

Supplementary Data

Review and Meta-Analysis of Biomarkers and Diagnostic Imaging in Alzheimer's Disease

Lisa M. Bloudek^a, D. Eldon Spackman^b, Michael Blankenburg^c and Sean D. Sullivan^{d,*}

^aAllergan, Inc., Irvine, CA, USA

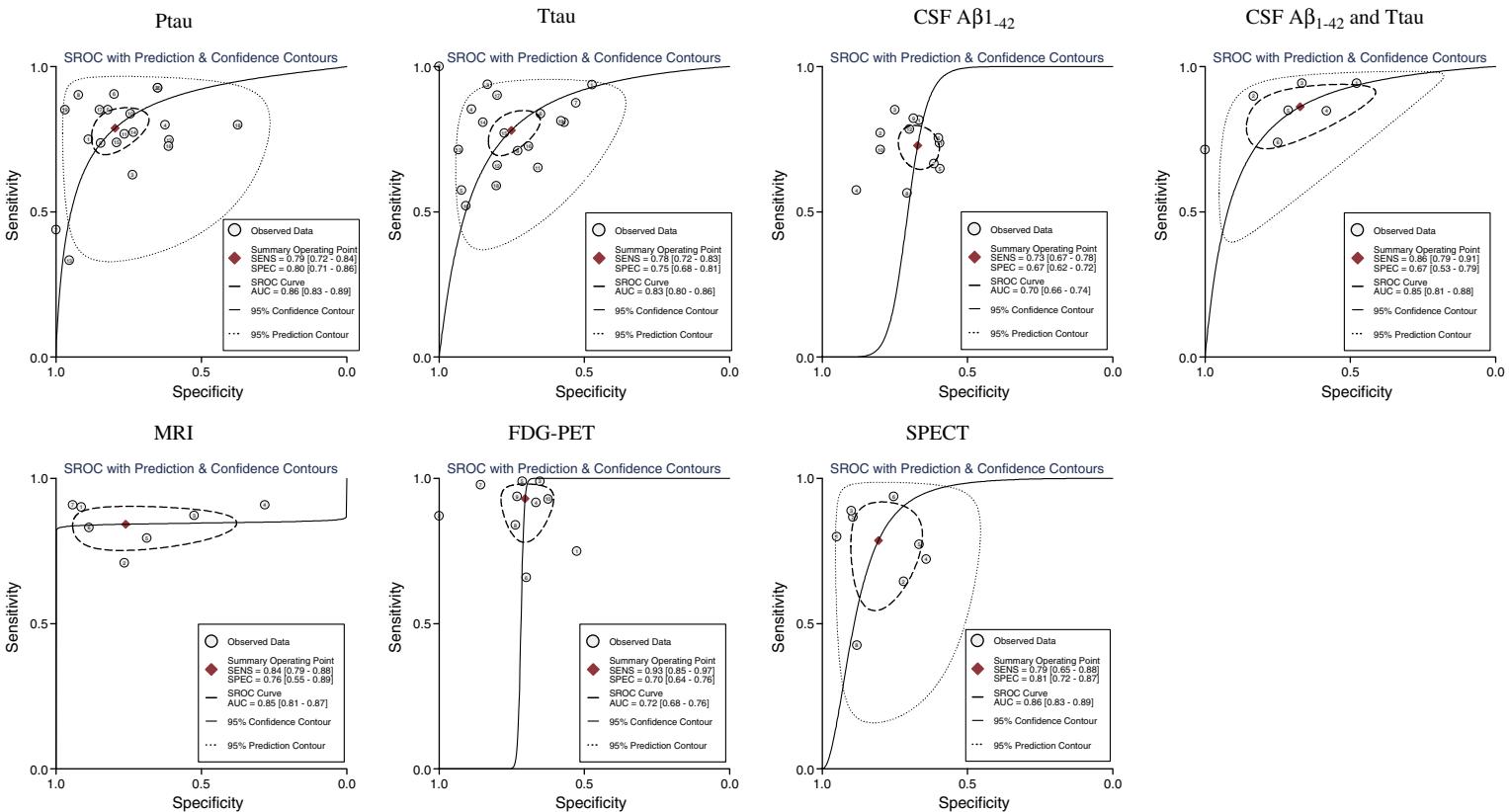
^bUniversity of York, Centre for Health Economics, York, United Kingdom

