interview,<br>Records Review:<br><br>Residents completed the GMHR<br>(overall comorbidity) and CSDD<br>(depression). Number of Medications<br>came from chart reviews and<br>interviews with the<br>resident, family informant, and facility<br>staff members who knew the resident<br>well. | Measures: MMSE, Trails A and B,<br>HVLT<br><br>Criteria: None<br><br>68% with cognitive impairment<br><br>Mean MMSE (p<.001)<br>CogImp: 14.64 (7.67)<br>No CogImp: 25.84 (5.50)<br><br>Mean HVLT: (p<.001)<br>CogImp: 0.54 (1.12),<br>No CogImp: 4.82 (3.32)<br><br>Mean Trails B (p <.001)<br>CogImp: 500.84 (157.74)<br>No CogImp: 270.17 (151.96) | Mean # of days from assessment to discharge, death, or<br>censoring:<br>CogImp: 521.55 (407.74)<br>No CogImp: 707.30 (429.09)<br>p=.004<br><br>% Discharged (including death)<br>CogImp: 78.4<br>No CogImp: 59.4<br>p=.005<br><br>Difference in medial survival time (CogImp vs No CogImp):<br>209 days (@7.5 months)<br>p=NR<br><br>All above analyses are unadjusted | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |



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| Author       | # Sample (N)       | Chronic Condition (CC): | Cognitive Impairment (CogImp)/Dementia (Dem):                  | Mortality Outcomes | Quality |
|--------------|--------------------|-------------------------|--|--------------------|---------|
| Yr           | Mean age (SD)      | Definition              |  |                    |         |
| Tracking #   | % Female           | Assessment              | Measures, Criteria, % with Dem/CogImp, Mean Scores, Assessment |                    |         |
| Study Design | Other Demographics | Who made assessment     |  |                    |         |
| Duration     | Location           |                         |  |                    |         |
|              | DB                 |                         |  |                    |         |

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|                            |  |  |  |  |  |
|----------------------------|--|--|--|--|--|
| 7. Lyketsos<br>(continued) |  |  | Information from MMSE, Trails, and HVLT (no cut-offs reported) was brought to a panel consisting of the team that evaluated the resident in his or her facility, another geriatric psychiatrist, a geriatric medicine physician, a neuropsychologist, and a registered nurse. The panel used a consensus process, based on all available information, to make diagnoses and render opinions about the completeness of evaluation and treatment for dementia on the basis community-care standards (e.g., minimize inappropriate medication use). |  |  |
|----------------------------|--|--|--|--|--|

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| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality  |
|----------------------------|---|--|---|---|--|
| 8. Guhne<br>2006<br>#1096  | N=908<br>83(4.7)<br>75% female<br>Low education (p<.001):<br>Dem: 45%, No Dem: 19%<br>Living location (p<0.001)<br>Nursing Home: Dem: 53%,<br>No Dem: 10%<br>Hom: Dem: 47%, No Dem:<br>90%<br>Marital status (p=NS)<br>INTL: Leipzig, Germany<br><br>Leipzig Longitudinal Study of<br>the Aged (LEILA75+) | Overall comorbidity (diabetes,<br>stroke, and/or myocardial infarction)<br><br>Self-Report:<br>Subjects or relative's reported<br>history of at least one CC<br><br>Who made assessment NR | Measures: MMSE, CDR, SIDAM<br><br>Criteria: SIDAM neuropsych-ological<br>battery (inc. MMSE and DSM-III-R<br>criteria), CDR<br><br>9% with dementia<br>-AD: 45, VaD: 18, Other: 15<br><br>SIDAM neuropsychological test battery<br>administered by<br>psychologist. SIDAM includes MMSE,<br>third-party information on psychosocial<br>impairment and a section for clinical<br>judgment including severity rating<br>according to DSM-III-R criteria.<br>Cognitive criteria for dementia diagnosis<br>based on SIDAM or CDR + proxy<br>interviews. Diagnosis made by<br>physician. | # / % Deceased (unadjusted):<br>Dem: 40 (51%); No Dem: 159 (19%)<br>p = NR<br><br>Mean (95% CI) survival (unadjusted):<br>Dem: CI: 3.1 years (2.8-3.4)<br>No Dem: 4.0 years (3.9-4.0)<br>p<0.001<br><br>Adjusted HR Mortality (95% CI):<br>Incident dementia = 2.42 (1.62-3.63) | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

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|--------------------------------|---|---|---|--|---|
| 9. Cacciatore<br>2005<br>#1557 | N=1332<br>74.1(6.4)<br>NR% female<br>Other Demog NR             | Hypertension, Overall Comorbidity<br><br>Clinical Exam/Diagnosis:<br>BP was measured according Joint<br>National Committee on Detection,<br>Evaluation and Treatment of High<br>Blood Pressure criteria: 3 BP<br>measurements at 2-min intervals<br>when subject had been sitting for @ 1<br>hr, using a standard mercury<br>sphygmomanometer. The<br>disappearance of sound (phase V)<br>was used for diastolic reading. Mean<br>value of the last two recorded<br>measures was considered for SBP<br>and DBP.<br><br>NR how overall comorbidity was<br>measured. Mean 2.4 (1.3) CC (for all<br>subjects)<br><br>A physician conducted the BP<br>assessments | Measures: MMSE<24<br><br>Criteria: None<br><br>Mean MMSE=25.4(4.8)<br>(all subjects)<br><br>A physician administered the MMSE | Adjusted RR mortality (referent=DBP 80-89):<br><i>DBP 90-99 (~ HTN)</i><br>CogImp: 1.51 (0.83-2.74), no CogImp: 1.39 (0.88-2.19)<br><i>DBP &gt;99 (~HTN)</i><br>CogImp: 3.41 (1.40-8.29), No CogImp: 1.60 (0.68-3.76)<br>DBP <80<br>CogImp: 2.84 (2.53-5.24), No CogImp: 1.64 (1.11-2.41)<br><br>Adjusted RR Mortality (referent = SBP 130-139)<br><i>SBP 140-159 (~ HTN)</i><br>CogImp: 0.66 (0.38-1.14), No CogImp: 1.09 (0.68-1.63)<br><i>SBP &gt;159 (~ HTN)</i><br>CogImp: 0.53 (0.28-1.02), No CogImp: 0.93 (0.55-1.57)<br><i>SBP &lt;130</i><br>CogImp: 0.53 (0.25-1.10), No CogImp: 0.85 (0.48-1.52) | SAMPLE: Yes<br>GENDER: NA<br>MCC: No<br>CC: Yes<br>DEM: No<br>MULTI: Yes<br>COV ASE: No |

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| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality  |
|--|---|--|---|---|--|
| 10. Fitzpatrick<br>2005<br>#1496<br><br>Prospective<br>Cohort<br><br>5 years | N=2798<br>75.1(NR)<br>~59% female<br>NoDem-AD-Mixed-VaD<br>Non-white (%):<br>9.3 - 17.1 - 13.9 - 17.7<br>Educ some HS (%):<br>19.2 - 37.1 - 27.8 - 27.4<br>Educ HS grad (%):<br>29.4 - 23.3 - 26.5 - 32.3<br>Educ some college (%):<br>26.4 - 18.4 - 21.2 - 16.1<br>Educ college or + (%):<br>25.0 - 21.1 - 24.5 - 24.2<br><br>U.S.: Forsyth county, NC,<br>Washington county, MD,<br>Sacramento county, CA,<br>Pittsburgh, PA<br><br>Cardiovascular Health<br>Cognition Study, and<br>offshoot of the<br>Cardiovascular Health Study | Stroke, Heart disease, Hypertension<br><br>Clinical Exam/Diagnosis, Records<br>Review:<br>This included data on demographics,<br>anthropometry, blood pressure,<br>psychosocial<br>interviews, depression, medical<br>history, health behaviors, physical<br>function, hematology, and<br>medications.<br><br>Who made assessment NR | Measures (screening): 3MS, Digit<br>Symbol Benton Visual Retention<br><br>Those failing the screening and still<br>living completed detailed<br>neuropsychological testing at the clinic<br><br>Measures (diagnosis): TICS, IQCODE,<br>DQ, Neuropsychiatric Inventory<br><br>Criteria: ~DSM-IV, NINCDS-ADRDA<br>(AD), CADDTC (VaD)<br><br>17% with dementia<br>AD: 245, VD: 62<br><br>A committee of neurologists and<br>psychiatrists evaluated data to<br>determine dementia diagnoses.<br>Participants were required to have<br>impairments in two cognitive domains,<br>which did not necessarily include<br>memory (this correlates very closely to<br>DSM-IV). Dementia type of dementia<br>was classified using NINCDS-ADRDA<br>(AD) and CADDTC (VaD), and MRIs. | Unadjusted HR Mortality (95% CI):<br>(referrant = no dementia):<br>Total dementia: 39.6% - 3.9 (3.2-4.6)<br>AD: 32.2% - 3.0 (2.3 - 3.8)<br>Mixed: 43.7% - 3.8 (2.9-4.9)<br>VaD: 53.2% - 5.7 (4.0-8.2)<br>p<0.0001<br><br>Adjusted HR Mortality (95% CI):<br>(referrent = no dementia):<br>Total dementia: 2.8 (2.3 - 3.4)¶<br>AD: 2.1 (1.6 - 2.7) ¶<br>Mixed: 2.5 (1.9 - 3.3)<br>VaD: 4.4 (3.1 - 6.3)<br>p<0.0001<br><br>Median Survival (95% CI) (adjusted):<br>No Dementia: 11.0 years (10.5-11.7)<br>VaD: 3.9 years (3.5-4.2)<br>AD: 7.1 years (6.7-7.5)<br>Mixed: 5.4 years (5.2-6.0) | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                                      | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality   |
|---|---|---|--|---|---|
| 11. Tschanz<br>2004<br>#1737<br><br>Longitudinal<br><br>5 years | N=4683<br>All 65+<br>Dem: 83.3(7)<br>No Dem: 74.7(6.8)<br>Dem: 64% female<br>No Dem: 56% female<br>Mean (SD) yrs education<br>Dem: 12.4(2.8)<br>No Dem: 13.3(2.9)<br>Dem: 28% in NH<br>No Dem: <1% in NH<br>99% Caucasian<br><br>U.S.: Cache County, UT<br><br>The Cache County Study<br>on Memory in Aging | PD, Asthma, CVD, CHF, HTN,<br>Diabetes, Head Injury, Pulmonary<br>Disease, Hypercholestermia,<br>Ulcer, Cerebrovascular Disease,<br>Depression, Smoking, Drinking<br><br>Self-Report (interview):<br>Self- or proxy report of common<br>CC. Cancer was not included b/c<br>primary intent was to identify risk<br>factors for dementia. Selected IW<br>questions to ascertain CC are<br>available on journal's website.<br><br>Who made assessment NR | Measures: 3MS, IQCODE, DQ<br><br>Criteria: DSM-III-R, NINCDS-<br>ADRDA, NINDS-AIREN<br><br>7.6% dementia<br>AD: 207 ,VD: 54, AD/VaD (Mixed):<br>31, OD (Other Dementia): 63<br><br>Subjects with DQ suggestive of<br>cognitive impairment received<br>clinical assessment, conducted<br>by research nurses and psycho-<br>metricians. These assessments<br>included a brief physical exam,<br>standardized BP measurement,<br>neurologic exam, and psycho-<br>metric testing. A board-certified geriatric<br>psychiatrist and neuro-psychologist<br>reviewed findings with the examiners<br>and assigned working diagnoses of<br>dementia (DMS-III-R) and rated<br>dementia severity using the CDR. One<br>year later, of subjects still living,<br><br>83.9% were examined by geriatric<br>psychiatrists and 65.9% of these<br>underwent MRI scanning and standard<br>lab tests for differential diagnosis. A<br>panel of experts reviewed all data for<br>those with suspected dementia and<br>assigned diagnoses of AD, VaD, or<br>other disorders using standard criteria. | Unadjusted Mortality Rate:<br>Dem: 82%<br>No Dem: 22%<br>p=.0001<br><br>Crude OR = 16.23, 95% CI 12.27-21.48, p = 0.0001<br><br>Adjusted RR Mortality: 2.99 (95% CI 2.53-3.53)<br><br>PAR% (Adjusted) = 16.6%<br><br>Dementia had risk of mortality 2-3x greater than other<br>life-threatening CC's, across all age groups. PAR% were<br>also higher than other CC's.<br><br>Relative hazard of death with dementia was highest at<br>ages 65 to 74 (RR = 7.3), but the high prevalence of<br>dementia after age 85 resulted in 27% PAR among the<br>oldest old.<br><br>Mortality increased substantially with severity of dementia.<br>AD shortened survival time most dramatically in younger<br>participants, but VaD posed a greater mortality risk among<br>the oldest old. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: No<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                 | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment | Mortality Outcomes   | Quality  |
|--|---|---|--|--|--|
| 12. Feil<br>2003<br>#1851                  | N=7482<br>65+ (58% age 65-74)<br>61.7% female<br>Education for CogImp:<br>40% < 8 yrs, 11% > 12 yrs                           | Overall comorbidity, Stroke, Cancer,<br>Heart Attack, HTN, Diabetes, Hip<br>Fracture<br><br>Self-Report, Clinical Exam/Diagnosis:   | Measures: modified SPMSQ (>= 2<br>errors)<br><br>Criteria: None<br><br>23.6% w/ cognitive impairment                         | RR mortality (adjusted): 1.68 (95% CI 1.53-1.86)<br>p<.0001<br><br>RR mortality due to CogImp was greater than RR for other<br>six CC (diabetes was closest at 1.62). Older age (75+) was<br>stronger predictor, and male gender was a<br>similar predictor.   | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Longitudinal<br>(mortality)<br><br>6 years | U.S.: East Boston, MA and<br>rural Iowa<br><br>Established Populations for<br>Epidemiologic Studies of the<br>Elderly (EPESE) | Subjects were asked about history of<br>CC. HTN was assessed from direct<br>measurement of BP according to<br>Hypertension Detection and Follow-Up<br>Program.<br><br>Interviews conducted by trained<br>interviewers (in-person and phone) | Trained interviewers conducted the<br>assessments  | The effect of CI and specific CC on mortality is mainly<br>additive (NS interactions between the survival curves). That<br>said, 6-year survival for CI + CC was 40%-50%, whereas no<br>CC and no CI was 80%.<br><br>The theory that the assoc btw CI and mortality is primarily<br>attributable to cognitive impairment aggravating chronic<br>medical illness was not supported in this study. CI predicted<br>mortality indep of the number of chronic illnesses. |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality  |
|------------------------------|--|--|---|--|--|
| 13. Nguyen<br>2003<br>#3430  | N=2625<br>72.9(NR)<br>58.5% female<br>Education > 12 years:<br>Mod-Severe: 3.1%<br>Mild: 1.8%<br>No CogImp: 14.5%  | Stroke, Cancer, CVD, Diabetes,<br>HTN, Depression, Hip Fracture<br><br>Self-Report, Standardized<br>Instrument:<br>Depression: CES-D by trained.<br>interviewer. All other CC were self-<br>report: "has a doctor ever told you<br>that you had any of the following<br>conditions?"                             | Measures: MMSE<24<br><br>Criteria: None<br><br>37% with cognitive impairment<br><br>MMSE 18-23 = "Mild" and 0-17 =<br>"Moderate-Severe" severity.<br>Serial-sevens item is not used in<br>the MMSE-Spanish. "Don't know"<br>and "refusals" were counted as<br>errors. Trained bilingual interviewers<br>conducted all interviews in Spanish or<br>English, depending on the respondent's<br>preference. | % Deceased at 5-year f/u (unadjusted):<br>No CogImp: 263 (16.0%)<br>Mild: 263 (16.0%)<br>Moderate-severe: 59 (45.7%)<br><br>20.3% of sample was deceased at follow-up<br><br>Adjusted HR Mortality (95% CI):<br>Mild: 1.56 (1.28-1.92)<br>Mod-Severe: 2.41 (1.74-3.34)<br>(data not reported)  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| 14. Ganguli<br>2002<br>#2037 | N=1064<br>74.9 (5.5)<br>57.5% female<br>58.4% >= H.S. grad<br>Largely blue-collar, low<br>income, European descent   | Depression, overall comorbidity<br><br>Self-Report, Standardized<br>Instrument:<br>Depression: mCES-D > 5<br>(interviewer-administered, max. 20<br>points)<br>Overall comorbidity: total # of<br>prescription medications was<br>used as a continuous measure of<br>morbidity, to control for medical<br>burden. | Measures: MMSE < 24<br><br>Criteria: None<br><br>8.2% with cognitive impairment<br><br>In-home screening interview including<br>the MMSE<br><br>Who made assessment NR  | Adjusted RR Mortality:<br>3-Year Mortality 1.43, p=.16<br>5-Year Mortality: 1.16, p=.48<br>10-Year Mortality:1.26, p=.12<br><br>In post hoc analyses, when baseline IADL disability was<br>excluded from the model, baseline MMSE was a significant<br>predictor of mortality:<br><br>3-Year Mortality 2.2, p<.001<br>5-Year Mortality 1.52, p=.04<br>10-Year Mortality 1.55, p = .002 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
|                              | U.S.: Texas, New Mexico,<br>Colorado, Arizona,<br>California.<br><br>Hispanic Established<br>Population for the<br>Epidemiological Study of the<br>Elderly | U.S.: SW Pennsylvania<br>Monongahela Valley<br>Independent Elders Survey<br>(MoVIES)   |   |  |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                          | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality   |
|---|---|---|---|---|---|
| 15. Helmer<br>2001<br>#2227                         | N=2923<br>65+<br>58.3% female                                   | Stroke, Cancer, Heart Disease, HTN,<br>Respiratory disease,<br>Diabetes, Dyspnea  | Measures: MMSE, other tests<br><br>Criteria: DSM-III-R, NINCDS-ADRDA,<br>Hachinski Scale  | % Died (unadjusted):<br>Dem: 39.1%, No Dem: 22.2%<br>p = NR   | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>CC: Yes |
| Prospective<br>population-<br>based cohort<br>study | INTL: Southwest France<br>(Gironde and Dordogne<br>region)      | Self-Report:<br>Self-reported diseases or symptoms.<br>Overall comorbidity = at least one of<br>these diseases or symptoms. | 10% with dementia<br>-AD: 189, VaD: 70, Other: 22   | Cause of death (unadjusted):<br>CVD, Stroke, Cancer, Respiratory, Other<br>Dem: 20%, 12.7%, 12.7%, 10%, 10.9%<br>No Dem: 32.3%, 9.2%, 24.6%, 6.4%, 7.2%<br>p=NR   | DEM: Yes<br>MULTI: Yes<br>COV: No                 |
| Duration NR   | Personnes Ages Quid<br>(PAQUID) cohort                          | Psychologist collected self-report<br>data  | MMSE and other tests administered by<br>psychologists to obtain info for<br>DSM-III-R criteria for dementia.<br>Participants who met DSM-III-R criteria<br>were seen by a senior neurologist, who<br>confirmed and completed DSM-III-R<br>criteria for dementia, NINCDS-ADRDA<br>criteria for AD, and the Hachinski score<br>for VaD. Informant was consulted when<br>available and all available info was<br>used. A consensus meeting was used to<br>definitely classify each case. | Adjusted RR Mortality (95% CI):<br>Dem: 1.80 (1.46, 2.21)<br>AD only: 1.72 (1.34, 2.21)<br><br>According to causes of death:<br>Adjusted RR Mortality for dementia (95% CI):<br>Cerebrovascular disease: 2.29 (1.26, 4.17) □<br>Respiratory disease: 2.78 (1.40, 5.51) □<br><br>Adjusted RR Mortality for those with AD (95% CI):<br>Respiratory disease: 2.82 (1.30, 6.17) □ |   |



SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality  |
|-----------------------------|---|---|---|---|--|
| 16. Bruce<br>1995<br>#3434  | N=3538<br>67.61 (11.69)<br>59% female<br><= 8 years education: 86%  | Overall comorbidity, PD, Stroke,<br>Arteriosclerosis, Myocardial Infarction,<br>Other Neurological Disease, Epilepsy,<br>seizures or head injuries  | Measures: MMSE<24<br>(<18=severe, 18-23 mild)<br><br>Criteria: None   | Unadjusted RR Mortality:<br>Men: mild: 1.96, severe: 3.91<br>p=0.0001 for both<br>Women: mild: 2.41, severe: 4.62<br>p=0.0001 for both  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Design NR<br>9 years        | U.S.: New Haven, CT<br><br>New Haven Epidemiologic<br>Catchment Area study  | Self-Report:<br>Subjects were asked if they had ever<br>been diagnosed with the CC<br><br>Who made assessment: NR   | 6% with cognitive impairment<br><br>Assessment details NR   | % Deceased within 9 years follow-up (unadjusted):<br>Men: CogImp: 74.07%, No CogImp: 32.41%<br>Women: CogImp: 73.53%, No CogImp: 25.72%<br>p=NR<br><br>Adjusted: For both men and women, lower scores on the<br>MMSE decreased the risk of survival (with the effect<br>stronger for younger respondents than older respondents)<br>(p<.05) |  |
| 17. Kukull<br>1994<br>#3067 | N=104<br>NR<br>% female:<br>Probable AD: 60%<br>Probable AD: 57.4%<br>Other Dem: 45.4%<br>No Dem: 58.8%                   | Stroke, Cancer, Ischemic Heart<br>Disease<br><br>Records Review:<br>ADRC participants completed a<br>medical history and clinical exam<br>when enrolled and at annual follow-up<br>visits | Measures: MMSE<br><br>Criteria: DSM-III (R), NINCDS-ADRDA<br><br>84% with dementia<br>-Probable AD: 55, Possible AD: 10,<br>Other: 22   | Underlying cause of death-cancers and CVD:<br>No Dementia: 59%<br>Possible AD: 80%<br>Probably AD: 35%<br>Other dementia: 36%<br><br>Probable AD cases died at similar ages, regardless of level<br>of cognitive impairment.  | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No    |
| Duration NR                 | Caucasian: 88-95%<br><br>U.S.: Seattle, WA<br><br>The University of<br>Washington Alzheimer's<br>Disease Patient Registry |   | MMSE was obtained from last follow-up<br>visit at AD Research Center was<br>obtained from the patient registry.<br><br>"No dementia" group were patient<br>registry participants who did not meet<br>dementia criteria. |   |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author                               | # Sample (N)  | Chronic Condition (CC):   | Cognitive Impairment (CogImp)/Dementia (Dem):   | Mortality Outcomes  | Quality   |
|--------------------------------------|---|---|---|---|---|
| Yr                                   | Mean age (SD)   | Definition  |   |   |   |
| Tracking #                           | % Female  | Assessment  | Measures, Criteria, % with  |   |   |
| Study Design                         | Other Demographics  | Who made assessment   | Dem/CogImp, Mean Scores, Assessment   |   |   |
| Duration                             | Location  |   |   |   |   |
|                                      | DB  |   |   |   |   |
| Clinic-based                         |   |   |   |   |   |
| 18. Aguero<br>-Torres, 1999<br>#2566 | N=989<br>84.3(4.3)<br>77% female<br>< 8 years education:<br>Dem: 74.6%<br>No Dem: 53.7% | Overall comorbidity (inc: cancer, stroke not followed by dementia hip fracture, heart disease<br><br>ICD Diagnosis, Records Review<br>Electronic inpatient records on hospital discharge diagnoses<br><br>Who made assessment: NR | Measures: MMSE, free recall & recognition of random words, digit span.<br><br>Criteria: DSM-III-R<br><br>12.8% with dementia<br>-AD: 102 ,VD: 21<br><br>Mean MMSE=25.2(5.3) (all subjects)<br><br>Clinical eval using standard protocol: family and personal history (nurses), clinical exam (physicians), psychological tests (psychologists). Informant used when subject unable to answer. Two indep diagnoses (examining physician; specialist) | % Died after 5-year follow-up (unadjusted):<br>Dementia: 70%, No Dementia: 35%, p=NR<br><br>Unadjusted Mortality Rates (100 person-yrs) (95% CI)<br>Men Age 77-84: Dem: 28.8 (9.2-67.3), no Dem: 8.1 (5.5-11.1)<br>Men 85+: Dem: 24.8 (10.8-48.9); No Dem: 14.6 (11.2-18.7)<br>Women 77-84: Dem: 17.5(8.4-32.3); no Dem: 3.9(2.8-5.3)<br>Women 85+: Dem: 23.6(18.2-30.0); no Dem: 10.0(8.4-11.8)<br>p=NR<br><br>Unadjusted RR Mortality:<br>Men: 77-84: 3.6(1.4-9.1); 85+: 1.7(0.8-3.5)<br>Women: 77-84: 4.5(2.2-8.9); 85+: 2.4(1.8-3.2)<br>p=NR<br><br>PAR% (unadjusted):<br>Men 77-84: 10(8-13); 85+: 5 (4-7)<br>Women 77-84: 17(13-20); 85+: 18(14-20)<br>p=NR | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Longitudinal (mortality)             |   |   |   |   |   |
| Duration NR                          | INTL: Stockholm, Sweden<br><br>Kungsholmen Project                                      |   |   |   |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author       | # Sample (N)       | Chronic Condition (CC): | Cognitive Impairment (CogImp)/Dementia (Dem): | Mortality Outcomes | Quality |
|--------------|--------------------|-------------------------|---|--------------------|---------|
| Yr           | Mean age (SD)      | Definition              |   |                    |         |
| Tracking #   | % Female           | Assessment              | Measures, Criteria, % with                    |                    |         |
| Study Design | Other Demographics | Who made assessment     | Dem/CogImp, Mean Scores, Assessment           |                    |         |
| Duration     | Location           |                         |   |                    |         |
|              | DB                 |                         |   |                    |         |

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18. Aguerro-Torres (continued)

Mean Survival Time from f/u (95% CI) (unadjusted):  
 Dem: 3.0 yrs (2.7-3.4); No Dem: 4.2 yrs (4.1-4.3); p<0.001  
 Similar findings for AD vs No Dem, and VaD vs No Dem

Adjusted RR of 5-yr Mortality (95% CI)  
 AD: 2.0 (1.5-2.7); VD: 3.3 (2.0-5.3)  
 p=NS (AD vs VD)

There were similar risk factors for death when stratifying by age and dementia types.

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics    | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality  |
|---|--|--|---|---|--|
| 19. Bursi<br>2006<br>#1273  | N=1832<br>Median: 82 (38-102 yrs)<br>NR % female<br>Other Demog NR | Myocardial infarction,<br>Hypertension, Lipidemia, Diabetes,<br>Substance use (smoking)  | Measures: NR<br><br>Criteria: DSM-IV, H-ICDA code   | % Survival at 10 years (unadjusted):<br>Dementia: 22% (95% CI: 20, 25)<br>No Dementia: 39% (95% CI: 36, 43)<br>p < 0.001  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: No<br>MULTI: Yes<br>COV ASE: No |
| Cohort study  | U.S.: Rochester, MN  | ICD Diagnosis, Records Review,   | 50% with dementia   | # Cardiac Deaths: (unadjusted)<br>Dementia: 313 (55% attributed to coronary disease)<br>No Dementia: 370 (58% attributed to coronary disease)<br>p = .01  |  |
| Mean f/u:<br>Dem: 6.1 years<br>(0.4-19.5),<br>No Dem: 8.2<br>years (1.1-<br>19.5)<br>p < .001 | Rochester Epidemiology<br>Project                                  | Standardized Instrument:<br><br>MI assessed using ICD-9 codes from<br>discharged diagnoses in medical<br>records. Trained nurse abstractors<br>validated the diagnosis of MI using<br>standardized criteria for definite or<br>probable MI (cardiac pain, biomarker<br>values, and Minnesota coding of the<br>electrocardiogram). NR how other<br>CC's were collected. | Searched for H-ICDA codes in medical<br>records. Each potential case (at least<br>one H-ICDA code) was screened by<br>trained nurse abstractors. A neurologist<br>confirmed the presence of dementia<br>using DSM-IV. | Adjusted HR Mortality: 1.67, p=NS (0.516)<br><br>Adjusted HR Cardiac Death (95% CI):<br>0.82, 0.70-0.95; p = 0.010<br><br># / % Sudden death: (adjusted)Dementia: 130 (42%)<br>No Dementia: 133 (36%)<br>p = NS (0.135) |  |
|   |  | Trained nurse abstractors reviewed<br>the records and compared against<br>standardized criteria.   |   | Survival free of cardiac death after accounting for the<br>competing risk of noncardiac death was better among<br>subjects with dementia (p = 0.014).   |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment                                 | Mortality Outcomes  | Quality   |
|-----------------------------|---|--|--|---|---|
| 20. Foley<br>#2604<br>1999  | N=2905<br>0% female<br>77.1(4.2)  | Stroke, Asthma, COPD, HTN,<br>Diabetes, Arthritis, CHD<br><br>Clinical Exam/Diagnosis, Records<br>Review, Structured IW:<br>HTN: measured SBP >= 160 mm Hg,<br>or measured DBP >= 95 mm<br>Hg, or use of antiHTN meds.   | Measures: CASI<74<br><br>Criteria: DSM-III-R<br><br>9.1% with cognitive impairment; 2.5%<br>with dementia<br><br>Note: 6 subjects w/ dementia had<br>CASI>74 | Adjusted OR for 3-year Mortality (95% CI):<br>(for cognitive impairment)<br>2.26 (1.64, 3.10) | SAMPLE: Yes<br>GENDER: No<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Longitudinal<br>(mortality) | 86.3% married (all subjects)  |  |  |   |   |
| Approx.. 3<br>years         | U.S.: Oahu, HI<br><br>Honolulu-Asia Aging<br>Study (HAAS), using<br>survivors from a cohort study<br>of cardiovascular disease<br>(Honolulu Heart Program<br>(HHP)) | COPD: structured interview data<br>indicating a physician's diagnosis of<br>emphysema at least 2 years earlier or<br>coughing or bringing up phlegm for<br>periods of at least 3 consecutive<br>months for 2 or more years.<br>History of a physician's diagnosis of<br>diabetes, arthritis, and asthma were<br>obtained from the structured interview.<br>CHD and Stroke (CVA) history:<br>surveillance data from<br>hospital/physician discharge records | Who made assessment NR   |   |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality  |
|--|--|---|---|--|--|
| 21. Freels<br>2002<br>#2055  | N=248<br>45+<br>% female NR<br>Other demog NR  | Stroke, Myocardial Infarction, Atrial<br>fibrillation, HTN, Lipidemia, Diabetes,<br>Depression, Substance use (smoking)   | Measures: MMSE, BOMCT, Formal<br>neuropsychological testing<br><br>Criteria: ~NINDS-AIREN   | Unadjusted:<br>Survival at 2 years:<br>AD: 80.9%, VaD: 89.4%, No Dem: 92.6%<br>Survival at 5 years:<br>AD: 54.6%, VaD: 58.6%, No Dem: 77.1%<br>Survival at 7 years:  | SAMPLE: No<br>MULTI: Yes<br>GENDER: NR<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>COV ASE: No |
| Cohort study<br><br>7 years (median<br>f/u: AD: 5.9<br>years, VaD:<br>5.8, No Dem:<br>6.1) | U.S.: Chicagoo. Illinois<br>Database NR<br>Patients were referred from<br>a hospital-based stroke<br>registry and an academic AD<br>center | Self-Report, Standardized<br><br>Instrument:<br>An epidemiologic interviewer<br>administered standardized<br>questionnaires to determine CC's.<br>Stroke: The study neurologist<br>completed the Stroke Data Bank<br>Neurologic Examination and an<br>unstructured neurologic interview with<br>informants. | 77% with dementia.<br>AD: 113 ,VD: 79<br><br>All study patients completed diagnostic<br>measures with trained interviewers and<br>informants completed a structured<br>neurologic interview. The following<br>criteria were used: AD = met dementia<br>criteria and no other conditions<br>contributing to cog impairment; VaD =<br>dementia + diagnosis of stroke by<br>Stroke Data Bank criteria,<br><br>and a temporal relationship between<br>stroke and dementia onset (VaD<br>diagnosis predated both of the currently<br>most commonly used diagnostic<br>systems today). These criteria are<br>consistent with NINDS-AIREN. Persons<br>meeting both VaD and AD criteria were<br>excluded.<br><br>The "No dementia" group included<br>patients who met criteria for neurologic<br>dysfunction due to vascular disease<br>(stroke) but did not meet criteria for<br>dementia. | Adjusted HR Mortality (95% CI):<br>AD vs VaD: HR 0.91, p= 0.685<br>Adjustment for age and CDR reverses the direction of the<br>HR to a protective, but NS, effect.<br><br>Adjusted HR Mortality (95% CI):<br>VaD vs No Dem: HR 1.30, p 0.450<br>The increase in risk for patients with VaD is lower and no<br>longer significant after adjusting for age.<br><br>Note: No Dem = No Dementia for this study |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #             | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes   | Quality   |
|--|---|---|--|--|---|
| 22. Kammoun<br>2000<br>#2307           | N=342<br>84.94(6.9)<br>62% female<br>Other Demog NR             | Overall comorbidity (CCI),<br>Cancer, CVD (heart failure,<br>myocardial infarction), pulmonary<br>embolism, Cerebrovascular<br>disease, CNS heaemorrhage,<br>cachexia, gastrointestinal diseases,<br>cancer, metabolic disorders,<br>depression. and renal failure. | Measures: NR<br><br>Criteria: DSM-IV criteria<br><br>35% with dementia<br>-AD: 21 ,VD: 34, MD (Mixed): 65  | Pulmonary infections and cardiovascular diseases were<br>the most common causes of death. NS differences in<br>causes of death between persons with and No<br>dementia. (unadjusted analysis, data not reported) | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>Multi: No<br>COV ASE: No |
| Retrospective<br>correlation<br>study. | INTL: Geneva, Switzerland                                       | ICD Diagnosis:<br>ICD- 10 for somatic diseases and<br>DSM-IV for mental illnesses.  | Subjects with comprehensive<br>clinical, neuropsychological and<br>neuropathological data were included.<br>Cases with major neuropsychiatric<br>illness, alcoholism, head trauma or PD<br>were excluded. Who made assessment<br>NR. |  |   |
| Duration NR                            | Database NR (university-<br>based, geriatric care center)       | Who made assessment NR  |  |  |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality  |
|--|---|---|---|--|--|
| 23. Lavretsky<br>2010<br>#A368   | N=498<br>74.5(NR)<br>49.8% female<br>73.3% Caucasian,<br>15.7% Asian,<br>6.0% African-American,<br>4.6% Hispanic                                      | Stroke, CVD (MI, CHF, HTN)<br>Diabetes, Hyperlipidemia,<br>Depression<br><br>Standardized Instrument, Clinical<br>Exam/Diagnosis:<br>No other details were provided<br>about the assessment | Measures: Neuropsychological<br>Battery (not specified)<br><br>Criteria: DSM-IV, NINDS-AIREN,<br>CADDTC<br><br>33% with dementia<br>AD: 87 ,VD: 36<br><br>Participants received a<br>comprehensive clinical evaluation.<br>Cognition was evaluated using<br>neuropsychological battery (details NR).<br>Dementia diagnoses were made at a<br>multidisciplinary case conference using<br>DSM-IV criteria and NINDS-AIREN for<br>AD and CADDTC for VaD) | Unadjusted Mortality rate:<br>Dem: 52.0%<br>No Dem: 21.1%<br>p<0.001<br><br>Unadjusted HR Mortality (95% CI):<br>7.4 (4.9-11.4)<br>p<0.001<br><br>Adjusted HR Mortality (95% CI):<br>4.7 (2.9-7.5)<br>p=NR | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Multicenter<br>longitudinal<br>study<br><br>12 years<br>(median 4.7 yrs) | Location<br>DB<br><br>U.S.: California<br><br>Ischemic Vascular<br>Dementia Program Project<br>(4 clinical sites from 3<br>academic dementia centers) |   |   |  |  |



SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #     | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality  |
|--------------------------------|---|---|--|---|--|
| 24. Magaziner<br>2005<br>#1287 | N=2153<br>Dem: 82.7 (7.3)<br>No Dem: 80.2 (7.7)<br>Female (%)<br>Dem: 69.4<br>No Dem: 72.8<br>White (%)<br>Dem: 76.2<br>No Dem: 84.7%<br>Married (%)<br>Dem: 26.8%<br>No Dem: 21.5%<br>Education <= 8 years (%):<br>Dem: 37.9%<br>No Dem: 28.9%<br><br>U.S.: Maryland (nursing<br>homes)<br><br>Database NR | Overall comorbidity, Fractures or<br>Injuries<br><br>Self-Report, Records Review:<br>Overall comorbidity: # of CC at NH<br>admission inc. CHD, CHF,<br>cerebrovascular disease, COPD, liver<br>disease, peripheral vascular disease,<br>seizure disorder, peptic ulcers,<br>arthritis, cancer, uncontrolled HTN<br>(SBP >=160 or DBP >=90 mmHg),<br>and malnutrition/underweight (BMI<br><=20). This information was obtained<br>from interviews with significant others<br>at admission, with the exception that<br>BMI and HTN information was<br>abstracted from the nursing home<br>chart. Two other comorbidity<br>measures (a modified CCI's and the<br>Diagnostic Cost Group/Hierarchical<br>Coexisting Condition risk adjustor,<br>with the dementia diagnosis omitted<br>from both measures) had the same<br>direction, magnitude, and SS of the<br>effect of dementia on any study<br>outcome so the comorbidity count only<br>is presented.<br><br>Fractures and injuries occurring in the<br>first week of NH stay were included<br>because these originated in the NH.<br>Study abstractors collected follow-up<br>data from NH home charts. | Criteria: NR<br><br>Diagnosis: DSM-III-R criteria<br><br>48.2% with dementia<br><br>An expert panel of geriatric psychiatrists<br>and neurologists, and a geriatrician<br>determined dementia status using DSM-<br>III-R criteria. 2 panelists gave indep<br>diagnoses: dementia, no dementia, or<br>difficult to diagnose. A larger panel<br>rendered a diagnosis if the two panelists<br>disagreed. Data was obtained from<br>medical records and interviews with<br>staff, family, and residents (conducted<br>by lay interviewers).<br><br>Note: No dementia includes subjects No<br>dementia and those that were difficult to<br>diagnose | Mortality rate per 100 person-days (unadjusted):<br>Dem: .09, No Dem: .14<br>p=NR<br><br>Unadjusted RR mortality (95% CI):<br>0.61 (0.53-0.71)<br>p <= 0.001<br><br>Adjusted RR mortality (95% CI)<br>0.63(0.51-0.77)<br>p <= 0.001<br><br>During the first 90 days of the nursing home stay, residents<br>with dementia had significantly lower rates of mortality if not<br>admitted for rehabilitative care under a Medicare qualifying<br>stay (RR 0.25, 95% CI 0.14-0.45) than residents No<br>dementia. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #    | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment                              | Mortality Outcomes   | Quality   |
|-------------------------------|---|---|---|--|---|
| 25. Meerman<br>2008<br>#640   | N=502<br>Dem: 81.7 (7.1)<br>No Dem: 81.1 (7)<br>68.5% female<br>Most subjects were low or<br>middle class   | CVD (heart failure, myocardial<br>infarction), Cerebrovascular Diseases<br>(CVA,<br>TIA), HTN, Diabetes (type 1 and 2),<br>Obesity (BMI>25) | Measures: NR<br>Criteria: DSM, NINCDS-ADRDA<br><br>50% with dementia  | Median survival (25-75%) (unadjusted):<br>Dem: 2.3 (1.3-3.6) years<br>No Dem: 3.7 (2.1-6.2) years<br><br>1-year: Dem: 73.7%, No Dem: 92.8%<br>2-year: Dem: 53.7%, No Dem: 79.7%<br>3-year: Dem: 30.2%, No Dem: 66.9%<br>Unadjusted HR Mortality (95% CI):<br>3.9 (2.3-3.9) | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Retrospective<br>case-control | INTL: Nijmegen, The<br>Netherlands<br><br>Continuous Morbidity<br>Registration database, Dept<br>of Family Medicine of<br>Radboud University<br>Nijmegen in the Netherlands | ICD Diagnosis, Records Review:<br>GP diagnoses coded as ICHPPC<br>codes in medical records.   | In the GP registration, dementia is<br>classified as a syndrome only; ICHPPC-<br>2 classification does not distinguish the<br>different types of dementia | Adjusted HR Mortality (95% CI):<br>(CC within 5 years prior to dementia diagnosis)<br>Cerebrovascular: 1.54 (1.13-2.09)<br>Cardiovascular: 1.91 (1.48-2.46)<br>Obesity: 0.68 (0.52-0.90)   |   |
| Maximum 23<br>years           |   |   |   |  |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality  |
|---|--|---|--|---|--|
| 26. Zekry<br>2009<br>#463<br><br>Prospective<br>cohort<br><br>Duration NR | N=435<br>85.3(6.7)<br>74% female<br>Level 1 education (lowest)<br>Dem: 55.6%<br>No Dem: 59.9%<br>Acute and rehab patients<br>in geriatric hospital<br><br>INTL: Geneva, Switzerland<br><br>Database NR<br>(acute and rehabilitation<br>geriatric hospital) | Overall comorbidity, Obesity<br><br>Standardized Instrument:<br>Overall comorbidity: The same<br>geriatrician calculated CCI for<br>each patient using the patient's<br>medical records. Higher scores =<br>greater comorbidity. Dementia was<br>not included in the calculation of CCI<br>or comorbidity. Obesity = BMI, using<br>the short version MNA administered<br>on admission, by the same nurse. | Measures: MMSE, Short Cognitive<br>Evaluation Battery, CDR<br><br>Criteria: DSM-IV-TR, NINCDS-<br>ADRD, NINDS-AIREN<br><br>44% with dementia<br>AD: 77, VD: 21, Mixed: 82, Other: 11<br><br>The same neuropsychologist assessed<br>all subjects for clinical dementia, at least<br>one week after admission, to avoid the<br>effects of concomitant delirium. Based<br>on screening results with the objective<br>measures, the same neuropsychologist<br>carried out a comprehensive<br>standardized<br><br>neuropsychological assessment, to<br>determine the etiology and severity of<br>clinical dementia. Dementia severity<br>was assessed with the CDR (2,<br>moderate, and 3, severe dementia).<br>DSM IV-TR was used for dementia<br>diagnosis. NINCDS-ADRD and<br>ADDTC used for AD, and NINDS-<br>AIREN used for VaD. Cerebral imaging<br>was also carried out. | # / % Death in Hospitals:<br>Dem: 7 (3.9%), No Dem: 12 (5.8%), p = 0.641<br><br>Unadjusted OR Death in Hospital<br>(No Dem / Normal Cognition = referant group)<br>AD: 0.21 (0.03-1.67); p=0.142<br>VaD: 0.81 (0.10-6.580); p=0.846<br>Mixed: 1.06 (0.36-3.09); p=0.360<br>CDR 2-3: 0.91 (0.31-2.67); p=0.868<br><br>Adjusted OR Death in Hospital<br>AD: 0.20 (0.01-3.81); p=0.286<br>VaD: VaD dropped bc of collinearity<br>MD: 0.70 (0.07-7.30); p=0.762<br>CDR 2-3: 1.28 (1.12 -13.52); p=0.837 | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #             | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics           | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality  |
|--|---|--|--|---|--|
| 27. Arfken<br>1999<br>#2622            | N=455<br>77.3(7.3)<br>68% female<br>Mean yrs educ: 9.5 (3.2)<br>71% Black | Depression, Overall comorbidity<br><br>Standardized Instrument; Records<br>Review:<br>GDS (depression); CCI Index (from 18<br>different CC diagnoses obtained from<br>medical records)               | Measures: DRS < 103<br><br>Criteria: None<br><br>14% w/ severe cog. Impairment<br><br>Mean DRS: 120.3 ±16.2 (all subjects)   | Adjusted OR Mortality:<br>2.13 (95% CI: 1.13,4.02)<br>p = .02   | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Design NR<br><br>Mean f/u 1.9<br>years | U.S.: Detroit, Michigan<br><br>Database NR<br>(rehabilitation hospital)   | Mean GDS: 7.91 ± 5.76 (range 0-28;<br>>10=depression)<br><br>Neuropsychology dept staff conducted<br>the GDS; Hospital Management<br>Information System staff for the<br>conducted the Record Review | Screen administered by staff from the<br>neuropsychology dept<br><br>"No CogImp" includes subjects without<br>cognitive impairment (53%) and mild<br>cognitive impairment (33%). |   |  |
| 28. Gale<br>1996<br>#3433              | N=921<br>65+<br>45% female<br>Other Demog NR                              | Stroke (ischemic), Cancer,<br>Cardiovascular Diseases<br><br>ICD Diagnosis<br>Who made assessment NR   | Measures: AMTS </= 7<br><br>Criteria: NA<br><br>6.1% with cognitive impairment   | Adjusted RR All-Cause mortality (95% CI):<br>Model 1: CogImp: 2.2 (1.6 to 2.9)<br>Model 2: CogImp: 2.0 (1.4 to 2.7) | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: No<br>MULTI: Yes<br>COV ASE: No |
| Cohort study<br><br>20 years           | INTL: Britain<br><br>Database NR<br><br>Family practices in 8 areas       |  | A geriatrician administered the AMTS   | Adjusted RR Mortality from stroke (95% CI):<br>Model 1: CogImp: 3.3 (1.7 to 6.2)<br>Model 2: CogImp: 2.8(1.4-5.5)   |  |
|  |   |  |  | *Model 1 adjusts for age and sex only   |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality  |
|-----------------------------|--|---|---|---|--|
| 29. Stump<br>2001<br>#3431  | N=2975<br>68.1(7.4)<br>69.2% female<br>Mean (SD) yrs education:<br>CogImp: 7.5 (3.8)<br>No CogImp: 9.1 (3.1)<br>% Black:<br>CogImp: 85.4%<br>No CogImp: 61.2%                      | Cancer, COPD, Artherosclerosis,<br>CAD, Cerebrovascular disease,<br>HTN, Diabetes, Arthritis,<br>Depression, Substance Abuse<br>(Problem Drinking), Smoking,<br>Obesity, High Cholesterol,<br><br>Standardized Instrument, Records<br>Review:<br>Depression was assessed using<br>CES-D >= 16). Alcohol drinking<br>problems were measured using the<br>CAGE>= 2 positive responses.<br>For the records review, data on<br>medical diagnoses, lab data, and<br>smoking status were assessed using<br>data routinely collected and stored in a<br>comprehensive electronic medical<br>record. Physicians order all laboratory<br>tests and procedures. A diagnosis or<br>diagnoses must be entered as the<br>reason for the patient's visit after each<br>encounter. | Measures: SPMSQ (>2 errors)<br><br>Criteria: None<br><br>5.2% with moderate-severe<br>cognitive impairment<br><br>Trained research assistants<br>collected the SPMSQ.<br><br>No other assessment details are<br>provided. | % Survived:<br>CogImp: 78.5%<br>No CogImp: 78.6%<br><br>% Died:<br>CogImp: 40.8%<br>No CogImp: 21.4%<br>p<.001<br><br>Unadjusted HR Mortality (95% CI):<br>CogImp: 2.31 (1.83-2.90) p<=.01<br><br>Adjusted HR Mortality (95% CI):<br>CogImp: 1.70 (1.32-2.19), p<=.0001 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Prospective<br>cohort study |  |   |   |   |  |
| 5-7 years                   |  |   |   |   |  |
|                             | U.S.: Indianapolis, Indiana<br><br>Regenstrief Medical<br>Record System: Over<br>600,000 outpatient visits<br>and 60,000 inpatient stays<br>per year across several<br>insitutions |   |   |   |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                            | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes   | Quality   |
|---|---|--|--|--|---|
| 8 studies   | N=4513  | Parkinson's Disease (PD)   |  |  | SAMPLE: 2<br>GENDER: 7<br>MCC: 0<br>DEM: 4<br>CC: 4<br>MULTI: 2<br>COV ASE: 2           |
| 30. Buter<br>2008<br>#603                             | N=233<br>75(8.4)<br>51.1% female<br>Other Demog NR              | Idiopathic PD<br>Clinical Exam:<br><br>Idiopathic PD required at least 2 of 4 cardinal signs (i.e., resting tremor, rigidity, akinesia, and postural instability); at least a moderate response to a dopaminergic agent; and no other evident potential cause of parkinsonism. Patients with neurologic symptoms or radiologically proven structural brain abnormalities compatible with brain diseases other than PD were excluded. | Measures: CDR, MMSE; If MMSE >16: Neuropsychological Battery visual memory, visuospatial ability, and executive functions)<br><br>Criteria: DSM-III-R, Caregiver-based dementia interview<br><br>Scores below the lowest quartile of population-based, age- and education-corrected normative cognitive test data were considered cognitive impairment.<br><br>60% with dementia | Life Expectancy, years (unadjusted):<br>Men Age 70: Dementia: 4.2, No Dementia: 8.0<br>Men Age 70: Dementia: 4.2, No Dementia: 8.0<br>Women Age 70: Dementia: 5.7, No Dementia: 11.0<br>Men Age 75: Dementia: 3.4, No Dementia: 6.2<br>Women Age 75: Dementia: 4.6, No Dementia: 8.7<br>Men Age 80: Dementia: 2.7, No Dementia: 4.7<br>Women Age 80: Dementia: 3.8, No Dementia: 6.8<br>Men Age 85: Dementia: 2.2, No Dementia: 3.6<br>Women Age 85: Dementia: 3.0, No Dementia: 5.2<br>Men Age 90: Dementia: 1.8, No Dementia: 2.4<br>Women Age 90: Dementia: 2.7, No Dementia: 3.9<br><br>P = NR | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Prospective<br>longitudinal<br>cohort<br><br>12 years | INTL: Norway<br><br>Database NR                                 | Assessment made by neurologist   | Independent trained raters (blind to eval) administered the tests in an interview with the patient and an informant; An experienced clinician (neurologist) conducted the neurologic exam and made the diagnosis.  |  |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality  |
|--|---|---|---|---|--|
| 31. Hobson<br>2010<br>#A217<br>NR<br>4 years           | N=166<br>Survivors: 74.9(8.3)<br>Deceased: 77.8(7.1)<br>Survivors: 36%<br>Deceased: 38%<br><br>INTL: England and Wales<br><br>Database NR | Parkinson's Disease<br><br>Clinical Exam/Diagnosis:<br>PD Brain Bank clinical diagnostic<br>criteria for probable PD.<br><br>Who made assessment NR | Measures: CAMDEX CAMCOG<br>Neuropsychological battery-not<br>specified<br><br>Criteria: DSM-IV<br><br>48.2% with dementia<br><br>Mean (SD) CAMDEX CAMCOG:<br>Alive 82.2(19.2)<br>Deceased: 73.9(16.8)<br>p=.01<br><br>Other assessment details NR | Standardised Mortality Ratios (unadjusted)<br>Dem: 3.10 (2.39-3.96)<br>No Dem 1.15 (0.75-1.69)<br>p<0.001<br><br>Univariate models of association with mortality (log<br>rank statistic):<br>Dementia = 42.8 (p<0.0001) (greatest predictor, more<br>than age and institutional placement)<br><br>Average LE (life expectancy) Ages 55 and 74:<br>Dem: 7.5 (SD 3), No Dem: 12.4 (SD 7)<br>Average AAD (avg age death) Ages 55 and 74:<br>Dem: 72.5 (SD 4), No Dem: 77.8 (SD 7)<br><br>Average LE (life expectancy) Ages 75+:<br>Dem: 2 (SD 1), No Dem: 4.7 (SD 3)<br><br>Average AAD (avg age death) Ages 75+:<br>Dem: 89.5 (SD 6), No Dem: 92.2 (SD 6)<br><br>p = NR | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: No<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment | Mortality Outcomes  | Quality   |
|---|---|--|--|---|---|
| 32. Levy<br>2002<br>#2017<br><br>Design NR<br><br>mean f/u was<br>3.9 years +/-<br>2.2 years) | N=180<br>71(10.3)<br>53.9% female<br>Avg 11.1(4.8) yrs educ<br>55.6% White, non-<br>Hispanic, 35.6% Hispanic,<br>8.9% Black, non-Hispanic<br>61.8% English as primary<br>language<br><br>U.S.: Manhattan, NY<br><br>Database NR | Idiopathic PD. Patients with<br>parkinsonism or a Parkinson-plus<br>syndrome were excluded, as were<br>patients who presented memory<br>loss or dementia before the motor<br>manifestations of PD.<br><br>Disease Registry:<br>Patients were identified through<br>the development of a registry for PD in<br>the community for individuals<br>considered to have PD or related<br>disorder. Patients were identified from:<br>Hospital admission and discharge<br>lists, lists from various ambulatory<br>care sites, and practitioners both in<br>the hospital and in the community.<br><br>Who made the assessment NR | Measures: NR<br><br>Criteria: DSM-III-R<br><br>29% with dementia<br><br>Other assessment details NR                          | RR Mortality (95% CI) (adjusted):<br>3.7 (2.0-7.2), p < 0.001<br><br>When both incident dementia and EPS (extrapyramidal<br>signs) severity were analyzed in a Cox model, RR for<br>incident dementia was:<br>Adjusted RR=2.2, 95% CI (1.1-4.5), p < 0.04<br>**This RR includes UPDRS score in the model**<br><br>RR's by severity of CDR are provided. CDR 1, 2, and 3 or 4<br>are significant (range from 3.6 to 5.0) | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: No<br>MULTI: Yes<br>COV ASE: Yes |



SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                                  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality   |
|---|---|--|---|---|---|
| 33. Marder<br>1991<br>#3225<br><br>Design NR<br><br>5 years | N=257<br>Dem: 69.6(7.9)<br>No Dem: 64 (9.9)<br>38.5% female<br>Age at onset of motor<br>signs:<br>Dem: 59.2(12.1)<br>No Dem: 54.8 (11.2)<br>p=0.006<br><br>U.S.: New York City<br><br>Database NR | Parkinson's Disease<br><br>Clinical exam/diagnosis:<br>2:4 cardinal motor manifestations<br>of PD (resting tremor, rigidity,<br>gait impairment, bradykinesia)<br><br>Who made assessment NR | Measures: UPDRS Item 1<br><br>Criteria: DSM-III-R<br><br>28% with dementia<br><br>Structured interview (some in-<br>person, some over phone).<br>Diagnoses made by neurologist<br>or psychiatrist or health<br>professional. UPDRS Item 1<br>(intellectual impairment) = 2 (if<br>performed by neurologist. | % Deaths (unadjusted)<br>Dem: 23.3%, No Dem: 15.2%, p= NS<br><br>% Deaths for only age 50+ w/ PD for 10 yrs:<br>Dem: 26.2%, No Dem: 11.8%, p < .02<br>No significant differences in causes of death (NS).<br>Pneumonia and MI most common causes.<br><br>Multivariate:<br>Dementia is not a SS predictor of death when age and<br>sex are included in model.<br>Dementia is significant predictor of death (p<0.005)<br>when only include age 50+ w/ PD for 10 years<br>Dementia group has SS decreased survival (p=.049) | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: No<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics          | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality  |
|---|--|--|---|--|--|
| 34. Parashos<br>2002<br>#3429   | N=178<br>73(NR)<br>41.6% female<br>36% < HS education<br>68.6% married   | Parkinson's Disease<br><br>Records Review:<br>PD diagnostic criteria: (1) 3: 4<br>cardinal signs (rest tremor, rigidity,<br>bradykinesia, and postural reflex<br>impairment), 2 cardinal signs +<br>diagnosis of typical PD by a<br>neurologist, OR 2 cardinal signs<br>and improvement of symptoms<br>with antiparkinsonian therapy (as<br>documented in the medical record);<br>(2) no other apparent cause of<br>parkinsonism; and (3) absence of<br>signs of more extensive nervous<br>system dysfunction (not expected in<br>PD). PD cases with evidence of<br>dementia preceding or within the first<br>year of motor symptoms were<br>excluded. | Measures: NR<br><br>Criteria: in Beard, 1995<br><br>26% with dementia<br><br>Dementia diagnosis required<br>fulfillment of 3 criteria: (1)<br>evidence of previously normal<br>intellectual and social function,<br>irreversible decline in intellectual<br>and social function, dementia as a<br>predominant symptom, and definite<br>evidence of memory impairment; (2) at<br>least 2 of the following symptoms<br>(patient must be fully alert):<br>disorientation, decline in personality<br>and/or behavior, dyscalculia, apraxia<br>and/or agnosia, problems with<br>language, and for impairment in<br>judgment and/or abstract thinking; and<br>(3) cognitive impairment lasting at least<br>6 months | Unadjusted RR for Mortality (95% CI)<br>(absence of dementia)<br>0.3; (0.2-0.5)<br>p<.001<br><br>Longer survival was associated w/ absence of<br>dementia. | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: No<br>MULTI: No<br>COV ASE: No |
| Historical<br>cohort study  | U.S.: Olmstead County,<br>Minnesota<br>Rochester Epidemiology<br>Project |  |   |  |  |
| Median f/u:<br>Dem: 7.0 yrs<br>(<1-15.5)<br>No Dem: 7.5 yrs<br>(<1-15.5 yrs). |  |  |   |  |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #    | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes   | Quality   |
|-------------------------------|---|--|--|--|---|
| 35. Mitchell<br>2000<br>#2454 | N=2426<br>All age 65+<br>NR% female<br>Other Demog NR           | Parkinson's Disease<br><br>Clinical Exam/Diagnosis, Records<br>Review, Medication list:<br>2:4 PD signs assessed by<br>physician in a neurological<br>exam: bradykinesia of either the<br>face or limbs, resting tremor;<br>increased tone in the limbs, and<br>abnormal gait and posture. The<br>precise nature of the abnormal gait or<br>posture was not noted. To exclude<br>subjects with spasticity, those with<br>increased tone in the presence of<br>abnormal reflexes or focal<br>neurological signs were excluded.<br>Subjects who met PD criteria but who<br>were on medications that commonly<br>cause extrapyramidal side effects<br>were excluded from the survival<br>analysis (any neuroleptic or<br>metoclopramide). Subjects with a<br>previous diagnosis of Idiopathic PD<br>were also excluded from the survival<br>analysis. | Measures: 3MS < 78<br><br>Criteria: DSM-III-R, NINCDS<br>-ADRDA<br><br>29.7% with AD   | Unadjusted RR Mortality (95% CI):<br>AD: 1.43 (1.09-1.87)<br>No Dem: 1.82 (1.31-2.53)<br><br>Adjusted RR Mortality (95% CI):<br>AD vs No Dem: 1.39 (1.13-1.72)<br><br>An interaction term between AD and PD to test for<br>effect modification was NS with mortality when added to the<br>model (RR, 0.87; 95% CI, 0.59-1.29). | SAMPLE: Yes<br>GENDER: NR<br>MCC: No<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Prospective<br>cohort         | INTL: Canada  |  |  |  |   |
| 5 years                       | Canadian Study of Health<br>and Aging (CSHA)                    |  | Following a comprehensive<br>evaluation by physician and nurse, a<br>consensus diagnosis was reached using<br>DSM-III-R. AD based on NINCDS-<br>ADRDA and includes probable or<br>possible AD. |  |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                    | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics             | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment | Mortality Outcomes  | Quality  |
|---|---|--|--|---|--|
| 36. Lo<br>2009<br>#52                         | N=573<br>70.5(7.4)<br>38.7% female<br>80.6% non-Hispanic<br>whites.         | Clinical subtypes of PD included<br>modified tremor dominant, PIGD,<br>or mixed based on the presence<br>of cardinal signs within 2 years of<br>Recent PD diagnosis  | Measures: MMSE < 24<br><br>Criteria: None<br><br>33% with cognitive impairment   | Kaplan-Meier curve showed dementia severity<br>correlated with HR. Severe cognitive impairment had<br>the strongest impact on survival among all predictors.<br><br>Adjusted HR Mortality (95% CI) (age, sex, ethnicity):<br>Severe: 2.71 (1.88-3.91) p=<.001 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: No<br>MULTI: Yes<br>COV ASE: Yes |
| Cohort study<br><br>Till death or<br>10 years | U.S.: Northern California<br><br>Parkinson Epidemiology<br>at Kaiser Study. | Records Review, Admitted to<br>Clinic/Program, Disease Registry,<br>Computerized pharmacy system:<br>Potential incident PD cases were<br>identified through comprehensive<br>clinical inpatient and outpatient<br>databases, a computerized pharmacy<br>system, and KPMCP physician<br>referrals. The records of all potential<br>cases were reviewed by a movement<br>disorder specialist for eligibility and<br>diagnostic status. > 90% subjects<br>were newly diagnosed with PD by a<br>neurologist (median 1 month since<br>first symptom onset). All eligible cases<br>met modified Core Assessment<br>Program for Intracerebral<br>Transplantation/ Hughes diagnostic<br>criteria at the time of diagnosis. | MMSE interviewer was trained by<br>a neurologist   | Adjusted HR Mortality (95% CI) (extended model):<br>Severe: 2.16 (1.31-3.55) p=.002   |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                                     | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes   | Quality   |
|---|---|---|---|--|---|
| 37. Ebmeier<br>1990<br>#3286<br><br>Design NR<br><br>Mean 3 yrs, 7<br>mo. | N=500<br>All 60+<br>NR% female<br>Other Demog NR<br><br>INTL: Aberdeen, Scotland<br><br>Database NR | 48.6% with idiopathic PD<br><br>Clinical Exam/Diagnosis:<br>Comprehensive exam by a<br>research psychiatrist with<br>experience in geriatric medicine | Measures: MSQ < 8<br><br>Criteria: None<br><br>54.7% with cognitive impairment<br><br>MSQ collected during the original PD<br>examination. NR who made the<br>assessment. | OR Mortality (95% CI) (unadjusted):<br>CogImp: 5.00 (2.18-13.60)<br>No CogImp: 2.83 (0.82-12.79) | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: No<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB                           | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality   |
|--|---|--|--|---|---|
| 6 studies  | N=2302  | Stroke   |  |   | SAMPLE: 0<br>GENDER: 3<br>MCC: 2<br>DEM: 4<br>CC: 4<br>MULTI: 5<br>COV ASE: 2             |
| 38. Desmond<br>2002<br>#2085<br><br>Longitudinal<br><br>Median f/u:<br>Dem: 2.6 yrs(0-10)<br>Database NR<br>No Dem: 4 yrs (0-10.5) | N=453<br>72(8.3)<br>52.5% female<br>36.4% education <= 8 yrs,<br>72.6% non-White<br><br>U.S.: New York City | Ischemic stroke<br><br>Clinical Exam/Diagnosis, Records<br>Review, Admitted to Clinic/<br>Program:<br>7-10 days after stroke onset,<br>neurologists specializing in stroke<br>administered a structured<br>neurologic examination and<br>documented any history of stroke,<br>TIA, or exposure to risk factors for<br>cerebrovascular disease based on<br>review of medical records and a<br>structured interview administered to all<br>patients and knowledgeable<br>informants. A comprehensive medical<br>history was also recorded. Based on<br>the review of clinical features and<br>brain imaging performed immediately<br>after stroke, patients were classified<br>by stroke syndrome using a<br>modification of the methods of the<br>Stroke Data Bank. | Measures: Neuropsychological<br>Battery- not specified<br><br>Criteria: DSM-III-R and similar<br>criteria as later proposed by<br>NINDRS-AIREN<br><br>26.3% with dementia<br><br>During baseline assessment and<br>annual examinations, all patients<br>were administered a comprehensive<br>battery of neuropsychological tests<br>developed for use in epidemiologic<br>studies of dementia, with testing<br>performed in either English or Spanish<br>based on the language spoken in the<br>subject's home. Knowledgeable<br>informants were administered the<br>Blessed Functional Activity Scale, which<br>taps the cognitive aspects of activities of<br>daily living. We required deficits in<br>memory and two or more additional | Unadjusted RR Mortality (95% CI):<br>2.98 (2.15-4.12), p = NR (SS)<br><br>Crude Incidence Mortality Rate<br>Dem: 15.9 deaths/100 person-years<br>No Dem: 5.37 deaths/100 person-years<br>p<0.0001 (log-rank test comparison of survival curves)<br><br>% Causes of death:<br>Pneumonia: Dem: 26.1%, No Dem: 20.7%, p = 0.516<br>Cerebrovasc.disease: Dem: 29.8%, No Dem: 23.0%,<br>p = 0.422<br>Cardiac disease: Dem: 18.2%, No Dem: 20.7%, p=0.752<br>Malignancies: Dem: 5.3%, No Dem: 19.0%, p=0.070.<br><br>The risk of death did not differ significantly between patients<br>with cerebrovascular disease as the primary basis for their<br>dementia syndrome (51.5% of whom had died or 14.37<br>deaths per 100 person-years) and patients with AD with<br>concomitant stroke (53.3% of whom had died or 18.51<br>deaths per 100 person-years) by a log-rank test.<br><br>Adjusted RR Mortality (95% CI):<br>2.37 (1.64-3.43) | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes | Quality |
|----------------------------|---|--|---|--------------------|---------|
| 38. Desmond<br>(continued) |   | Other CC: Atrial fibrillation, CHF, MI, HTN, Prior Stroke, Diabetes, Consistent drinking, Consistent smoking<br><br>Same assessment process as described above | cognitive domains as determined in the neuropsychological evaluation as well as functional impairment not solely related to physical disability documented with the Blessed scale.<br><br>When patients were aphasic, we required that they exhibit evidence of nonverbal memory impairment. We defined impairment within any cognitive domain as any neuropsychological test score within that domain falling below a predetermined cutoff that was selected in a pilot study. In previous work, we noted that a diagnosis of dementia based on this paradigm had the greatest validity as a predictor of death among stroke patients, while diagnoses based on less stringent operationalized criteria, the MMSE, and the examining neurologist's clinical judgment were only weakly related. |                    |         |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                                | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality   |
|---|---|---|--|---|---|
| 39. Liebtrau<br>2003<br>#1838<br><br>Design NR<br>3 years | N=494<br>85+<br>71.1% female<br>25% higher education<br><br>INTL: Gothenburg, Sweden<br><br>Database NR | Stroke (first-ever)<br><br>Self-Report:<br>First-ever stroke was obtained from<br>self-reports (n =235 for 85-88)<br>and key informant interviews (n =190)<br>for age > 88.<br><br>Who made assessment NR | Measures: NR<br><br>Criteria: DSM-III-R<br><br>30% with dementia<br><br>The study included nurse home<br>visits, physical examinations by<br>geriatricians (including assessment of<br>physical disorders), neuro-psychiatric<br>examinations, key informant interviews<br>(to assess cognitive measures, also by<br>neuropsychiatrist), and lab tests | Unadjusted mortality rate, 3-year (95% CI):<br>Dem: 67.9% (53.6-79.7)<br>No Dem: 30.0% (17.1-46.7)<br><br>Unadjusted RR mortality:<br>Dem: 2.3 (1.1-4.7)<br>No Dem: 1.5 (0.7-3.1) | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: No<br>MULTI: No<br>COV ASE: No |



SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment                 | Mortality Outcomes   | Quality   |
|----------------------------|--|---|--|--|---|
| 40. Melkas<br>2009<br>#307 | N=451<br>72(7.7)<br>49% female<br><= 6 yrs formal educ.  | Ischemic stroke (intracerebral or<br>subarachnoid haemorrhage<br>excluded)  | Measures: MMSE, Neuropsych-<br>Battery-not specified<br><br>Criteria: DSM-III  | Median survival (95% CI) (unadjusted):<br>Dem: 5.1 years, 4.1 - 6.0<br>No Dem: 8.8 years (7.8 - 9.9)<br>p<0.001  | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: No |
| Design NR<br><br>12 years  | Dem: 48.2%<br>No Dem: 25.1%<br><br>INTL: Helsinki, Finland<br><br>Helsinki Stroke Aging<br>Memory (SAM) cohort | Admitted to Clinic/Program:<br>Patients w/ suspected stroke<br>admitted to university hospital<br><br>Other CC: Stroke, AF, MI, CHF,<br>Arterial HTN, Diabetes, Smoking<br>Peripheral arterial disease<br><br>Self-Report, Records Review:<br>Detailed medical and neurological<br>history. History of CC assessed by<br>reviewing all available hospital charts<br>and conducting a structured interview<br>with the patient and a knowledgeable<br>informant.<br>HTN = SBP >= 160 mm Hg or greater<br>and DBP >= 95 mm Hg. Diabetes<br>included previously documented<br>diagnosis, current use of insulin or oral<br>hypoglycemic medication, or fasting<br>blood glucose greater than 7.0 mmol/l. | 25.5% with dementia<br><br>Clinical mental status examination by<br>the neurologist, including MMSE.<br>Dementia<br>diagnosed using DSM-III. | Median survival brain death (95% CI) (unadjusted):<br>Dem: 6.8 years (4.6 - 8.9)<br>No Dem: 11.1 years (10.2 -11.9)<br>p<0.001<br><br>HR Mortality (95% CI) (adjusted):<br>1.53 (1.15 to 2.04)<br>p=.003 | MULTI: Yes<br>COV ASE: No                                   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality   |
|----------------------------|---|--|--|---|---|
| 41. Oksala<br>2009<br>#170 | N=409<br>71.2(7.7)<br>49.4% female<br>< 6 years educ: 31.8%     | Stroke (acute) (80% first-ever)<br><br>Clinical Exam/Diagnosis, Admitted<br>to Clinic/Program:<br>Detailed medical and<br>neurological history. Stroke<br>severity was assessed using the<br>modified Rankin score at 3 mths.<br>(40% had poor Rankin score) | Measures: MMSE, Stroop<br>colour naming test, WMS-Digit<br>span subtest, WMS-R Logical<br>memory and Visual reproduction<br>subtests, WCST, verbal fluency<br>test, FOME, Token test, the Boston<br>naming test, BDAE, WAIS-R Block<br>design subtest, clock test, and by<br>copying a triangle, a flag, a three-<br>dimensional cube and a Greek cross,<br>Trail making test<br><br>Criteria: DSM-III<br><br>MMSE <=25 in 28.6%<br><br>28.1% with dementia<br><br>Impairment in each cognitive domain<br>was judged using normative, age-<br>specific data from a random healthy<br>Finnish population (2 SD or, if more<br>than one test, 1 SD below the level of<br>the norm if one test). Structured<br>interview with the patient and a<br>knowledgeable informant. Dementia<br>diagnosis using DSM-III criteria. | Unadjusted RR Mortality (95% CI):<br>Dem: 4.4 (3.7 -5.1)<br>No Dem: 9.3 (8.3 -10.4)<br>p < 0.0001<br><br>Adjusted HR Mortality (95% CI) for dementia:<br>Model inc. memory: 1.45 (1.07 to1.96) p=0.017<br>Model inc. language: 1.46 (1.09 to 1.97) p=0.012<br>Model inc. executive: 1.44 (1.06 to 1.94) p=0.019<br><br>Model inc. visuospatial: 1.28 (0.94-1.75) p=.116<br><br>No different findings when recurrent stroke excluded | SAMPLE: No<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #     | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Mortality Outcomes  | Quality   |
|--------------------------------|---|--|---|---|---|
| 42. Tatemichi<br>1994<br>#3044 | N=251<br>>= 60<br>NR% female<br>Other demog NR                  | Acute ischemic stroke<br><br>Clinical Exam/Diagnosis, Admitted<br>to Clinic/Program:<br>Consecutively stroke patients<br>admitted within 30 days of onset to<br>university medical center. Stroke<br>diagnosis was supported by CT<br>scan obtained as part of the<br>clinical evaluation.<br><br>Who made assessment NR | Measures: Neuropsychological<br>Battery-not specified<br><br>Criteria: DSM-III-R<br><br>26% with dementia<br><br>Neuropsychological battery<br>details not available. DSM-III-R<br>criteria to diagnose dementia.<br>BFAS was administered with<br>informants to assess functional<br>impairment. In evaluation of<br>memory deficits in aphasic subjects who<br>were testable, impairment in nonverbal<br>memory was required. | Mortality rate (unadjusted):<br>Dem: 19.8/100 person-years<br>No Dem: 6.9/100 person-years<br>p = NR<br><br>NS differences in causes of death btw dementia and<br>no dementia<br><br>Median Survival (%) (unadjusted):<br>Dem: 38.9(+/- 0.08)<br>No Dem: 74.51(0.04%)<br>p<.001<br><br>Adjusted RR Mortality (95% CI):<br>Model A: 3.11 (1.79-5.41)▯<br>Model B: 3.21 (1.64-6.25)▯<br><br>When the baseline MMSE score was used instead of the<br>diagnosis of dementia, the▯<br>results of the model were similar: RR of 3.99 (95% CI, 1.50<br>to 10.65) for a score of < 12 versus >24. | SAMPLE: No<br>GENDER: NR<br>MCC: No<br>DEM: Yes<br>CC: No<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Mortality Outcomes  | Quality   |
|--|---|--|--|---|---|
| 43. Desmond<br>1998<br>#2760<br><br>Prospective<br><br>Median f/u<br>4.7 years | N=244<br>71.7(8.5)<br>NR% female<br>Mean yrs education:<br>10.1 (4.5)<br><br>U.S.: New York City<br><br>Database NR | Ischemic Stroke<br><br>Admitted to Clinic/Program, Other:<br>Stroke was defined as the acute<br>onset of a focal neurological<br>deficit attributable to cerebro-<br>vascular disease and supported<br>by CT scan (normal or<br>relevant infarct).<br><br>Who made assessment NR | Measures: MMSE, Neuropsych.<br>Battery: MMSE orientation scale,<br>SRT, BVT, BNT, BVRT,<br>repetition and complex ideation<br>subtests of BDAE, other language<br>tests, RDT, WAIS-R similarities<br>subset, MDRS Oddities subtest,<br>Blessed Functional Activity Scale<br><br>Criteria: None<br><br>Comprehensive neurological exam by<br>neurologist specializing in CVD and<br>dementia. | All of the cognitive impairment paradigms were SS<br>predictors of mortality using log-rank tests and Cox<br>proportional hazards analysis. The neuropsychological<br>testing was superior to the use of MMSE and of clinical<br>judgement, particularly when memory impairment was<br>required.<br><br>*This study compared multiple methods for assessing<br>dementia (we are only including those that included<br>memory + 1 other cognitive domain in their assessment).<br>The lowest prevalence (11%) was found for using<br>multiple single-cognitive domain items + functional<br>impairment. The highest prevalence (38%) was found for<br>MMSE < 24. | SAMPLE: No<br>GENDER: NR<br>MCC: No<br>DEM: No<br>CC: No<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Service Utilization Outcomes  | Quality   |
|-----------------------------|--|--|--|---|---|
| 3 studies                   | N = 3423   | Multiple Chronic Conditions  |  | Length of stay-Institution  |   |
| 23. Rothman<br>2008<br>#356 | N=754<br>78.4(5.3)<br>64.6% female<br>Mean (SD) yrs education:<br>12.0 (2.9)<br>90.5% non-Hispanic white<br>39.5% live alone | Inurious Falls, Overall Comorbidity<br><br>Self-Report, Records Review:<br>Participants were asked about<br>overnight hospital stays during the<br>past month and to provide the<br>primary reason for admission.<br>An independent review of hospital<br>records of 44 participants indicated<br>high reliability of self-reported<br>information (kappa value of 0.89).<br><br>An injurious fall was defined as a<br>fall leading to a hospital admission<br>, head injury, or hematoma or<br>bruise of the face or scalp.<br><br>Overall CC: Mean 1.9(1.3) CCs. Self-<br>reported, physician-diagnosed CC's:<br>HTN, MI, CHF, diabetes mellitus,<br>arthritis, hip fracture, stroke, chronic<br>lung disease, cancer (other than<br>minor skin cancers).<br><br>Trained research staff conducted the<br>assessments | Measures: MMSE<24<br><br>Criteria: None<br><br>11.4% with cognitive impairment<br><br>Mean MMSE (SD):<br>CogImp: 21.7(1.5) (min. = 16)<br>No CogImp: NR<br><br>MMSE administered by trained<br>research staff. Other details NR. | Adjusted HR Long-Term Nursing Home Stay (95% CI):<br>Model 1 adjusted for demographics and CC:<br>3.7 (2.5 - 5.4)<br>Model 2 (~Model 1 + 6 other frailty criteria):<br>2.6 (1.7 - 4.0)<br><br>Notes:<br>"Long-term NH stay" = >= 90 days<br><br>Frailty criteria: low physical activity, cognitive<br>impairment, depressive symptoms, weight loss, gait<br>speed, falls.<br><br>In Model 1 and 2, slow gait speed was the strongest predictor of<br>long-term NH stay (out of the 6 frailty criteria)<br><br>The associations for the other frailty criteria were weaker<br>(and of comparable magnitude). | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Prospective<br>cohort study |  |  |  |   |   |
| 7.5 years                   | U.S.: New Haven, CT<br><br>Precipitating Events<br>Project (PEP)<br>(community-dwelling)                                     |  |  |   |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #     | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Service Utilization Outcomes  | Quality  |
|--------------------------------|---|--|---|---|--|
| 24. Magaziner<br>2005<br>#1287 | N=2153<br>Dem: 82.7 (7.3)<br>No Dem: 80.2 (7.7)<br>Female (%)<br>Dem: 69.4<br>No Dem: 72.8<br>White (%)<br>Dem: 76.2<br>No Dem: 84.7%<br>Married (%)<br>Dem: 26.8%<br>No Dem: 21.5%<br>Education <= 8 years (%):<br>Dem: 37.9%<br>No Dem: 28.9% | Overall comorbidity, Fractures or Injuries<br><br>Self-Report, Records Review:<br>Overall comorbidity: # of CC at NH admission: CHD, CHF, cerebrovascular disease, COPD, liver disease, peripheral vascular disease, seizure disorder, peptic ulcers, arthritis, cancer, uncontrolled HTN (SBP >=160 or DBP >=90 mmHg), and malnutrition/underweight (BMI <=20). Information was obtained from interviews with significant others at admission, with the exception that BMI and BP was from NH charts.<br><br>Two other comorbidity measures (modified CCI's and the Diagnostic Cost Group/Hierarchical Coexisting Condition risk adjustor, with the dementia diagnosis omitted from both measures) had the same direction, magnitude, and SS of the effect of dementia on any study outcome so the comorbidity count only is presented.<br><br>Fractures and injuries occurring in the first week of NH stay were included. Study abstractors collected follow-up data from NH home charts. | Criteria: NR<br><br>Diagnosis: DSM-III-R criteria<br><br>48.2% with dementia<br><br>An expert panel of geriatric psychiatrists and neurologists, and a geriatrician determined dementia status using DSM-III-R criteria. 2 panelists gave indep diagnoses: dementia, no dementia, or difficult to diagnose. A larger panel rendered a diagnosis if the two panelists disagreed. Data was obtained from medical records and interviews with staff, family, and residents (conducted by lay interviewers).<br><br>Note: No dementia includes subjects No dementia and those that were difficult to diagnose | Discharged home rate per 100 person-days (unadjusted):<br>Dem: .04, No Dem: .20<br><br>p=NR<br><br>Adjusted RR for Discharged Home (95% CI)<br>0.23 (0.17-0.31)<br><br>p <= 0.001 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Service Utilization Outcomes   | Quality  |
|---|---|--|---|--|--|
| 25. Smith<br>2000<br>#2446  | N=516<br>Dem: 80.8 (7.1)<br>No Dem: 81.6 (6.9)<br>66% female  | Overall comorbidity<br><br>Records Review, Standardized<br>Instrument:<br>Healthcare records from Mayo<br>integrated record-linkage system<br>were reviewed to calculate the<br>Charlson (CCI) scores, excluding<br>dementia | Measures: NR<br><br>Criteria: DSM-III-R, NINCDS-<br>ADRDA (AD)<br><br>42.6% with dementia<br>AD: 164, Other Dementia: 150   | Median length of stay in nursing home (unadjusted):<br>Dem: 946 days (range 46-4473)<br>No Dem: 579 days (range 46-3211)<br>P = NR | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV: Yes |
| Retrospective<br>case control<br><br>6-10 year<br>retrospective<br>record<br>review | % Married:<br>Dem: 46%, No Dem: 38%<br>Mean (SD) education:<br>Dem: 11.7 years (3.6)<br>No Dem: 11.8 years (3.4)<br><br>U.S.: Rochester MN<br><br>Rochester Epidemiology<br>Project (REP) | Who made assessment NR   | First step, search of REP<br>integrated record-linkage system<br>for diagnostic terms that might be<br>related to dementing illness (search<br>continued 6-10 years after 1980-84<br>to capture delayed diagnoses).<br>Individual health care records<br>subsequently reviewed for<br>descriptive features of dementia:<br>including physician's notes,<br>correspondence, lab studies, nursing<br>home notes, etc. Then assessment<br>of probable cause by a neurologist<br>using NINDS-ADRDA criteria.<br><br>Potential controls' records screened<br>similarly for absence of features of<br>dementia. |  |  |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration                         | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes   | Quality  |
|--|--|---|--|---|--|
| 19 studies   | N = 37,446   | Multiple Chronic Conditions   |  |   | SAMPLE: 14<br>GENDER: 18<br>MCC: 19<br>DEM: 9<br>CC: 17<br>MULTI: 11<br>COV ASE: 6         |
| 1. Rothman<br>2008<br>#356<br><br>Prospective<br>cohort study<br><br>7.5 years | N=754<br>78.4(5.3)<br>64.6% female<br>Mean (SD) yrs education:<br>12.0 (2.9)<br>90.5% non-Hispanic white<br>39.5% live alone<br><br>U.S.: New Haven, CT<br><br>Precipitating Events<br>Project (PEP)<br>(community-dwelling) | Inurious Falls, Overall Comorbidity<br><br>Self-Report, Records Review:<br>Self-reported, physician-diagnosed<br>CC's were HTN, MI, CHF, stroke,<br>diabetes mellitus, arthritis, hip<br>fracture, chronic lung disease, and<br>cancer (other than minor skin<br>cancers). There were a mean<br>(SD) of 1.9(1.3) CC.<br><br>Trained research staff conducted<br>the assessments | Measures: MMSE<24<br><br>Criteria: None<br><br>11.4% with cognitive impairment<br><br>Mean MMSE (SD):<br>CogImp: 21.7(1.5) (min. = 16)<br>No CogImp: NR<br><br>MMSE administered by trained<br>research staff. Other details NR. | Adjusted HR Chronic Disability (95% CI):<br>Model 1 adjusted for demographics and CC:<br>2.1 (1.7 - 2.8)<br>Model 2 (~Model 1 + 6 other frailty criteria):<br>1.8 (1.4 - 2.4)<br><br>In Model 1 and 2, slow gait speed was the strongest<br>predictor of chronic disability (out of the 6 frailty criteria:<br>low physical activity, cognitive impairment, depressive<br>symptoms, weight loss, gait speed, and falls).<br><br>The associations for the other frailty criteria were<br>weaker (and of comparable magnitude). | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

How function was assessed:

"Chronic disability" = new ADL disability that persists  
for at least 3 consecutive months



SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                          | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                                    | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment | Function Outcomes  | Quality   |
|---|--|--|--|--|---|
| 2. Nikolova<br>2011<br>#29                          | N=1164<br>82(7.3)<br>>50% female<br>>50% had >= h.s. educ  | Overall comorbidity<br><br>Standardized Instrument:<br>A 16-item questionnaire on<br>prevalence of common CC.<br>Number of diseases was coded<br>as: 0-1 disease, 2-3, 4-5, and<br>more than 6 diseases. | Measures: SPMSQ >= 5<br><br>Criteria: None<br><br>30% with cognitive impairment<br><br>Assessment details NR                 | Adjusted OR IADL disability (95% CI):<br>2.64 (0.47;14.90)<br>p=NS<br><br>Adjusted OR ADL disability (95% CI):<br>7.10 (1.12;44.85)<br>p < 0.05  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Secondary<br>analysis of<br>SIPA RCT<br><br>3 years | INTL: Montreal, Canada<br><br>SIPA (Research Program<br>on Integrated Services for<br>the Elderly) | Trained Interviewer  |  | <u>How function was assessed:</u> Data on physical functional limitations were collected using a 7-item scale based on a portion of Roslow and Breslau's Functional Health Scale and some measures of physical performance adapted from Nagy's work on disability. Total score varied from 7 to 28 points with higher scores denoting worse function. The activities referred to were climbing stairs, walking outdoors for 1.5 km, carrying an object of 5 kg, bending and kneeling, pulling or pushing large objects like a chair, handling or picking up small objects, raising arms above shoulders. A cut-off for severity of functional limitations was set as four or more limitations, based on Roslow and Breslau (1966). |   |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration            | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes   | Quality  |
|---|--|---|---|---|--|
| 3. Millan-<br>Calenti<br>2010<br>#A28<br>Design NR<br>Duration NR | N=579<br>75.1 (7.5)<br>56.3% female<br>No formal education:<br>CogImp: 87.4%<br>No CogImp: 83.2%<br><br>Older subjects (85+) were<br>more cognitively impaired<br>(29.4%) and presented more<br>co-occurring CogImp and<br>depression (22.1%) than<br>other ages.<br><br>INTL: Naron Council, A<br>Coruna, Spain<br><br>Database NR (community-<br>dwelling) | Overall comorbidity, Visual /hearing<br>impairments<br><br>Self-Report, Standardized<br>Instrument, Clinical Exam/ Diagnosis,<br>Records Review:<br><br>Medical histories were collected by a<br>physician or a trained nurse in charge<br>of the participant during the research.<br>Participants report was given by the<br>patient or their relatives according to<br>medical records. Comorbidity<br>conditions were defined according to<br>the CCI. | Measures: MMSE<br><br>Criteria: None<br>20.6% with cognitive impairment<br><br>Mean (SD) MMSE: 23.9 (5.6)<br>(all subjects)<br><br>MMSE cut-offs based on pop-based<br>age-education adjusted norms: <20<br>for 80+ w/ 0 -4 yrs of education to<br><29 for age 65-69 college-educated.<br>Majority of subjects (>80%) had no<br>formal education. MMSE<br>administered by a psychologist. | IADL dependence (%)<br>Cog Imp: 75.3%<br>No CogImp: 40.8%<br>p < 0.001<br><br>ADL dependence (%)<br>CogImp: 42.5%<br>No CogImp: 21.7%<br>p < 0.001<br><br><u>How function was assessed:</u> Functional status was<br>measured using the ADL (Katz et al., 1963) and IADL<br>(Lawton and Brody, 1969) scores. Participants were asked<br>by a physician or a trained nurse if they had any difficulty<br>performing each task without the help of another person.<br>Individuals who were unable to perform any one of the<br>activities were considered to be functionally incapacitated<br>in that activity (ADL or IADL dependent). | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                            | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes   | Quality  |
|----------------------------|--|---|---|---|--|
| 4. Zekry<br>2009<br>#463   | N=435<br>85.3(6.7)<br>74% female   | Overall comorbidity, Obesity<br><br>Standardized Instrument:<br>Overall comorbidity: The same geriatrician calculated CCI for each patient using the patient's medical records. Higher scores = greater comorbidity. Dementia was not included in the calculation of CCI or comorbidity. Obesity = BMI, using the short version MNA administered on admission, by the same nurse. | Measures: MMSE, Short Cognitive Evaluation Battery, CDR<br><br>Criteria: DSM-IV-TR, NINCDS-ADRDA, NINDS-AIREN<br><br>44% with dementia<br>AD: 77, VD: 21, Mixed: 82, Other: 11  | Premorbid ADL: Mean (SD)<br>Dem: 4.51 (1.31)<br>Non-Dem: 5.27 (0.86)<br>p< 0.001<br><br>Premorbid IADL: Mean (SD)<br>Dem: 3.47 (2.26)<br>Non-Dem: 5.33 (2.01)<br>p< 0.001<br><br><u>How function was assessed:</u> ADL and IADL (Katz et al., 1963; Lawton and Brody, 1969) scores were determined as a function of patient status 2 weeks before admission, based on the patient's medical history or information supplied by an informal or formal carer. ADL assesses ability to manage activities such as bathing, dressing, going to the toilet, continence, feeding, and transfer (6 items). IADL assesses ability to use the telephone, to shop, to use transport, to cook, to do housework, to take medication and to handle finances (8 items). For both scales, 0 indicates total dependence and the maximum score (6 or 8) indicates total independence. | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Prospective cohort         | Dem: 55.6%<br>No Dem: 59.9%<br>Acute and rehab patients in geriatric hospital              |   |   |   |  |
| Duration NR                | INTL: Geneva, Switzerland<br><br>Database NR (acute and rehabilitation geriatric hospital) |   | The same neuropsychologist assessed all subjects for clinical dementia, at least one week after admission, to avoid the effects of concomitant delirium. Based on screening results with the objective measures, the same neuropsychologist carried out a comprehensive standardized neuropsychological assessment, to determine the etiology and severity of clinical dementia. Dementia severity was assessed with the CDR (2, moderate, and 3, severe dementia). DSM IV-TR was used for dementia diagnosis, NINCDS-ADRDA and ADDTC for AD, and NINDS-AIREN for VaD. Cerebral imaging was also carried out. |   |  |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                                      | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes  | Quality   |
|---|--|--|---|--|---|
| 5. Lyketsos<br>2007<br>#805                                     | N=198<br>CogImp: 86.1(6.73)<br>No CogImp: 84.9 (10.76)<br>% Female   | Overall comorbidity, Depression,<br>Polypharmacy/High risk medications   | Measures: MMSE, Trails A andB,<br>HVLT<br><br>Criteria: None  | Mean (SD) PGDRS scores:<br>CogImp: 14.24 (8.81)<br>No CogImp: 8.42 (6.12)<br>p<.001  | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Prospective<br>cohort study<br>4 years<br>(median 1.5<br>years) | CogImp: 80.6%<br>No CogImp: 75%<br>% Caucasian<br>CogImp: 79.9%<br>No CogImp: 89.1%<br>p=NS<br><br>% in large facility<br>CogImp: 70.9%<br>No CogImp: 85.9%<br>p=.02 | Self-Report, Standardized Interview,<br>Records Review:<br>Residents completed the GMHR<br>(overall comorbidity) and CSDD<br>(depression). Number of Medications<br>came from chart reviews and<br>interviews with the<br>resident, family informant, and facility<br>staff members who knew the resident<br>well. | 68% with cognitive impairment<br><br>Mean MMSE (p<.001)<br>CogImp: 14.64 (7.67)<br>No CogImp: 25.84 (5.50)<br><br>Mean HVLT: (p<.001)<br>CogImp: 0.54 (1.12),<br>No CogImp: 4.82 (3.32)<br><br>Mean Trails B (p <.001)<br>CogImp: 500.84 (157.74)<br>No CogImp: 270.17 (151.96)   | Mean (SD) Get up and Go scores:<br>CogImp: 39.44 (57.48)<br>no CogImp: 29.66 (18.76)<br>p = 0.13<br><br>High activities, hours ( 75th percentile) %:<br>CogImp: 23.5<br>No CogImp: 38.1%<br>p = 0.03<br><br>Mean (SD) total activity hours:<br>CogImp: 69.45 (62.56)<br>No CogImp: 91.29 (74.89)<br>p = 0.03   |   |
|   | U.S.: Central Maryland<br><br>The Maryland Assisted<br>Living Study (MD-AL)  |  | Information from MMSE, Trails, and<br>HVLT (no cut-offs reported) was<br>brought to the panel (the team that<br>evaluated the resident in his or her<br>facility, another geriatric psychiatrist,<br>a geriatric medicine physician, a<br>neuropsychologist, and a registered<br>nurse). The panel used a consensus<br>process to make diagnoses and<br>render opinions about the<br>completeness of evaluation and<br>treatment for dementia on the basis<br>community-care standards (e.g.,<br>minimize inappropriate medication<br>use). | <u>How function was assessed:</u><br>-Psychogeriatric Dependency Rating Scale (PGDRS) is a<br>measure of ADL impairment; higher scores indicate greater<br>ADL dependency.<br>-Get Up and Go is a direct assessment of physical<br>functioning, measuring the time it takes to rise from a chair<br>and walk 5 m, assessing gait speed and balance. Higher<br>scores (longer time) indicates greater impairment.<br>-Caregiver ratings of hours of activity participation in the<br>previous month based on self-report and available records<br>kept at the facility. |   |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes  | Quality  |
|--|---|---|--|--|--|
| 6. Magaziner<br>2005<br>#1287<br>Epidemiological<br>cohort<br>Up to 2 years<br>or until<br>discharge | N=2153<br>Dem: 82.7 (7.3)<br>No Dem: 80.2 (7.7)<br>Female (%)<br>Dem: 69.4<br>No Dem: 72.8<br>White (%)<br>Dem: 76.2<br>No Dem: 84.7<br>Married (%)<br>Dem: 26.8<br>No Dem: 21.5<br>Education <= 8 years (%)<br>Dem: 37.9%<br>No Dem: 28.9%<br><br>U.S.: Maryland (nursing<br>homes)<br><br>Database NR | Overall comorbidity<br><br>Self-Report, Records Review:<br>Overall comorbidity: # of CC at NH<br>admission inc. CHD, CHF,<br>cerebrovascular disease, COPD, liver<br>disease, peripheral vascular disease,<br>seizure disorder, peptic ulcers,<br>arthritis, cancer, uncontrolled HTN<br>(SBP >=160 or DBP >=90 mmHg),<br>and malnutrition/ underweight (BMI<br><=20). Information on CC's was<br>obtained from interviews with<br>significant others at NH admission,<br>with BMI and HTN information<br>abstracted from NH charts. Two other<br>comorbidity measures (modified<br>CCI's and Diagnostic Cost Group/<br>Hierarchical Coexisting Condition risk<br>adjustor, with dementia omitted from<br>both measures) had the same<br>direction, magnitude, and SS of the<br>effect of dementia on any study<br>outcome so the comorbidity count<br>only is presented. | Criteria: NR<br><br>Diagnosis: DSM-III-R criteria<br><br>48.2% with dementia<br><br>An expert panel of geriatric<br>psychiatrists and neurologists, and a<br>geriatrician determined dementia<br>status using DSM-III-R criteria. 2<br>panelists gave indep diagnoses:<br>dementia, no dementia, or difficult to<br>diagnose. A larger panel rendered a<br>diagnosis if the two panelists<br>disagreed. Data was obtained from<br>medical records and interviews with<br>staff, family, and residents<br>(conducted by lay interviewers).<br><br>Note: No dementia includes subjects<br>No dementia and those that were<br>difficult to diagnose | Mean (SD) number of ADL dependencies:<br>Dem: 4.3(1.7)<br>No Dem: 3.2(2.1)<br>p < .01<br><br>>= 4 ADL dependencies:<br>Dem: 73.1<br>No Dem: 50.1<br>p < .01<br><br>% Bedbound:<br>Dem: 5.5<br>No Dem: 10.2<br>p < .01<br><br>% Chairbound:<br>Dem: 36.8<br>No Dem: 38.5<br>p=NS (0.41)<br><br><u>How function was assessed:</u> Baseline physical function<br>was assessed using a modified Katz ADL scale, which<br>measures ability in bathing, dressing, toileting, transferring,<br>feeding, and continence. Each ADL domain was scored as<br>dependent or fully independent; a summary measure<br>indicated the number of domains in which residents were<br>dependent. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes  | Quality  |
|----------------------------|---|--|--|--|--|
| 7. Stewart<br>2009<br>#185 | N=1890<br>Ages 77 - 96<br>0% female                             | Diabetes, HTN, Depression,<br>Smoking<br><br>32% with diabetes, 46% w/ HTN,<br>58% past smoker   | Measures: NR ("Cognitive<br>prescreening and neuropsych-<br>ological testing")<br><br>Criteria: DSM-III (R), NINCDS<br>-ADRDA, ADDTC, Hachinski<br>Ischemic Scale<br>59% with dementia<br>AD: 74 ,VD: 15, Other: 25  | % with impaired physical function:<br>Incident Dementia: 9%<br>Incident AD: 4%<br>Incident VaD: 13%<br>No Dementia: 2%       | SAMPLE: Yes<br>GENDER: No<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV: No |
| Longitudinal               | U.S.: Oahu, HI  | Self-Report, Clinical Exam/<br>Diagnosis:<br>Diabetes was assessed at the<br>fourth clinical examination using<br>World Health Organization criteria.<br>BP was measured with the same<br>standardized protocol at each<br>examination. After the participant had<br>been seated for 10 minutes, SBP and<br>DBPs were measured on 3 occasions<br>5 minutes apart on the left arm of a<br>seated participant using a mercury<br>sphygmomanometer with a standard<br>cuff. Diastolic pressure was recorded<br>as the fifth phase. Repeated readings<br>were averaged for each examination.<br>Depressive symptoms were<br>measured using the CES-D. Smoking<br>status was ascertained from previous<br>exams from the study. | The 3-stage procedure included a<br>cognitive prescreening,<br>neuropsychological testing, proxy<br>interview, neurological examination,<br>and neuroimaging. Consensus<br>diagnoses were made by a<br>neurologist and 2 physicians.<br>Diagnosis was made according to<br>DSM-III-R, with NINCDS-ADRDA<br>(AD) and ADDTC (VaD). Probable<br>VaD diagnosis required dementia,<br>computed tomography/MRI evidence<br>of 1 infarct outside of the cerebellum,<br>and then either clinical/imaging<br>evidence of 2 ischemic strokes or a<br>single stroke with a clear temporal<br>relationship to the onset of dementia.<br>Additional support is allowed if there<br>is evidence of multiple infarcts in<br>brain regions known to affect<br>cognition, multiple transient ischemic<br>attacks, a history of vascular risk<br>factors, and an elevated Hachinski<br>Ischemic Scale score. | <u>How function was assessed:</u> Impaired physical<br>function = inability to rise from a chair or walking<br>speed 0.4 m/s |  |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                               | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                                   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes  | Quality   |
|--|---|---|--|--|---|
| 8. Eriksson<br>2008<br>#808<br><br>NR<br><br>6-month f/u | N=186<br>83.6(6.6)<br>72.6% female<br>Other Demog NR<br><br>Intl: Umea, Sweden<br><br>Database NR | Stroke (32%) , Heart disease,<br>Depression (28%), Falls (53%),<br>Urinary Incontinence (29%), Visual<br>and hearing impairments, Sleeping<br>disorder, Diabetes<br><br>Records Review, Other:<br>6 specialists in geriatric medicine,<br>who were also the resident's ordinary<br>physicians, registered the diagnoses<br>and current medication of each<br>participant based on previous<br>knowledge and chart review. The<br>staff was instructed to register every<br>fall they observed on a structured fall<br>incidence report, or when they found<br>residents unaccountably on the floor<br>or ground. A fall was defined as an<br>event in which the resident<br>unintentionally came to rest on the<br>floor, regardless of whether or not an<br>injury was sustained. This included<br>falls as a consequence of acute<br>illness (e.g., stroke, an epileptic<br>seizure). The fall reports were<br>collected once a week by the<br>research team. In order to optimize<br>the collection of fall reports, the<br>resident' charts were reviewed after 3<br>months and at the end of the study.<br>The staff was obliged to note down<br>important events in the charts, such<br>as a fall or a change in a resident's<br>health status. | Measures: MMSE < 24<br><br>Criteria: DSM-IV<br><br>55.4% with dementia<br><br>Dementia was one of the diagnoses<br>registered by the six specialists<br>described in the CC assessment<br>section. After the study was<br>terminated, the diagnoses of<br>dementia were re-evaluated,<br>according to the DSM IV criteria,<br>through chart review and a review of<br>the baseline assessments by a<br>specialist in geriatric medicine. All<br>participants with an MMSE < 24<br>(administered by a physiotherapist)<br>with no diagnosis of dementia were<br>included in the re-evaluation (N =<br>82). An explanation for the low<br>MMSE score was explored, which<br>could be due to different diagnosis or<br>difficulties originating from these<br>diagnoses (e.g., depression, stroke,<br>impaired vision, paresis affecting<br>writing, and aphasia these<br>diagnoses). If no alternative<br>explanation regarding the low MMSE<br>score was found, the findings still<br>had to fit the symptom profile of<br>dementia, including memory deficit<br>and social dependency, to be<br>diagnosed with dementia. | Mean (SD) Barthell ADL:<br>Dem: 11.1 (5.7) , No Dem: 14.5 (4.8), p = NS<br><br># (%) unable to walk:<br>Dem: 14 (13.6) , No Dem: 16 (19.3), p = NS<br><br><u>How function was assessed:</u> Nurses' aides and licensed<br>practical nurses who knew the resident were questioned by<br>physiotherapists about ADLs using the Barthell Index. The<br>physiotherapists also assessed walking with and without an<br>aid. | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #     | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes  | Quality   |
|--------------------------------|---|---|---|--|---|
| 9. Lyketsos<br>2005<br>#1365   | N=695<br>Dem: 83.89 (6.29)<br>No Dem: 79.93 (6.73)<br>% female  | Overall comorbidity, Stroke, HTN,<br>Diabetes, Chronic Pain, Arthritis,<br>GI disease, Thyroid disease,<br>Heart Attack, Stroke, High<br>cholesterol  | Measures: MMSE, IQCODE<br><br>Criteria: DSM-IV<br><br>53.8% with dementia   | Mean DSRS-ADL score, by GMHR rating<br>GMHR = Fair:<br>Dem: ~12 (10.5 - 14), No Dem: ~2 (1 - 3.5)<br>GMHR = Good:<br>Dem: ~9 (7.5 - 11.5), No Dem: ~1 (0.5-1.5)<br>GMHR = Excellent<br>Dem: ~4.5 (3 - 6), No Dem: ~0.2 (0.1 - 0.3)   | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Nested, case-<br>control study | Dem: 64, No Dem: 54.8<br>99% White  | Mean number of CC's:<br>Dem: 4.1 (2.5), No Dem: 3.7 (2.3)   | Mean (SD) MMSE:<br>Dem: 20.47 (5.92)<br>No Dem: 28.30 (1.86)  | Data are estimated from Figure 2A. p values are not reported. Less serious comorbidity (higher ratings on GMHR) was associated with less impairment (lower mean DSRS- ADL).  |   |
| Cross-sectional                | Mean (SD) yrs education:<br>Dem: 12.95 (3.13)<br>No Dem: 13.58 (2.90)<br>% married:<br>Dem: 43, No Dem: 63.2<br><br>U.S: Cache County, Utah<br><br>Cache County Study | Self-Report, Standardized<br>Instrument, Records Review:<br>A detailed review of systems was used to identify each participant's medical illnesses, with follow-up questions to clarify diagnoses and treatments. The GMHR was administered to rate seriousness of non-cognitive medical comorbidity in persons with cognitive disorders. It was developed as a clinician rating. Ratings of 4 indicate little-to-no comorbidity; 3: mild-to-moderate comorbidity; 2: moderate-to-severe comorbidity; and 1: serious comorbidity. In this study, GMHR ratings were assigned by a geriatric psychiatrist on the basis of direct and proxy interviews by the nurse, as well as a brief physical and neurological exam. To assess the reliability of these ratings, we calculated the agreement between two raters in a random sample of 150 cases and found it to be high (p < 0.001) | The subsample and all other participants with suspected dementia underwent a comprehensive clinical assessment for dementia in their place of residence by a research nurse, a psychometric technician, and a geriatric psychiatrist. Data from these evaluations were used to classify participants at consensus conferences that included two geriatric psychiatrists, a board-certified neurologist, a senior neuropsychologist, and a cognitive neuroscientist. | <u>How function was assessed:</u> A knowledgeable informant rated each participant on the Dementia Severity Rating Scale (DSRS), an 11-item scale of signs and symptoms associated with dementia. Six of the 11 items refer to ADLs, including: engagement in social activities, household responsibilities, personal care, meals/feeding, incontinence, and mobility. Each ADL is assessed on a scale from 0 to 4 (except mobility, which is assessed on a scale from 0 to 6). A rating of "0" indicates that there is no impairment, and the highest rating, of "4" (or "6") indicates complete dependency or loss of ability to perform the ADL. The sum of the six ADL ratings (DSRS-ADL) was used as an indicator of cumulative ADL impairment. |   |



SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #         | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes   | Quality  |
|------------------------------------|--|--|---|---|--|
| 10. Zekry<br>2008<br>#711          | N=349<br>85.2(6.7)<br>76% female<br>% < 12 yrs education:<br>Dem: 57, No Dem: 63.1<br>% live alone:<br>Dem: 50, No Dem: 65 | Overall comorbidity, Stroke, COPD,<br>Cancer, AF, CHF, HTN, Arthritis,<br>Cerebrovascular disease,<br>Substance use (drinking, smoking),<br>BMI, Nutrition<br><br>Mean (SD) CCI:<br>Dem: 4.87 (2.56)<br>No Dem: 4.50 (2.79)  | Measures: MMSE, Neuropsychological<br>Battery - Not Specified, Short<br>Cognitive Evaluation<br><br>Criteria: "Formal clinical criteria"<br><br>53.8% with cognitive impairment<br><br>AD: 61 ,VD: 17, Mixed: 62, Other: 11<br>(DLB: 3, PDD: 2, Creutzfeld-Jacob<br>disease: 1, cortico-basal: 1, fronto-<br>temporal: 1, hydrocephaly with<br>normal pressure: 1, glioblastoma: 1,<br>cerebral measures: 1)<br><br>Mean (SD) MMSE Scores:<br>AD: 16.3 (4.7)<br>Mixed dementia: 15.6 (4.9)<br>VaD: 17.5 (6.7)   | Mean (SD) Functional Independence Measure (FIM):<br>(Assessment at discharge)<br>Dem: 84.87 (27.88)<br>No Dem: 99.56 (28.81)<br>p < 0.0001<br><br><u>How function was assessed:</u> The Functional Independence<br>Measure (FIM). FIM scores range from 18 (completely<br>dependent) to 126 (completely independent). | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Prospective<br>study<br><br>1 year | Location<br>DB   | Self-Report, Records Review:<br>The Charlson (CCI) was determined<br>by extensive review of the patient's<br>medical records for diagnoses<br>established at/or before enrollment in<br>this study, higher scores indicating<br>greater comorbidity. The study also<br>included HTN, AF, stroke, and<br>hypercholesterolaemia (CC's not<br>included in the CCI). The various<br>classes of medication taken before<br>admission were also listed, as well as<br>functional status. ADL and IADL were<br>determined by the same nursing<br>team on the admission day of the<br>patient. The information regarding the<br>previous 2 weeks was supplied by<br>the patient when he was capable of<br>answering and by an informal and/or<br>formal caregiver.<br><br>Who made assessment NR | The same neuropsychologist<br>assessed all subjects at least one<br>week after patient inclusion. The<br>following neuropsychological battery<br>was applied: the MMSE and the<br>short cognitive evaluation. Based on<br>this screening, a comprehensive<br>standardised neuropsychological<br>battery used in our routine clinical<br>practice was carried out by the same<br>neuropsychologist, with formal<br>clinical criteria used to determine the<br>aetiology and severity of clinical<br>dementia (Appendix 3). Cerebral<br>imaging was also carried out. |   |  |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                              | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment | Function Outcomes   | Quality   |
|----------------------------|--|---|--|---|---|
| 11. Feng<br>2010<br>#A441  | N=146<br>77.8(10.8)<br>69.9% female<br>Other Demog NR  | Overall comorbidity<br><br>Mean (SD) CCI: 2.1(1.9)  | Measures: MMSE < 24<br><br>Criteria: None  | Mean (SD) modified Barthel Index<br>Baseline:<br>CogImp: 50.3 (16.6), No CogImp: 57.3 (21.2)<br>Discharge:<br>CogImp: 74.6 (16.5), No CogImp: 83.8 (15.0)<br>12-month:<br>CogImp: 87.2 (27.9), No CogImp: 94.4 (7.4)  | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: No<br>MULTI: Yes<br>COV: No |
| Cohort study<br><br>1 year | INTL: Singapore<br><br>Database NR (hip fracture patients in a rehabilitative care facility) | Standardized Instrument, Records Review:<br>Overall comorbidity was obtained from medical charts and included diabetes, HTN, stroke, dementia, congestive cardiac failure, ischemic heart disease, COPD and arthritis. A modified Charlson comorbidity scale was created that excluded dementia and HTN as disease conditions.<br><br>Nurses reviewed charts and completed the CCI. | 79% with cognitive impairment<br><br>Nurses administered the MMSE  | Mean (SD) Ambulatory status<br>Baseline:<br>CogImp: 2.3 (0.6), No CogImp: 2.7 (0.7)<br>Discharge:<br>CogImp: 2.8(0.8), No CogImp: 3.6(1.2)<br>12-month:<br>CogImp: 4.7 (1.1), No CogImp: 4.1 (1.3)<br><br>p = NR  |   |
|                            |  |   |  | <u>How function was assessed:</u> The Modified Barthel Index (MBI) is a 10-item questionnaire used to assess independence in ADLs. A summary score was computed by adding the point of each single item. Possible scores range from 0 to 100, with a higher score indicating greater independence in ADLs. Ambulatory status was determined based on functional walking categories defined by Perry et al. (1995). It includes six walking categories, each of which was characterized according to observation of the participants' ambulation, including the walkingspeed test. |   |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #      | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes  | Quality   |
|---------------------------------|--|---|---|--|---|
| 12. Cankurtaran<br>2008<br>#779 | N=1436<br>72.7(6.9)<br>65.8% female<br>Living status (%)<br>Alone: Dem 51.7%,<br>No Dem 54.1%  | Stroke, CVD, HTN, Diabetes,<br>Depression, Lipidemia,<br>Cerebrovascular disease,<br>Substance use (drinking, smoking)  | Measures: MMSE, CDT<br><br>Criteria: DSM-IV, NINCDS<br>-ADRDA, Hachinski Ischemic<br>Score for VaD  | Mean (SD) IADL score:<br>AD: 13.5 (3.5)<br>No Dem: 14.3 (2.9)<br>p=0.001<br><br>ADL score data was not reported.                           | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Design NR<br>Cross-sectional    | Living With family:<br>AD 46.3%, VaD 47.9%, no<br>Dem 41.3%<br>Nursing Home: AD 2.0%,<br>VaD 4.3%, no CI 4.7%<br><br>Intl: Ankara, Turkey<br><br>Database NR | Clinical Exam/Diagnosis,<br>Lab exams:<br>A comprehensive geriatric<br>assessment (CGA) was<br>conducted. "Normal": vitamin B12<br>160 pg/ml and over; total cholesterol<br>200 mg/dl and lower; triglycerides<br>200 mg/dl and lower; LDL-C 130<br>mg/dl and lower, HDL-C 40 mg/dl<br>and over | 21% with dementia<br>AD: 203 ,VD: 73<br><br>Global Deterioration Scale (GLDS)<br>was used to stage severity of<br>dementia:<br>60% mild (GLDS stage 3), 30%<br>moderate (stage 4), and 10% severe<br>(stage 6). | <u>How function was assessed:</u> The Instrumental ADL Scale<br>(Lawton and Broody, 1969) and the ADL<br>Scale (Mahoney and Barthel, 1965) |   |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration               | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes   | Quality  |
|--|--|--|--|---|--|
| 13. Gombojav<br>2011<br>#A5<br><br>Longitudinal<br><br>Mean 11.8 yrs | N=2496<br>73.6(5.9)<br>57.7% female<br>% no formal education:<br>CogImp: 80%<br>No CogImp: 50%<br><br>INTL: South Korea<br><br>Database NR | 1/3 with a chronic disease, 1/3 with<br>HTN (may be overlap btw these<br>2 categories), BMI, Smoking,<br>Drinking,<br><br>Clinical Exam/Diagnosis, Self-<br>Report:<br>Avg 2 BP measurement using<br>standard mercury sphygmo-<br>manometer, assessed by<br>trained interviewer. SBP/DBP:<br><= 140/90 mm Hg.<br>Overall comorbidity: Subjects<br>answered yes or no to the<br>question "do you have any<br>chronic disease or past accident<br>or injury due to which you feel<br>uncomfortable in your daily life<br>including work? | Measures: MMSE<br><br>Criteria: None<br><br>44% with CogImp<br><br>Mean MMSE (SD):<br>mild severity CogImp: 17.6 (1.1)<br>severe CogImp: 11.9 (3.1)<br>no CogImp: 24.1 (2.9)<br><br>MMSE administered by the<br>investigation team | % with ADL disability:<br>No CogImp: 0.8<br>Mild Severity CogImp: 1.6<br>Severe CogImp: 7.4<br>p = 0.000<br><br>% w/ IADL disability:<br>No CogImp: 49.7<br>Mild Severity CogImp: 63.7<br>Severe CogImp: 73.4<br>p = 0.000<br><br><u>How function was assessed:</u> ADLs included:<br>bathing or showering, dressing, going to the toilet,<br>transferring (in and out of bed or chair), and eating.<br>A 3-category outcome score was used to assess<br>disability in ADL: (1) can perform the activity independently,<br>(2) can perform the activity with assistance, and (3) unable<br>to perform the activity. ADL disability was defined as being<br>unable to perform the activity. In order to define IADL, the<br>following activities were addressed: housework, meal<br>preparation, traveling by car or public transportation,<br>shopping food or clothes (regardless of transport),<br>medication use (preparing and taking correct dose),<br>making telephone calls and managing money. Each activity<br>was graded on a 3-part scale: independent, assistance<br>needed and dependent. Participants who performed the<br>activities dependently or with assistance were considered<br>as having an IADL disability. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration                        | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes  | Quality   |
|---|--|--|---|--|---|
| 14. Covinsky<br>2003<br>#1965<br><br>Retrospective<br>analysis<br><br>2 years | N=917<br>Age at death:<br>CogImp: (85.3 (7.7)<br>No CogImp: 80.9 (9.6)<br>% Female (CogImp/No):<br>70.7%, 65.9%<br>% Race/Ethnicity Cog<br>Imp/No Cog Imp:<br>White: 41.7 / 53.9<br>African Americ: 21.3 / 14.7<br>Hispanic: 4.5 / 9.6<br>Asian: 31.4 / 20.7<br>Other: 1.2 / 1.2<br><br>U.S.: 12 PACE<br>demonstration centers in the<br>U.S.<br><br>DataPACE<br><br>PACE cares for frail older<br>people who meet criteria for<br>nursing home placement,<br>with the goal of keeping the<br>patient at home. | Parkinson's Disease (12%),<br>Cancer (19%), Diabetes (27%),<br>Stroke (43%), CHF (37%),<br>CAD (46%)<br><br>Records Review:<br>DataPACE includes data about<br>demographics, functional status, and<br>comorbid conditions for patients<br>enrolled at the PACE demonstration<br>sites. Sources of data include<br>patients, caregivers, nurses, social<br>workers, and physicians. Principles<br>underlying PACE data collection<br>include a consistent set of variables<br>collected by all sites, consistent<br>guidelines for recording data across<br>sites, and centralized training and<br>quality assurance procedures.<br>Detailed procedure manuals outlined<br>specific definitions and protocols for<br>each data element, and data<br>collection staff were trained to a<br>standard of reliability. Staff from the<br>coordinating center visited each<br>PACE site on a yearly basis to<br>monitor data collection and perform<br>additional reliability checks. | Measures: SPMSQ<br><br>Criteria: None<br><br>63.6% with cognitive impairment<br><br>Mean (SD) SPMSQ errors:<br>CogImp: 8.5 (1.5)<br>No CogImp: 1.9 (1.7)<br><br>Cognitive status was defined as six<br>or more errors on the 10-item<br>SPMSQ. PACE nurses or social<br>workers assessed SPMSQ scores on<br>enrollment and every 3 months<br>thereafter. For these analyses, the<br>SPMSQ score that was closest to 24<br>months before death was used. | Unadjusted OR for Incidence of Functional Decline<br>(95% CI) for with vs without CogImp:<br>Bathing: OR: 3.4 (2.3-5.2)<br>Eating: OR: 2.7 (1.9-3.9)<br>Mobility: OR: 2.2 (1.6-3.2)<br>Continence: OR: 3.1 (2.2-4.6)<br>Mean (SD) Occurrence of Decline, Days Before Death:<br>Bathing:<br>CogImp: 338 (332) / No Cog Imp: 194 (175), p=.83<br>Eating:<br>CogImp: 325 (222) / No CogImp: 173 (174), p<.001<br>Mobility:<br>CogImp: 331 (281) / No CogImp: 187 (186), p=.05<br>Continence:<br>CogImp: 337 (270) / No CogImp: 191 (175), p=.016<br><br>Adjusted OR for Functional Decline (95% CI):<br>Bathing: OR: 3.6 (2.3-5.8)<br>Eating: OR: 2.6 (1.7-3.8)<br>Mobility: OR: 2.4 (1.6-3.7)<br>Continence: OR: 3.5 (2.3-5.3)<br><br>For each measure, patients with cogimp were much more<br>likely than patients without cogimp to be fully dependent 2<br>years before death. People with cogimp were also more<br>likely to have the maximal level of dependence in the 0- to<br>3-month window before death. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author       | # Sample (N)       | Chronic Condition (CC): | Cognitive Impairment (CogImp)/Dementia (Dem): | Function Outcomes | Quality |
|--------------|--------------------|-------------------------|---|-------------------|---------|
| Yr           | Mean age (SD)      | Definition              |   |                   |         |
| Tracking #   | % Female           | Assessment              | Measures, Criteria, % with                    |                   |         |
| Study Design | Other Demographics | Who made assessment     | Dem/CogImp, Mean Scores, Assessment           |                   |         |
| Duration     | Location           |                         |   |                   |         |
|              | DB                 |                         |   |                   |         |

Covinsky (cont)

How function was assessed: PACE nurses assessed each functional measure at the time of PACE enrollment and every 3 months thereafter. For bathing, eating, and mobility (walking), patients were classified as independent (able to do the activity without the assistance of another person all of the time), partially dependent (needs help for part of the activity some or all of the time), or fully dependent (needs help for all of the activity all of the time). For mobility, patients who used an assistive device other than a wheelchair were classified as independent as long as they did not need the help of another person. Continence was defined based on the combination of bladder and bowel incontinence. For each, incontinence was defined as one or more episodes of incontinence per week. A hierarchical classification was used. Patients were defined as continent, bladder incontinent, or bowel incontinent. Bowel incontinence was classified as a higher level of functional impairment because, in most patients, bladder incontinence proceeds bowel incontinence and because bowel incontinence generally confers greater caregiving needs than bladder incontinence.

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes   | Quality   |
|----------------------------|--|--|--|---|---|
| 15. Sousa<br>2009<br>#31   | N=14,981<br>71.3 - 75.2 among 11 sites<br>53 - 66% female<br>% with educ attainment<br>ranged from 34% - 97% | Median (range) prevalence:<br>HTN: 62.6% (28.5 - 75.4)<br>Diabetes: 14.0% (1.0 - 24.5)<br>Stroke: 7.1% ( 1.1 - 8.7)<br>COPD: 5.8% (1.6 - 7.6)<br>Depression : 4.7% ( 0.3 - 13.8)<br>Heart Disease: 4.4% ( 1.2 - 14.2)<br>Arthritis/Rheumatism: 18.2% (1.9-51.1)<br><br>Self-Report, Standardized<br>Instrument, Clinical Exam/Diagnosis:<br>Depression: ICD-10 criteria<br>ascertained with the structured GMS<br>clinical interview.<br>HTN: European Society of<br>Hypertension criteria (SBP >= 140<br>mm Hg or DBP >= 95 mm Hg) or a<br>positive answer to the question 'have<br>you ever been told by a doctor that<br>you have hypertension?'<br>COPD: People who "usually cough<br>up phlegm from their chest first thing<br>in the morning" "more than 3 months"<br>per year.<br>Diabetes: Persons who had ever<br>been told they have diabetes. | Measures: "Extensive multidomain<br>cognitive testing" (not specified)<br><br>Criteria: DSM-IV, 10/66 dementia<br>diagnosis algorithm<br><br>8.7% with dementia<br><br>Dementia was ascertained according<br>to the cross-culturally validated 10/66<br>dementia diagnosis algorithm and<br>the DSM-IV dementia criterion after<br>extensive multidomain cognitive<br>testing and clinical and informant<br>interview. | Adjusted RR for association btw disability and dementia:<br>All Countries: 1.88 (1.79-1.98)<br>RR's ranged from low (rural India, 1.32 (1.18-1.47)) to a<br>high from Rural Peru: 2.66 (2.25-3.15)<br><br>Compared to other significant CCs, such as depression,<br>stroke, diabetes, heart problems, myocardial<br>infarction/angina, COPD and Hypertension, the RR of<br>disability and self-reported impairments is higher for<br>subjects with dementia. (e.g., the next highest RR was<br>1.39 for depression and for stroke).<br><br><u>How function was assessed:</u> Disability was measured with<br>the 12-item WHODAS 2.0. This short version covers all six<br>domains of the full 36-item version. The schedule has five<br>activity-limitation domains: understanding or<br>communication, getting around (mobility), self-care, getting<br>along with people (interpersonal interaction), and<br>life activities. A sixth domain, participation in society,<br>assesses broad social aspects of disability. Each domain is<br>covered by two questions, with scores ranging from 0 (no<br>difficulty) to 4 (extreme difficulty or cannot do). The<br>standardised global score ranges from 0 (non-disabled) to<br>100 (maximum disability). | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                     | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes  | Quality  |
|--|---|---|---|--|--|
| 16. Zhu<br>1998<br>#2680                       | N=1810<br>82.5(5.2)<br>71.6% female<br>53% 4-7 yrs education    | Stroke (8.4%), hip fracture (10.8%)<br>heart disease (13.4%), cancer<br>(11.9%)   | Measures: MMSE>24<br>Criteria: DSM-III-R  | Adjusted OR for ADLs for dementia vs no dementia:<br>OR for Bathing: Stroke: 3.4 (2.3&#x2013;5.1)<br>OR for Dressing: 8.7 (6.2 - 12.2)<br>OR for Toileting: 5.3 (3.7 - 7.4)<br>OR for Transfer: 10.3 (6.5 - 16.3)<br>OR for Continence: 6.2 (4.2-9.0)  | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: No<br>MULTI: Yes<br>COV ASE: Yes |
| Cross-sectional survey of a longitudinal study | Intl: Stockholm, Sweden<br>Kungsholmen Project                  | ICD Diagnosis:<br>Cases of stroke (ICD-8 codes 430 to 438) were identified through the computerized inpatient register system. There are no private hospitals that treat patients with stroke in this area. Persons with any stroke event recorded in the system before the date of the interview were considered prevalent stroke cases. In Sweden 90% of patients who suffer from a stroke are admitted to a hospital. Most of the stroke patients who are not hospitalized are those who die at home or on the way to the hospital. All kinds of heart disease (ICD-8 codes 390 to 429), cancer (ICD-8 codes 140 to 208 and 230 to 239), and hip fracture (ICD-8 code 820) were detected from the same source. | 12.4% with dementia<br>AD: 121 ,VD: 52, Other: 52<br>Dementia cases were detected by a screening phase (health interview and MMSE) and a clinical exam phase (all subjects with MMSE<24 and a sample of subjects with MMSE > 24). Dementia diagnosis was made according to DSM-III-R. The cognitive exam explored memory (facts of general knowledge and past personal information), language (object naming and comprehension), abstract thinking (problem solving and proverbs), praxis (examining simple motor activities-dressing, pantomime), and visuospatial skills (copying figures). A preliminary diagnosis was first given by the examining physician. All cases were independently reviewed by a neurologist, and a second preliminary diagnosis was made. If there was agreement between the two, this was the final diagnosis. In case of disagreement, a third opinion was asked, and the concordant diagnosis was accepted. For patients with aphasia, a close informant was asked about a patient's everyday behavior and ADLs. Patients who were behaviorally unchanged and routinely attempted to use residual functions were not considered | Population-Attributable Risk Percentages (PAR%) of Dementia to Disability:<br>Bathing: 48.7<br>Dressing: 48.9<br>Toileting: 34.7<br>Transfer: 53.6<br>Continence: 39.2<br>p values are not reported<br>Dementia produced the largest OR of disability compared to hip fracture, stroke, heart disease (OR range 1 - 3), and was the biggest contributor to disability (PARs).<br><u>How function was assessed:</u> Functional disability was assessed according to Katz Index of independence in ADL. The subjects were asked questions regarding their ability to bathe, dress, go to toilet, transfer, maintain continence, and feed. Any dependent performance of these activities was recorded as disability in correspondent items. |  |



SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #<br>Study Design<br>Duration | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics<br>Location<br>DB   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes   | Quality  |
|--|---|---|---|---|--|
| 17. Agüero-Torres, 2002 #2023                          | N=570<br>No Dem: 84.0 (5.2)<br>Dem: 85.0 (5.5)<br>% female (No Dem/Dem) 81.4% / 80.9%<br>% education < 8 yrs: No Dem: 59.5 Dem: 74.4<br>% Single, Divorced, Widow: No Dem: 79.4 Dem: 79.1<br>Intl: Stockholm, Sweden<br>Kungsholmen Project | Cancer (12%), Hip fracture (19%), Cerebrovascular disease (12%), Heart disease (19%)<br><br>ICD Diagnosis:<br>The information was obtained from the Computerized Stockholm Inpatient Register System, which is a register of hospital discharge diagnoses from 1969 to 1987. Disease diagnoses were based on the ICD-8 codes: heart diseases (410-414, 427, 428); cerebrovascular diseases (430 - 438); hip fracture (820) and; cancer (140 - 208 and 230 - 239). A new variable was created to identify when at least one of these somatic disorders, or target chronic conditions (CC) was present.<br><br>Who made assessment NR | Measures: MMSE < 24 (Swedish), Neuropsychological battery (not specified)<br><br>Criteria: DSM-III-R<br><br>48.4% with dementia<br><br>Psychologic tests included the MMSE, episodic, and primary memory tasks as free recall and recognition of random words, and digit span. Dementia was diagnosed using the DSM III-R criteria. As the clinical diagnosis of AD and other dementias presents particular difficulties because of the lack of specific markers, the diagnostic process was double: first, a preliminary diagnosis was made by the examining physician, and all the cases were independently reviewed by a geriatrician who made a second preliminary diagnosis. In case of agreement between the physicians, this was the final diagnosis. In case of disagreement, a third opinion was sought (neurologist, expert in dementia research) before the final diagnosis was accepted. A complete neuropsychologic assessment was also carried out. | Functional disability (%)<br>In at least one ADL:<br>No Dem: 24.1%, Dem: 77.6%<br>In at least one IADL:<br>No Dem: 68.0%, Dem: 97.1%<br><br>Demented subjects had the highest prevalence of dysfunction in all ADL and IADL activities. In ADL, the distribution of dysfunction was similar among demented and nondemented subjects; bathing being the most affected item for all subjects. In IADL, non-demented subjects had most dysfunction in cleaning, while demented subjects had most difficulties in handling economy.<br><br>Adjusted OR (95% CI) for ADL and IADL disability for combined effect of CC + Dementia (reference group = nondemented + no target CC):<br>ADL disability:<br>Dementia w/ no target CC: 4.9 (2.7-8.8)<br>Dementia w/ at least 1 target CC: 26.5 (14.0-49.9)<br>IADL disability:<br>Dementia w/ no target CC: 14.2 (4.8-41.9)<br>Dementia w/ at least 1 target CC: 45.2 (10.7-191.0)<br><br><u>How function was assessed:</u> Functional status was measured as ability to perform basic ADL and IADL. The ADL Katz scale was used, which includes 6 activities: bathing, dressing, toileting, transferring, continence, and feeding. IADL included 5 activities: cleaning, cooking, using public transportation, handling finances, and shopping. Information on both ADL and IADL was obtained directly from the subjects if MMSE > 23, and from a close relative for subjects with MMSE < 24. If the person lived in an institution, the personnel in charge were interviewed. Any dysfunction in the performance of these activities was recorded as dependence in the correspondent item. Because IADL items are often gender-specific, we considered not only the current ability to perform each item, but also the potential capability in case of necessity. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: No<br>MULTI: Yes<br>COV ASE: Yes |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #      | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                    | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Function Outcomes  | Quality   |
|---------------------------------|--|--|---|--|---|
| 18. Agüero-Torres, 1998 #2674   | N=1745<br>81.7(4.9)<br>76% female<br>% high level education:<br>Dem:24%, NoDem:40% | Cancer (12%), Hip fracture (11%), Cerebrovascular disease (9%), Heart disease (17%)<br><br>ICD Diagnosis:<br>The information was obtained from the Computerized Stockholm Inpatient Register System, which is a register of hospital discharge diagnoses from 1969 to 1987. Disease diagnoses were based on the ICD-8 codes: heart diseases (410-414, 427, 428); cerebrovascular diseases (430 - 438); hip fracture (820) and; cancer (140 - 208 and 230 - 239). A new variable was created to identify when at least one of these somatic disorders, or target chronic conditions (CC) was present.<br><br>Who made assessment NR | Measures: MMSE < 24 (Swedish), Neuropsychological battery (not specified)<br><br>Criteria: DSM-III-R<br><br>12% with dementia<br><br>Mean (SD) MMSE:<br>Dem: 12.6 (8.2)<br>No Dem: 26.8 (3.3)<br><br>Psychologic tests included the MMSE, episodic, and primary memory tasks as free recall and recognition of random words, and digit span. Dementia was diagnosed using the DSM III-R criteria. As the clinical diagnosis of AD and other dementias presents particular difficulties because of the lack of specific markers, the diagnostic process was double: first, a preliminary diagnosis was made by the examining physician, and all the cases were independently reviewed by a geriatrician who made a second preliminary diagnosis. In case of agreement between the physicians, this was the final diagnosis. In case of disagreement, a third opinion was sought (neurologist, expert in dementia research) before the final diagnosis was accepted. A complete neuropsychologic assessment was also carried out. | Adjusted OR (95% CI) for Functional Dependence (Katz ADL Index > 1):<br>Cerebrovascular disease and Dementia: 2.3 (0.8-7.3)<br>Heart disease and Dementia: 5.4 (1.4-21.2)<br>Cancer and Dementia: 0.5 (0.1-2.1)<br>Hip fracture and Dementia: 1.7 (0.5-5.2)<br><br>ADL Dependence (Requiring personal assistance in at least 1 of the 6 basic activities) (%):<br>Dem: 77.6, No Dem: 26.1, p value = NR<br><br>Adjusted OR (95% CI) for Risk Factors for Developing Functional Dependence (Katz ADL Index >1 ) after a 3-Year Follow-up Interval:<br>Dementia: 25.2 (9.6-66.4)<br><br>Adjusted OR for 3-Year Functional Decline for Those Who Already Had Functional Dependence at Baseline:<br>Dementia: 2.2 (1.1-4.5)<br><br><u>How function was assessed:</u> Functional dependence = Katz ADL Index > 1. ADL Dependence = Requiring personal assistance in at least 1 of the 6 basic ADL activities. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: Yes<br>COV ASE: Yes |
| Prospective longitudinal cohort | Int: Stockholm, Sweden   |  |   |  |   |
| Mean follow-up: 3.36 years      | Kungsholmen Project  |  |   |  |   |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                        | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Function Outcomes  | Quality   |
|--|--|---|--|--|---|
| 19. Wang<br>2010<br>#A43   | N=4942<br>CogImp: 84.6 (7.4)<br>No CogImp: 84.1 (7.7)<br>~69% female                   | HTN (39%), CVD (20%), Stroke (4%), Arthritis (8%), Depression (10%), Vision impairments (19%), Bladder incontinence (12%), Bowel incontinence (9%)  | Measures: MDS-COG<br><br>Criteria: None<br><br>36.6% with cognitive impairment   | Mean (SD) Total ADL score<br>Baseline:<br>CogImp: 16.5 (7.1) , No CogImp: 12.4 (7.1)<br><br>Follow-Up:<br>CogImp: 16.6 (7.4) , No CogImp: 11.5 (8.0)<br><br>Totally dependent, n (%):<br>Baseline:<br>CogImp: 75 (4.1) , No CogImp: 7 (0.2)<br>Follow-up:<br>CogImp: 84 (4.6), No Cog Imp: 44 (1.4)<br><br>Eating, % (supervision, assistance or dependence):<br>Baseline:<br>CogImp: 62.2%, No Cog Imp: 25.6%<br>Follow-up:<br>Cog Imp: 66% , No CogImp: 28.2%<br><br>Toilet Use, % (supervision, assistance or dependence):<br>Baseline:<br>Cog Imp: 93.1%, No CogImp: 79.5%<br>Follow-Up:<br>CogImp: 90.7%, No CogImp: 71%<br><br>Personal Hygiene, % (supervision, assistance or dependence):<br>Baseline:<br>CogImp: 97%, No CogImp: 83.2%<br>Follow-up:<br>CogImp: 96.7%, No CogImp: 77.6% | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: Yes<br>COV ASE: No |
| Non-experiment<br>longitudinal coh<br>study  | % < 12 yrs education:<br>Dem: 57, No Dem: 63.1<br>% live alone:<br>Dem: 50, No Dem: 65 | *CC prevalence is reported for each<br>CC individually. Subjects may have<br>more than one CC. Mean CC's = 1.5  | The study sample was divided into<br>high and low cognitive function<br>based on the MDS-COGS, which is<br>constructed from eight MDS<br>cognitive function items measured at<br>baseline (range 0-10). Residents<br>with a MDS-COGS score of 0 to 4<br>were categorized as having high<br>cognitive function and those with a<br>MDS-COGS score of 5 or higher<br>were categorized as having low<br>cognitive function. |  |   |
| 4 - 9 months   | U.S.: Minnesota<br><br>Database NR   | Self-report, Standardized Instrument:<br>Participants responding "yes" to both<br>having the CC and reporting having a<br>related treatment were classified as<br>having the CC.<br>The Chinese GDS-SF was used to<br>evaluate the depressive symptoms of<br>the elderly subjects in the past one<br>week. GDS-SF has shown good<br>sensitivity and specificity for<br>predicting depressive disorders in<br>different settings. A cutoff value of<br>>= 5 (0 - 15 range) was used to<br>define geriatric depression.<br><br>All CC data was collected during<br>home visits with well-trained<br>interviewers. | Who made assessment NR   |  |   |
| p = NR for above comparisons. Notes: Participants with no cognitive impairment were less dependent in all ADL measures at baseline and follow-up than those with cognitive impairment. |  |   |  |  |   |