^cBayer HealthCare Pharma, Berlin, Germany

^dUniversity of Washington, Pharmaceutical Outcomes Research and Policy Program, Seattle, WA, USA

Accepted 6 May 2011

*Correspondence to: Sean D. Sullivan, Ph.D., Professor and Director, Pharmaceutical Outcomes Research and Policy Program, University of Washington, 1959 NE Pacific Ave, H-375Q, Box 357630, Seattle, WA 98195-7630, USA. Tel.: +1 206 685-8153; Fax: +1 206.543.3835; E-mail: sdsull@u.washington.edu.



Supplementary Table
Method of Establishing True AD Diagnosis for All Included Studies

| Author (Year) | Method | Comparator Group | True Diagnosis | Other Notes |
|--------------------|----------------------------------------------------------------------------------------------------------------|-------------------|-----------------------|-------------------------------------------------------------------------|
| Mulder (2010) | CSF A β _{1–42} , Ttau, Ptau ₁₈₁ | ND | NINCDS-ADRDA criteria | |
| Brys (2009) | CSF Ttau, Ptau ₂₃₁ | MCI | Clinical diagnosis | Predicted 2 yr conversion of MCI to AD. Multiple MTL measures presented |
| Brys (2009) | MRI of MTA, Ptau ₂₃₁ , Ttau | MCI | NINCDS-ADRDA criteria | Predicted 2 yr conversion of MCI to AD |
| Burton (2009) | MRI of MTA | NADD | Autopsy | |
| Frisoni (2009) | MRI of MTA, FDG-PET, CSF A β _{1–42} | MCI | Clinical diagnosis | |
| Haense (2009) | FDG-PET | ND | NINCDS-ADRDA criteria | |
| Koopman (2009) | CSF A β _{1–42} , Ttau, and Ptau | NADD | Autopsy | ADNI cohort |
| Mattsson (2009) | CSF A β _{1–42} , Ttau, and Ptau | MCI | Clinical diagnosis | |
| McEvoy (2009) | Volumetric MRI of MTA | ND | NINCDS-ADRDA criteria | |
| Roher (2009) | CSF A β _{1–42} | ND, NADD | Autopsy | |
| Shaw (2009) | CSF A β _{1–42} , Ttau, and Ptau | ND | Autopsy | |
| Smach (2009) | CSF A β _{1–42} , Ttau, A β _{1–42} + Ttau, | ND, NADD | NINCDS-ADRDA criteria | |
| Tapiola (2009) | CSF A β _{1–42} , Ttau, A β _{1–42} + Ttau, Ptau | NADD | Autopsy | |
| Vemuri (2009) | MRI of MTA, CSF Ptau ₂₃₁ , A β _{1–42} , Ttau, A β _{1–42} + Ttau | ND | Clinical diagnosis | ADNI cohort |
| Boban (2008) | CSF Ttau, Ptau ₁₈₁ , Ptau ₁₉₉ | VD | Clinical diagnosis | |
| Brandt (2008) | CSF A β _{1–42} , Ttau, Ptau ₁₈₁ | MCI, NADD, DEP | NINCDS-ADRDA criteria | |
| Colliot (2008) | Volumetric MRI of MTA | MCI | NINCDS-ADRDA criteria | Automated volumetric MRI scans |
| Duara (2008) | MRA of MTA | MCI | NINCDS-ADRDA criteria | Conversion of MCI to AD at 1-year |
| Engelborghs (2008) | CSF A β _{1–42} , Ttau, and Ptau | ND, NADD | Autopsy | |
| Habeck (2008) | FDG-PET | ND | Clinical diagnosis | |
| Lewczuk (2008) | CSF A β _{1–42} , Ttau, Ptau ₁₈₁ | NADD | NINCDS-ADRDA criteria | Also presents analysis to discriminate MCI-AD from MCI-NADD |
| McMurtray (2008) | FDG-PET | ND | NINCDS-ADRDA criteria | |
| Mosconi (2008) | FDG-PET | ND, MCI, FTD, DLB | NINCDS-ADRDA criteria | |
| Ravaglia (2008) | CSF Ptau ₁₈₁ | ND, VD | NINCDS-ADRDA criteria | |
| Barkhof (2007) | MRI of MTA | NADD | DSM-IIIR | Post-mortem MRI of Patients >85 years old |
| Ewers (2007) | CSF Ptau ₁₈₁ | MCI | Clinical diagnosis | Longitudinal. Multiple sites presented separately |
| Foster (2007) | FDG-PET | FTD | Autopsy | |
| Jagust (2007) | Clinical diagnosis, FDG-PET | NADD, ND | Autopsy | Result from final evaluation used in analysis |
| Matsunari (2007) | FDG-PET, MRI of MTA | ND | Clinical diagnosis | |
| McNeill (2007) | HMPAO SPECT | FTD | Autopsy | |
| Nihashi (2007) | FDG-PET | ND | NINCDS-ADRDA criteria | |
| Wada-Isoe (2007) | CSF Ptau | ND | NINCDS-ADRDA criteria | |
| Bonte (2006) | FDG-PET | NADD | Autopsy | |
| Ibach (2006) | CSF A β _{1–42} , Ttau, A β _{1–42} + Ttau, Ptau ₁₈₁ | ND, NADD | Clinical diagnosis | |