SIP-37 Cognitive Impairment and Co-occurring Chronic Conditions: Evidence Table

| Author       | # Sample (N)       | Chronic Condition (CC): | Cognitive Impairment (CogImp)/Dementia (Dem): | Function Outcomes | Quality |
|--------------|--------------------|-------------------------|---|-------------------|---------|
| Yr           | Mean age (SD)      | Definition              |   |                   |         |
| Tracking #   | % Female           | Assessment              | Measures, Criteria, % with                    |                   |         |
| Study Design | Other Demographics | Who made assessment     | Dem/CogImp, Mean Scores,                      |                   |         |
| Duration     | Location           |                         | Assessment                                    |                   |         |
|              | DB                 |                         |   |                   |         |

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Wang  
(continued)

NH random effects were much stronger for residents with a no cognitive impairment. For those with no cognitive impairment, NH random effects were SS for total ADLs, toileting, and personal hygiene, whereas for those with cognitive impairment, NH effect was significant only for eating function.

How function was assessed: Total ADL scores at baseline and follow-up were aggregated from seven MDS items (bed mobility, transfer, locomotion on unit, dressing, eating, toilet use, and personal hygiene), each item rated from 0 (totally independent) to 4 (totally dependent), resulting in a total ADL score ranging from 0 to 28.

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #             | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes   | Quality   |
|--|--|--|---|---|---|
| 6 studies                              | N = 8236   | Hypertension   |   | Anti-HTN medications  |   |
| 15. Huang<br>2009<br>#221              | N=782<br>93.62(NR)<br>All in their 90's and 100's<br>67.5% female              | % with hypertension:<br>CogImp: 56.99%,<br>No CogImp: 57.10%   | Measures: MMSE<19<br><br>Criteria: NA   | % with antihypertensive medication (unadjusted)<br>CogImp: 4.52%<br>No CogImp: 4.41%<br>p=0.947 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Cross-sectional<br>design and duration | Intl: Dujiangyan, China<br><br>Project of Longevity and<br>Aging in Dujiangyan | Self-Report/Interview, Clinical<br>Exam/Diagnosis:<br>Trained study personnel conducted<br>standardized face-to-face<br>interviews, including self-reported<br>medical history, medication,<br>anthropometric measurements,<br>standardized physical examination,<br>and a 12-lead electrocardiogram. BP<br>was taken twice, to nearest 2 mm Hg,<br>using standard mercury<br>sphygmomanometer. The mean value<br>of the two measurements was used to<br>calculate SBP and DBP according to<br>the Joint National Committee VII<br>criteria. HTN = SBP>140 mm Hg<br>and/or DBP>90 mm Hg and/or<br>receiving antihypertensive treatment.<br><br>Participants with confirmed HTN and<br>no identified cause of secondary HTN<br>were diagnosed with essential HTN<br>Persons with cancer, type 2 diabetes,<br>secondary HTN, severe heart failure,<br>and terminal stage COPD were<br>excluded . | 59.5% with cognitive impairment<br>3% had vascular dementia.<br><br>Mean (SD) MMSE: 14.95 (5.99)<br>CogImp: 11.63 (4.38)<br>No CogImp: 21.03 (2.49)<br><br>Trained professional physicians<br>administered the MMSE during the<br>face-to-face interview. In addition to<br>the MMSE the physicians also asked<br>about dementia diagnoses during<br>the face-to-face interview. Vascular<br>dementia was defined as people with<br>dementia combined with a history or<br>clinical evidence of stroke.<br><br>"No CogImp" included people with<br>mild cognitive impairment (MMSE 19-<br>24, 27% of the sample). |   |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #          | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes  | Quality  |
|-------------------------------------|---|--|---|--|--|
| 16. Stewart<br>2009<br>#185         | N=1890<br>Ages 77 - 96<br>0% female                             | % with hypertension:<br>No Dem: 46%, Incident Dem: 49%<br>AD: 43%, VaD: 80%  | Measures: NR ("Cognitive<br>prescreening and neuropsych-<br>ological testing")  | % Antihypertensive medication use (unadjusted)<br>No Dem: 40%<br>Incident Dementia: 41%<br>Incident AD: 37%<br>Incident VaD: 73%<br>p = NR | SAMPLE: Yes<br>GENDER: No<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Longitudinal<br><br>Mean f/u 32 yrs | U.S.: Oahu, HI<br><br>Honolulu Heart Program and                | Self-Report, Clinical Exam/<br>Diagnosis, Standardized<br>Instrument:<br>BP was measured with the same<br>standardized protocol at each<br>examination. After the participant had<br>been seated for 10 minutes, SBP and<br>DBPs were measured on 3 occasions,<br>5 minutes apart on the left arm of a<br>seated participant using a mercury<br>sphygmomanometer with a standard<br>cuff. DBP was recorded as the fifth<br>phase. Repeated readings were<br>averaged for each examination.<br><br>Other CC: Diabetes, Depression,<br>Smoking<br><br>Depressive symptoms were<br>measured using the CES-D. Smoking<br>status was ascertained from previous<br>exams from the study. | Criteria: DSM-III (R), NINCDS<br>-ADRDA, ADDTC, Hachinski<br>Ischemic Scale<br>59% with dementia<br>AD: 74 ,VD: 15, Other: 25<br>3-stage procedure for dementia case<br>finding: cognitive prescreening,<br>neuropsychological testing, proxy<br>interview, neurological examination,<br>and neuroimaging. Consensus<br>diagnoses were made by neurologist<br>and 2 physicians. Diagnosis made<br>according to DSM-III-R (dementia),<br>NINCDS-ADRDA (AD), and ADDTC<br>(VaD). The criteria for probable VaD<br>require dementia, computed<br>tomography/MRI evidence of 1<br>infarct outside of the cerebellum,<br>and then either clinical/ imaging<br>evidence of 2 ischemic strokes or a<br>single stroke with a clear temporal<br>relationship to the onset of<br>dementia. Additional support is<br>allowed if evidence of multiple<br>infarcts in brain regions known to<br>affect cognition, multiple TIAs,<br>history of vascular risk factors, and<br>elevated Hachinski Ischemic Scale<br>score. |  |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes   | Quality  |
|------------------------------|---|---|---|---|--|
| 17. Vinyoles<br>2008<br>#404 | N=1579<br>70.1(6.7)<br>55.6% female<br>52% w/ incomplete<br>primary education<br>60% married/partner<br>17% living alone<br>77% rural<br><br>Intl: Spain<br><br>COGNIPRES Study | 100% w/ arterial hypertension<br><br>Duration: Mean 9.5 (6.7) years<br><br>Self-report, Clinical Exam/<br>Diagnosis:<br>Hypertension = SBP >= 140 mm<br>Hg or DBP < 90 mm Hg. BP was<br>determined by two measurements<br>spaced two minutes apart, with the<br>patient in the resting position, and<br>using a sphygmomanometer from<br>each clinic (conventional mercury<br>device or validated automated<br>oscillometric device) in order to<br>reproduce the conditions of routine<br>clinical practice. When differences<br>between the first and second<br>measurements were over 5 mmHg for<br>systolic or diastolic pressure, a third<br>measurement was made. | Measures: MMSE<br><br>Criteria: None<br><br>12.3% with cognitive impairment<br><br>25.9 (4.8)<br><br>The main study variable was the<br>prevalence of cognitive impairment,<br>which was assessed by the MMSE,<br>adjusted for educational level in<br>order to avoid false positive or false<br>negative results. The cut-off points<br>were 17/18 for no education, 20/21<br>for incomplete primary education,<br>and 23/24 for primary education and<br>over 27. The validated Spanish<br>version of the MMSE was used and<br>administered by a physician during<br>the clinical exam. | High Blood Pressure treatment % (N) (unadjusted)<br>CogImp: 12.3% (194)<br>No CogImp: 87.7% (1379)<br>p=0.028<br><br>Monotherapy<br>CogImp: 1 0.1% (65)<br>No CogImp: 89.9% (576)<br>p=NR<br><br>Combined therapy<br>CogImp: 13.8% (129)<br>No CogImp: 86.2% (803)<br>p=NR<br><br>Compliance to antihypertensive treatment:<br>Compliant patients<br>CogImp: 9.1% (94)<br>No CogImp: 90.9% (942)<br><br>Non-Compliant<br>CogImp: 19.0% (100)<br>No CogImp: 81.0% (425)<br><br>p<0.001 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: Yes<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author              | # Sample (N)       | Chronic Condition (CC):  | Cognitive Impairment (CogImp)/Dementia (Dem):                                   | Medication Outcomes   | Quality |
|---------------------|--------------------|--|---|---|---------|
| Yr                  | Mean age (SD)      | Definition   | Measures, Criteria, % with Dem/CogImp, Mean Scores, Assessment                  |   |         |
| Tracking #          | % Female           | Assessment   |   |   |         |
| Study Design        | Other Demographics | Who made assessment  |   |   |         |
| Duration            | Location           |  |   |   |         |
|                     | DB                 |  |   |   |         |
| 17. Vinyoles (cont) |                    | <p>Other CC: PD, Epilepsy, Stroke, Cancer, Lipidemia, Diabetes, Depression, Psychosis, Personality disorders, Substance use (drinking)</p> <p>Self-Report, Clinical Exam/Diagnosis:<br/>Assessment details: Data collection was performed during a single visit by completing a normalized questionnaire on demographic data, cardiovascular risk factors, previous clinical cardiovascular disease and current treatment.</p> | Note: Persons with previously diagnosed dementia were excluded from this study. | <p>Notes: Non-compliant patients with anti HTN treatment was defined as those who answered affirmatively in response to the Haynes-Sackett questionnaire item: "Most patients have difficulties in taking all their tablets, do you have difficulties in taking yours?". Patients yielding a negative answer to the same question, with three or four incorrect answers in the Morisky-Green test were also regarded as non-compliers. The Morisky-Green test is the most specific adherence questionnaire applied to hypertensive populations.</p> <p>The most frequent monotherapy comprised angiotensin II receptor antagonists (28.6%), diuretics (25.5%), angiotensin-converting enzyme inhibitors (ACEIs) (20.3%) and a fixed combination of angiotensin II receptor antagonists and diuretics (20%).</p> |         |



SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking # | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                             | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Medication Outcomes  | Quality  |
|----------------------------|---|---|--|--|--|
| 18. Hanon<br>2006<br>#1065 | N=1241<br>78(8)<br>67% female<br>% primary education:<br>No Dem: 27%<br>AD: 31%<br>VaD: 37% | 100% with hypertension<br><br>Clinical Exam/Diagnosis:<br>SBP and DBP were measured<br>during the consultation, by nurses,<br>3 times in each patient, after at<br>least 5 minutes of rest seated, on<br>the left arm, using a validated<br>electronic device. The average of the<br>3 measurements was used to<br>determine the BP level. Individuals<br>with SBP 140 mmHg or greater and<br>DBP 90 mmHg or greater or<br>those taking antihypertensive<br>medication were considered to be<br>hypertensive.<br><br>Other CC: CHD, Stroke, Heart<br>Failure, Atrial Fibrillation<br><br>Self-Report, Admitted to<br>Clinic/Program: Collected during<br>consultations at a geriatric memory<br>clinic. | Measures: MMSE, CEP<br>Criteria: DSM-IV, NINDS-AIREN<br>49% with dementia<br>AD: 524 ,VD: 85<br><br>Mean MMSE:<br>AD: 19 (+/-6), VaD 19 (+/-5)<br>No Dem: 28(+/-1)<br><br>Mean CEP scores (max 100)<br>AD: 29 (+/-15), VaD: 28 (+/-17)<br>No Dem: 75 (+/-8)<br><br>Subjects were screened during the<br>consultation by a physician. At the<br>end of the evaluation by<br>psychologists and physicians,<br>patients were classified into<br>four subgroups: AD (using,<br>DSM-IV), VaD (using<br>NINDS-AIREN), MCI and normal<br>cognitive function. The normal<br>group comprised patients with no<br>disease known to alter cognitive<br>function. They had normal scores on<br>the CEP according to age, sex and<br>education (score > mean 1.5 SD)<br>and were autonomous in their<br>activities of daily living. | % using antiHTN therapy:<br>No Dem: 60%<br>AD: 53%<br>VaD: 65%<br>p = NR<br><br>Treated hypertensive patients had better<br>cognitive function than untreated patients<br>(adjusting for covariates), and this association<br>was observed independently of the cognitive<br>status (in normal, AD and VaD hypertensive patients).<br><br>Note: AntiHTN therapy = diuretics, beta-blockers, calcium<br>antagonists, angiotensin-converting enzyme (ACE)<br>inhibitors, angiotensin receptor blockers (ARB) and other<br>drugs. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>CI: Yes<br>CC:<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Medication Outcomes   | Quality   |
|--|--|---|--|---|---|
| 19. Gombojav<br>2011<br>#A5<br><br>Longitudinal<br><br>Mean 11.8 yrs | N=2496<br>73.6(5.9)<br>57.7% female<br>% no formal education:<br>CogImp: 80%<br>No CogImp: 50%<br><br>INTL: South Korea<br><br>Database NR | 18% w/ hypertension (same % for<br>CogImp and NoCogImp)<br><br>Clinical Exam/Diagnosis:<br>SBP/DBP: <= 140/90 mm Hg<br>Avg 2 BP measurement using<br>standard mercury sphygmo-<br>manometer, assessed by<br>trained interviewer<br><br>Other CC: Overall comorbidity,<br>Smoking (former and current),<br>Drinking, BMI<br>Overall comorbidity: Subjects<br>answered yes or no to the<br>question "do you have any<br>chronic disease or past accident<br>or injury due to which you feel<br>uncomfortable in your daily life<br>including work?" | Measures: MMSE<br><br>Criteria: None<br><br>44% with CogImp<br><br>Mean MMSE (SD):<br>mild severity CogImp: 17.6 (1.1)<br>severe CogImp: 11.9 (3.1)<br>no CogImp: 24.1 (2.9)<br><br>MMSE administered by the<br>investigation team | % ever taking antiHTN medications:<br><br>No CogImp: 18.3<br>Mild Severity CogImp: 18.8<br>Severe CogImp: 18.6<br><br>% never taking antiHTN medications:<br><br>No CogImp: 61.1<br>Mild Severity CogImp: 62.1<br>Severe CogImp: 57.1<br><br>p = 0.76 | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: No<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics                                 | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes  | Quality   |
|---|---|--|---|--|---|
| 20. Freels<br>2002<br>#2055<br><br>Cohort study                     | N=248<br>45+<br>% female NR<br>Other demog NR<br><br>U.S.: Chicago, Illinois<br><br>Database NR | 46.5% AD, 74.7% VaD, 85.3% No<br>Dem with HTN<br><br>Self-Report, Standardized Instrument:<br>An epidemiologic interviewer<br><br>administered standardized<br>questionnaires to determine CC's. | Measures: MMSE, BOMCT, Formal<br>neuropsychological testing<br><br>Criteria: ~NINDS-AIREN<br><br>77% with dementia.<br>AD: 113 ,VD: 79  | Taking antihypertensive medication (%)<br>AD: ~47%<br>VaD: ~48%<br>No Dem: ~48%<br>p = NR<br><br>Adjusted HR for survival for VaD:<br>Taking antiHTN meds: 4.69 (p = 0.0001) | SAMPLE: No<br>GENDER: NR<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br><br>MULTI: Yes<br>COV ASE: Yes |
| 7 years (median<br>f/u: AD: 5.9 years,<br>VaD: 5.8, No Dem:<br>6.1) | Patients were referred from<br>a hospital-based stroke<br>registry and an academic<br>AD center | Other CC: Stroke, MI, AF, Lipidemia,<br>Diabetes, Depression, Smoking<br><br>Same methods as described above   | All study patients completed<br>diagnostic measures with trained<br>interviewers and informants<br>completed a structured neurologic<br>interview. The following criteria were<br>used: AD = met dementia criteria<br>and no other conditions contributing<br>to cog impairment; VaD = dementia<br>+ diagnosis of stroke<br><br>by Stroke Data Bank criteria, and a<br>temporal relationship between stroke<br>and dementia onset (VaD diagnosis<br>predated both of the currently most<br>commonly used diagnostic systems<br>today). These criteria are consistent<br>with NINDS-AIREN. Persons<br>meeting both VaD and AD criteria<br>were excluded.<br><br>The "No dementia" group included<br>patients who met criteria for<br>neurologic dysfunction due to<br>vascular disease (stroke) but did not<br>meet criteria for dementia. |  |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Medication Outcomes                               | Quality  |
|------------------------------|--|--|--|---|--|
| 21. Barzilay<br>2008<br>#552 | N=2316<br>Age 65 - 75 / > Age 80<br>Dem: 14.5% / 49.5 %<br>No Dem: 38.7% / 22.1%   | 49% with hypertension<br><br>Self-report, Clinical Exam/Diagnosis:<br>At annual clinic visits, participants<br>underwent baseline blood testing,<br>cardiac and carotid artery ultrasound<br>testing, electrocardiography, ankle-<br>brachial index measurement, and<br>completion of medical history and<br>clinical examination with their<br>physician. | Measures: 3MS < 80, TICS < 28<br>IQCODE > 3.6<br><br>Criteria: NINCDS-ADRDA, NINDS-AI p = 0.002<br><br>27% with dementia   | Any antiHTN use (%)<br>Dem: 7.6%<br>No Dem: 55.8% | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Design NR<br>Cross-Sectional | % Female<br>Dem: 59%<br>No Dem: 59.1%<br><br>Race: (%)<br>Nonwhite: (%)<br>Dementia: 19.4%<br>No Dem: 9.9%<br>White: %<br>Dementia: 80.6%<br>No Dem: 90.1%<br><br>< HS education:<br>Dem: 33.9%<br>No Dem: 18.3%<br><br>U.S.: Cardiovascular Health<br>Study Centers<br><br>Cardiovascular Health<br>Study | Other CC: Stroke, TIA, CHF, CHD,<br>Substance use (drinking, smoking),<br>Obesity<br><br>Same methods as described above   | Participants completed the 3MS<br>annually at clinic visits or the TICS<br>by phone if they did not come to the<br>clinic. The IQCODE was used for<br>additional information. If person is<br>unable to complete 3MSE or TICS, a<br>physician provided additional<br>information. Diagnosis of dementia<br>was based on a deficit in<br>performance in 2 or more cognitive<br>domains of sufficient severity to<br>affect ADLs and a history of normal<br>intellectual function before the<br>cognitive decrease. Diagnosis was<br>made by an adjudication committee<br>of neurologists with expertise in<br>dementia. Medical charts were also<br>reviewed for diagnosis of dementia. |   |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #               | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Medication Outcomes  | Quality  |
|--|---|--|--|--|--|
| 8 studies                                | N = 5245  | CVD  |  | CVD Medications (besides anti-HTN)   |  |
| 22. Lopponen<br>2006<br>#1143            | N=462<br>All age 75+<br>Dem: 79.8 (4.4)<br>No Dem: 84.4 (5.7)   | HTN, CAD, CHF, Diabetes,<br>AF, Hypercholesterolaemia<br><br>Mean (SD) CVD's:<br>Dem: 2.6(1.2), No Dem: 2.2(1.1)   | Measures: MMSE < 24<br><br>Criteria: DSM-IV, NINCDS-<br>ADLDA, NINDS-AIREN, CDR,<br>Hachinski Ischaemic Scale  | Adjusted OR CVD meds for CVD patients:<br>Dem: 0.31 (95% CI 0.12-0.82)<br><br>Cumulative OR for receiving a growing number<br>of medications:<br>Moderately demented: 0.39 (95% CI 0.20-0.76)<br>Severely demented: 0.50 (95% CI 0.19-1.34)  | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Longitudinal<br>epidemiological<br>study | <= 6 yrs education:<br>Dem: 78%<br>No Dem: 69%  | Self-Report, Clinical Exam/<br>Diagnosis, ICD Diagnosis,<br>Records Review:<br>Study protocol consisted of an<br>interview, lab visit, clinical exam, and<br>records review. 2 specially trained<br>research nurses conducted IWs to<br>assess physical, mental and social<br>health. BP was measured with a<br>mercury sphygmo-manometer after 5-<br>min rest with subject sitting. The mean<br>value of 2 measurements was used. A<br>close informant or nursing staff were<br>interviewed if participant was unable<br>to provide info. | 79.7% with dementia<br>AD: 40 ,VD: 35<br><br>Mean MMSE (SD):<br>Dem: 14.9 (7.7)<br>No Dem: 27.1 (2.5)<br><br>A 2-stage design was applied to<br>assess the occurrence of dementia.<br>First, the MMSE was performed to<br>screen cognitive functioning, and the<br>cut-off point of 23/24 was used for<br>further evaluation. Persons with an<br>MMSE score of 24–30 having a<br>previous history of a dementing<br>disorder in their medical records or<br>clinical suspicion of dementia in the<br>interview or clinical | (N) % Taking CVD Medications (unadjusted):<br><br>Antithrombotic agents (B01A):<br>Dementia (n=34): 40.0<br>Without Dementia (n=113): 34.9<br>p = NS<br><br>Nitrates in ischaemic heart disease (C01D)<br>Dementia (n=23): 27.1<br>Without Dementia (n=74): 22.8<br>p=NS<br><br>Beta-Blockers (C07A)<br>Dementia (n=13): 15.3<br>Without Dementia (n=88): 27.2<br>p=NS |  |
| Cross-Sectional                          | Unmarried:<br>Dem: 78%<br><br>No Dem: 57%<br><br>53% of people with<br>dementia lived in an<br>institution, compared with<br>1% of those without<br>dementia. |  |  |  |  |
|  | Intl: Lieto, Finland<br><br>Database NR   |  |  |  |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes   | Quality |
|-----------------------------|---|---|---|---|---------|
| 22. Lopponen<br>(continued) |   | <p>Two research physicians (experienced GP's) reviewed available primary-care medical records, registered medications used, and all diagnosed diseases and other relevant conditions. Diagnoses were coded according ICD-10. They also carried out clinical examinations, in which all info from interviews, medical records, and lab tests were available.</p> <p>In addition to looking at diagnoses in the medical records, CC's were defined as:<br/>                     *HTN = entitled to reimbursements from the NHI for HTN and/or had SBP&gt;160 mm Hg and/or DBP &lt; 100 mm Hg.<br/>                     *CHD = history of coronary by-pass operation or angioplasty and/or entitled to reimbursements from the NHI for CHD medication and/or had ischaemic ECG findings (major/moderate Q/QS item as a sign of MI, and/or a minor Q/QS item, S-T depression, T wave inversion or left bundle branch block as a sign of possible CHD)<br/>                     *CHF = entitled to reimbursements from the NHI for CHF medication.</p> | <p>examination were also included in the second stage. In the case of institutionalised or hospitalised patients, caregivers or nursing staff were interviewed. The interview was semi-structured and covered the items of the Hachinski Ischaemic Scale and the CDR Scale.</p> <p>The latter was used to stage the severity of the dementia. Finally, dementia was assessed in the clinical examination according to the DSM-IV criteria, diagnosis of possible AD according to NINCDS-ADRDA criteria and the diagnosis of possible VaD according to the NINDS-AIREN criteria. In cases of disagreement, a consensus was reached between the research physicians and the geriatrician.</p> | <p>Cardiac glycosides (C01A)<br/>Dementia (n=18): 21.1<br/>Without Dementia (n=36): 11.1<br/>p=NS</p> <p>Calcium antagonists, cardioselective (C08D)<br/>Dementia (n=9): 10.6<br/>Without Dementia (n=40): 12.4<br/>p=NS</p> <p>ACE inhibitors (C09A)<br/>Dementia (n=5):5.9<br/>Without Dementia (n=41): 12.7<br/>p=NS</p> <p>Minor analgesics and antipyretics (N02B)<br/>Dementia (n=15):17.7<br/>Without Dementia (n=29): 9.0<br/>p=NS</p> <p>Laxatives (A06A)<br/>Dementia (n=18): 21.2<br/>Without Dementia (n=22): 6.8<br/>p=NS</p> <p>Thyroxin (H03A)<br/>Dementia (n=12): 14.1<br/>Without Dementia (n=27): 8.3<br/>p=NS</p> |         |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment  | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment | Medication Outcomes   | Quality |
|-----------------------------|---|---|--|---|---------|
| 22. Lopponen<br>(continued) |   | <p>*AF = had AF at ECG.</p> <p>*Stroke = subjective history of stroke with neurological symptoms persisting for more than 24 h, verified in the clinical examination.</p> <p>*Hypercholesterolaemia = treated with lipid-lowering agents (ATC code C10) and/or had a fasting serum total cholesterol concentration 66.5 mmol/l.</p> <p>*Diabetes = treated with antidiabetic agents (ATC code A10) and/or had a fasting plasma glucose level 67.0 mmol/l.</p> |  | <p>Peptic ulcer and reflux drugs (A02B)<br/>Dementia (n=14): 16.5<br/>Without Dementia (n=25): 7.7<br/>p=NS</p> <p>Antiglaucoma preparations and miotics (S01E)<br/>Dementia (n=6): 7.1<br/>Without Dementia (n=31): 9.6<br/>p=NS</p> <p>Calcium antagonists, angioselective (C08C)<br/>Dementia (n=7): 8.2<br/>Without Dementia (n=29): 9.0<br/><br/>p=NS</p> <p>Potassium (A12B)<br/>Dementia (n=6): 7.1<br/>Without Dementia (n=27): 8.3<br/>p=NS</p> <p>Antigout preparations (M04A)<br/>Dementia: 7.1<br/>Without Dementia: 8.3<br/>p=NS</p> |         |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author                      | # Sample (N)       | Chronic Condition (CC): | Cognitive Impairment (CogImp)/Dementia (Dem):                  | Medication Outcomes   | Quality |
|-----------------------------|--------------------|-------------------------|--|---|---------|
| Yr                          | Mean age (SD)      | Definition              |  |   |         |
| Tracking #                  | % Female           | Assessment              | Measures, Criteria, % with Dem/CogImp, Mean Scores, Assessment |   |         |
| Study Design                | Other Demographics | Who made assessment     |  |   |         |
| Duration                    | Location           |                         |  |   |         |
|                             | DB                 |                         |  |   |         |
| 22. Lopponen<br>(continued) |                    |                         |  | <p>Sulphonamides and trimethoprim (J01E)<br/>Dementia (n=15): 17.7<br/>Without Dementia (n=16): 4.9<br/>p &lt; 0.001</p> <p>Loop diuretics (C03C)<br/>Dementia (n=8): 9.4<br/>Without Dementia (n=20): 6.2<br/>p=NS</p> <p>Potassium-sparing diuretics (C03E)<br/>Dementia (n=31): 36.5<br/>Without Dementia (n=93): 28.7<br/>p=NS</p> <p>NSAIDs (M01A):<br/>Dementia (n=19): 22.4<br/>Without Dementia (n=11): 34.3<br/>p &lt; 0.05</p> <p>Notes: This article aimed to examine medication use in persons with CVD both with and without dementia. Depression prevalence was not reported.</p> <p>Cardiovascular medications were defined as ATC groups C01 (cardiac glycosides, anti-arrhythmics, nitrates), C02 (antihypertensives), C03 (diuretics), C04 peripheral vasodilators), C07 (-blockers), C08 (calcium channel blockers), C09 (ACE inhibitors, angiotensin II receptor blockers), C10 (lipid-lowering agents) and B01</p> |         |



SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #  | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Medication Outcomes  | Quality  |
|---|---|--|--|--|--|
| 23. Bursi<br>2006<br>#1273  | N=1832<br>Median: 82 (38-102 yrs)<br>NR % female<br>Other Demog NR  | Myocardial infarction (MI)<br><br>ICD Diagnosis, Records Review,<br>Standardized Instrument:<br>MI assessed using ICD-9 codes from   | Measures: NR<br><br>Criteria: DSM-IV, H-ICDA code<br><br>50% with dementia   | There were NS differences between subjects with and without dementia in thrombolysis, ace inhibitors, beta blockers, and aspirin use after myocardial infarction.<br><br>Unadjusted analysis, data not reported. | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: No<br>CC: No<br>MULTI: No |
| Mean f/u:<br>Dem: 6.1 years (0.4-19.5),<br>No Dem: 8.2 years (1.1-19.5)<br>p < .001 | U.S.: Rochester, MN<br>Rochester Epidemiology Project (using Mayo clinic and othe records of Rochester residents) | discharged diagnoses in medical records. Trained nurse abstractors validated the diagnosis of MI using standardized criteria for definite or probable MI (cardiac pain, biomarker values, and Minnesota coding of the electrocardiogram).<br><br>Trained nurse abstractors reviewed the records and compared against standardized criteria.<br><br>Other CC: HTN, Lipidemia, Diabetes, Substance use (smoking)<br><br>Assessment details and who made the assessment of other CC's were NR | Searched for H-ICDA codes in medical records. Each potential case (at least one H-ICDA code) was screened by trained nurse abstractors. A neurologist confirmed the presence of dementia using DSM-IV. |  | COV ASE: No  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #    | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes  | Quality  |
|-------------------------------|--|--|---|--|--|
| 24. Andersson<br>2008<br>#798 | N=349<br>Median age: AD: 76,<br>DLB: 77, No Dem: 73<br>% female: AD: 69%,<br>DLB: 44%, No Dem: 66%<br>Other demog NR | CVD (MI, CAD, Peripheral Artery<br>Disease), HTN, Arteriosclerosis<br><br>Clinical exam:<br>All participants completed a physical<br>examination and cognitive testing.<br>This included the collection of data on<br>current medications and BP<br>measurement. The drugs were<br>classified by the Anatomical<br>Therapeutic Chemical Classification<br>system (ATC) recommended by the<br>World Health Organization (WHO)<br>(2001). All the patients attended the<br>Neuropsychiatric Clinic, Malmö<br>University Hospital.<br><br>Who made assessment NR<br><br>This article targeted orthostatic<br>hypotension (not a target chronic<br>condition of this review) and looked at<br>other chronic conditions that influence<br>blood pressure. | Measures: MMSE, ADAS<br><br>Criteria: NINCDS-ADRDA (AD),<br>McKeith (probable DLB)<br><br>82.2% with dementia<br>AD: 235, DLB: 52<br><br>Mean (SD) MMSE Scores:<br>AD: 21 (5), DLB: 22 (5)<br>No Dem: 29 (1)<br><br>Detailed clinical investigation:<br>anamnesic data, physical and<br>neuropsychiatric exam, cognitive<br>measures, blood and cerebrospinal<br>fluid sampling, brain CT, regional<br>cerebral blood flow, EKG and BP<br>measurements. Probable AD mild or<br>moderate were included. All patients<br>diagnosed with DLB during the study<br>were included. Dementia diagnosis<br>was given prospectively. Controls:<br>recruited through advertisements,<br>completed physical examination and<br>cognitive testing. Inclusion criteria =<br>absence of memory complaints or<br>any other cognitive symptoms,<br>preservation of general cognitive<br>functioning and no active<br>neurological or psychiatric disease. | This article included medications that may influence<br>orthostatic hypotension (OH) / blood pressure.<br><br>N (%) AntiHTN / cardiac therapy<br>AD: 79 (34%), p = NS<br>DLB: 19 (37%), p = NS<br>No Dem: 23 (37%)<br><br>N (%) antidepressant medications<br>AD: 98 (42%), p < 0.001<br>DLB: 26 (51%), p < 0.001<br>No Dem: 3 (5%)<br><br>N (%) Antipsychotics, Anxiolytics, Sedatives/Hypnotics<br>AD: 74 (32%), p < 0.001<br>DLB: 28 (55%), p < 0.001<br>No Dem: 1 (2%)<br><br>Note: Specific CVD medications are not described in the<br>article. This article looked at use of anti HTN and other<br>CVD medications since these meds have been known to<br>influence blood pressure. | SAMPLE: No<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #                                 | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics   | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes   | Quality   |
|--|---|--|---|---|---|
| 25. Freels<br>2002<br>#2055<br><br>Cohort study            | N=248<br>45+<br>% female NR<br>Other demog NR<br><br>U.S.: Chicago, Illinois                              | Stroke, Myocardial Infarction, Atrial fibrillation, HTN, Lipidemia, Diabetes, Depression, Substance use (smoking)  | Measures: MMSE, BOMCT, Formal neuropsychological testing<br><br>Criteria: ~NINDS-AIREN  | Taking aspirin or antiplatelet/anticoagulant medication (%):<br>AD: ~41%<br>VaD: ~37%<br>No Dem: ~50%<br>p = NR | SAMPLE: No<br>GENDER: NR<br>MCC: Yes<br>DEM: Yes<br>CC: Yes |
| 7 years (median f/u: AD: 5.9 years, VaD: 5.8, No Dem: 6.1) | Database NR<br><br>Patients were referred from a hospital-based stroke registry and an academic AD center | Self-Report, Standardized Instrument:<br>An epidemiologic interviewer administered standardized questionnaires to determine CC's.<br>Stroke: The study neurologist completed the Stroke Data Bank Neurologic Examination and an unstructured neurologic interview with informants. | 77% with dementia.<br>AD: 113 ,VD: 79<br><br>All study patients completed diagnostic measures with trained interviewers and informants completed a structured neurologic interview. The following criteria were used: AD = met dementia criteria and no other conditions contributing to cog impairment; VaD = dementia + diagnosis of stroke by Stroke Data Bank criteria, and a temporal relationship between stroke and dementia onset (VaD diagnosis predated both of the currently most commonly used diagnostic systems today). These criteria are consistent with NINDS-AIREN. Persons meeting both VaD and AD criteria were excluded.<br><br>The "No dementia" group included patients who met criteria for neurologic dysfunction due to vascular disease (stroke) but did not meet criteria for dementia. | Adjusted HR for survival for VaD:<br>Taking aspirin or antiplatelet/anticoagulant meds: 0.30 (p = 0.0084)       | MULTI: Yes<br>COV ASE: Yes                                  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #   | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment   | Medication Outcomes                                  | Quality  |
|------------------------------|--|--|--|--|--|
| 26. Barzilay<br>2008<br>#552 | N=2316<br>Age 65 - 75 / > Age 80<br>Dem: 14.5% / 49.5 %<br>No Dem: 38.7% / 22.1%   | 49% with hypertension<br><br>Self-report, Clinical Exam/Diagnosis:<br>At annual clinic visits, participants<br>underwent baseline blood testing,<br>cardiac and carotid artery ultrasound<br>testing, electrocardiography, ankle-<br>brachial index measurement, and<br>completion of medical history and<br>clinical examination with their<br>physician.<br><br>Other CC: Stroke, TIA, CHF, CHD,<br>Substance use (drinking, smoking),<br>Obesity<br><br>Same methods as described above | Measures: 3MS < 80, TICS < 28<br>IQCODE > 3.6<br><br>Criteria: NINCDS-ADRDA, NINDS-AI p = 0.005<br><br>27% with dementia<br><br>Participants completed the 3MS<br>annually at clinic visits or the TICS<br>by phone if they did not come to the<br>clinic. The IQCODE was used for<br>additional information. If person is<br>unable to complete 3MSE or TICS, a<br>physician provided additional<br>information. Diagnosis of dementia<br>was based on a deficit in<br>performance in 2 or more cognitive<br>domains of sufficient severity to<br>affect ADLs and a history of normal<br>intellectual function before the<br>cognitive decrease. Diagnosis was<br>made by an adjudication committee<br>of neurologists with expertise in<br>dementia. Medical charts were also<br>reviewed for diagnosis of dementia. | ACE inhibitor use (%)<br>Dem: 18.4%<br>No Dem: 13.1% | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Design NR<br>Cross-Sectional | % Female<br>Dem: 59%<br>No Dem: 59.1%<br><br>Race: (%)<br>Nonwhite: (%)<br>Dementia: 19.4%<br>No Dem: 9.9%<br>White: %<br>Dementia: 80.6%<br>No Dem: 90.1%<br><br>< HS education:<br>Dem: 33.9%<br>No Dem: 18.3%<br><br>U.S.: Cardiovascular Health<br>Study Centers<br><br>Cardiovascular Health<br>Study |  |  |  |  |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

| Author<br>Yr<br>Tracking #       | # Sample (N)<br>Mean age (SD)<br>% Female<br>Other Demographics  | Chronic Condition (CC):<br>Definition<br>Assessment<br>Who made assessment   | Cognitive Impairment<br>(CogImp)/Dementia (Dem):<br><br>Measures, Criteria, % with<br>Dem/CogImp, Mean Scores,<br>Assessment  | Medication Outcomes  | Quality   |
|----------------------------------|--|--|---|--|---|
| 27. Cankurtaran<br>2008<br>#779  | N=1436<br>72.7(6.9)<br>65.8% female<br>Living status (%)<br>Alone: Dem 51.7%,<br>No Dem 54.1%  | Stroke, CVD, HTN, Diabetes,<br>Depression, Lipidemia,<br>Cerebrovascular disease,<br>Substance use (drinking, smoking)   | Measures: MMSE, CDT<br><br>Criteria: DSM-IV, NINCDS<br>-ADRDA, Hachinski Ischemic<br>Score for VaD  | For people with VaD, intake of two NSAID's and of<br>were not significantly associated with VaD<br>(Data NR) | SAMPLE: Yes<br>GENDER: Yes<br>MCC: Yes<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Design NR<br><br>Cross-sectional | Living With family:<br>AD 46.3%, VaD 47.9%, no<br>Dem 41.3%<br>Nursing Home: AD 2.0%,<br>VaD 4.3%, no CI 4.7%<br><br>Intl: Ankara, Turkey<br><br>Database NR | Clinical Exam/Diagnosis,<br>Lab exams:<br>A comprehensive geriatric<br>assessment (CGA) was<br>conducted. "Normal": vitamin B12 160<br>pg/ml and over; total cholesterol 200<br>mg/dl and lower; triglycerides 200<br>mg/dl and lower; LDL-C 130 mg/dl<br>and lower, HDL-C 40 mg/dl and over | 21% with dementia<br>AD: 203 ,VD: 73<br><br>Global Deterioration Scale (GLDS)<br>was used to stage severity of<br>dementia:<br>60% mild (GLDS stage 3), 30%<br>moderate (stage 4), and 10% severe<br>(stage 6). |  |   |

SIP-37 Cognitive Impairment and Co-Occurring Chronic Conditions: Evidence Table

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|---|--|--|--|--|--|
| 28. Rastas<br>2007<br>#897  | N=553<br>All 85+<br>% female:<br>Dem: 81.3 / No Dem:79.5   | Heart failure (61%), HTN (25%),<br>Diabetes (20%), Stroke (21%),<br>AF (22%), MI (14%)   | Measures: MMSE, SPMSQ<br><br>Criteria: DSM-III-R, CDR  | # (%) Warfarin Use:<br>Dem: 2 (0.9)<br>No Dem: 9 (2.7)<br>p=NS | SAMPLE: Yes<br>GENDER: Yes<br>MCC: No<br>DEM: Yes<br>CC: Yes<br>MULTI: No<br>COV ASE: No |
| Prospective,<br>longitudinal,<br>population based<br>study<br><br>9 years | Mean (SD) yrs education:<br>Dem: 3.9(3.0)<br>No Dem: 4.2(2.9)<br><br>Intl: Vantaa, Finland<br><br>Vantaa 85+ Study | Clinical Exam/Diagnosis,<br>Records Review:<br>The evaluation included an interview<br>of a participant and a knowledgeable<br>informant by a trained nurse and a<br>clinical examination by a physician.<br>Information concerning health, health-<br>related behavior, medical history,<br>including all the illnesses and<br>medication, was obtained from an<br>electronic primary health care<br>database that contains all primary<br>care health records. The diagnosis of<br>clinical stroke was based on the<br>history of previous transient ischemic<br>attack or stroke in the medical records<br>and the presence of clinical<br>neurological focal signs indicating<br>previous stroke examined by a<br>neurologist. We also included in the<br>stroke group 27 subjects without a<br>history of cerebrovascular disease<br>who had focal signs indicating stroke.<br>The diagnosis of AF was made if 12-<br>lead ECG at rest or a short Holter<br>ECG monitoring during the exam<br>showed AF. Because these may fail to<br>detect paroxysmal AF, individuals with<br>a history of chronic AF in the health<br>records were included. | 38.7% with dementia<br><br>Mean MMSE (SD):<br>Dem: 8.3 (7.2)<br>No Dem: 23.2 (4.8)<br><br>Cognitive impairment was assessed<br>during the same evaluation and<br>electronic primary health care<br>records database as was used to<br>assess chronic conditions. The<br>diagnosis of dementia according to<br>the DSM-III R criteria was based on<br>data collected during the study: the<br>neurologist's clinical examination,<br>the MMSE and SPMSQ tests, and the<br>CDR, ADL and IADL. Besides the<br>subject, also the relatives, nurses,<br>and other persons taking care of the<br>subject were interviewed. Medical<br>history was also available. The<br>duration of the cognitive symptoms<br>had to be at least 3 months to<br>exclude, eg, delirium. The<br>consensus of 2 neurologists was<br>needed for the dementia diagnosis. |  |  |