Supplementary Table
(Continued)

| Author (Year) | Method | Comparator Group | True Diagnosis | Other Notes |
|------------------------|------------------------------------------------------------------------|------------------|--------------------------------------|------------------------------------------------|
| Vanderstichele (2006) | CSF A β ₁₋₄₂ , Ttau, Ptau ₁₈₁ | ND, DLB | Clinical diagnosis | |
| Anchisi (2005) | FDG-PET | MCI | NINCDS-ADRDA criteria | Predicts conversion of MCI to AD |
| Brettschneider (2005) | CSF A β ₁₋₄₂ | ND | NINCDS-ADRDA criteria | |
| Drzezga (2005) | FDG-PET | MCI | NINCDS-ADRDA criteria | Predicts conversion of MCI to AD |
| Ivanoiu (2005) | CSF A β ₁₋₄₂ , Ttau, A β ₁₋₄₂ + Ttau | MCI, DEP | Clinical diagnosis | |
| Hanyu (2005) | MTR of MTA | DLB | NINCDS-ADRDA criteria | |
| Herukka (2005) | CSF A β ₁₋₄₂ , A β ₁₋₄₂ + Ttau | ND, MCI | Undetermined | |
| Kapaki (2005) | CSF Ttau, A β ₁₋₄₂ , A β ₁₋₄₂ + Ttau | ND, ARCD | NINCDS-ADRDA criteria | |
| Stefani (2005) | CSF A β ₁₋₄₂ , Ttau, and Ptau | VD | NINCDS-ADRDA criteria | |
| Bonte (2004) | HMPAO SPECT | FTD | Mixed Clinical diagnosis and Autopsy | |
| Hampel (2004) | CSF A β ₁₋₄₂ , Ttau, Ptau | MCI | NINCDS-ADRDA criteria | Predicts conversion of MCI to AD |
| Rossi (2004) | CT of MTA | ND | Autopsy | |
| Schoonenboom (2004) | CSF A β ₁₋₄₂ , Ttau, Ptau ₁₈₁ | ND, FTD | NINCDS-ADRDA criteria | |
| Wang (2004) | MRI of MTA | ND, VD | NINCDS-ADRDA criteria | |
| Clark (2003) | CSF Ttau, A β ₁₋₄₂ + Ttau | ND, NADD | Clinical guidelines | |
| Buerger (2003) | CSF Ptau ₂₃₁ | DEP | NINCDS-ADRDA criteria | |
| Kapaki (2003) | CSF Ttau, A β ₁₋₄₂ , A β ₁₋₄₂ + Ttau | ND, NADD, VD | NINCDS-ADRDA criteria | |
| Maddalena (2003) | CSF A β ₁₋₄₂ + Ptau | ND, NADD | Clinical diagnosis | |
| Schonknect (2003) | CSF Ttau, Ptau ₁₈₁ | ND | Undetermined | Abstract only |
| Sunderland (2003) | CSF A β ₁₋₄₂ + Ttau | VD | Mixed Clinical diagnosis and Autopsy | |
| Bottino (2002) | MRI of MTA | ND, MCI | NINCDS-ADRDA criteria | |
| Buerger (2002) | CSF Ttau, Ptau ₂₃₁ | ND, FTD, DLB, VD | NINCDS-ADRDA criteria | |
| Herholz (2002) | FDG-PET | ND | NINCDS-ADRDA criteria | |
| Frisoni (2002) | CT and MRI of MTA | ND | Clinical diagnosis | |
| Riemenschneider (2002) | CSF A β ₁₋₄₂ + Ttau | MCI | NINCDS-ADRDA criteria | Predicts conversion of MCI to AD |
| Shoji (2002) | CSF Ttau | ND, NADD | NINCDS-ADRDA criteria | |
| Varma (2002) | MRI of MTA, HMPAO SPECT | NADD | NINCDS-ADRDA criteria | |
| Andreasen (2001) | CSF A β ₁₋₄₂ + Ttau | ND | NINCDS-ADRDA criteria | |
| Arnaiz (2001) | FDG-PET | MCI | NINCDS-ADRDA criteria | Predict conversion of MCI to AD |
| Hanyu (2001) | MRI of MTA | ND | NINCDS-ADRDA criteria | |
| Itoh (2001) | CSF Ptau ₁₉₉ | NADD | NINCDS-ADRDA criteria | |
| O'Brien (2001) | MRI and SPECT | ND | NINCDS-ADRDA criteria | |
| Parnetti (2001) | CSF Ttau, A β ₁₋₄₂ , Ptau | ND, DLB | Clinical diagnosis | AB42 results not significant and not presented |
| Silverman (2001) | FDG-PET | NADD | Mixed Clinical diagnosis and Autopsy | |
| Sjogren (2001) | CSF Ttau, Ptau | ND, FTD, PD | NINCDS-ADRDA criteria | |
| Denihan (2000) | CT of MTA | NADD | NINCDS-ADRDA criteria | |
| Higuchi (2000) | FDG-PET | ND, DLB | NINCDS-ADRDA criteria | |
| Hoffman (2000) | NINCDS-ADRDA criteria | NADD | Autopsy | |
| Kahle (2000) | CSF Ttau | ND | Mixed clinical diagnosis and Autopsy | |
| Killiany (2000) | MRI of MTA | ND, MCI | NINCDS-ADRDA criteria | Predicts conversion of MCI to AD |
| Kohnken (2000) | CSF Ptau ₂₃₁ | NADD | NINCDS-ADRDA criteria | |
| O'Brien (2000) | CT of MTA | DEP, DLB, VD | NINCDS-ADRDA criteria | |
| Ohyama (2000) | FDG-PET | ND | Clinical diagnosis | Abstract only |
| Andreasen (1999) | CSF Ttau | ND | Clinical diagnosis | |

Supplementary Table
(Continued)

| Author (Year) | Method | Comparator Group | True Diagnosis | Other Notes |
|--------------------|-------------------------------------------------------------------------------|------------------|--------------------------------------|-----------------------------------|
| Golebiowski (1999) | MRI of MTA | ND | Undetermined | |
| Defebvre (1999) | HMPAO SPECT | PD, DLB | NINCDS-ADRDA criteria | |
| Lim (1999) | NINCDS-ADRDA criteria | NADD | Autopsy | |
| Holmes (1999) | NINCDS-ADRDA criteria | NADD | Autopsy | |
| Hulstaert (1999) | CSF A β_{1-42} , Ttau | ND, NADD | NINCDS-ADRDA criteria | |
| Jobst (1998) | NINCDS-ADRDA criteria, DSM-IV criteria, SPECT, CT of MTA | ND | Autopsy | |
| Kurz (1998) | CSF Ttau | ND | Clinical diagnosis | |
| Laakso (1998) | MRI of MTA | ND | NINCDS-ADRDA criteria | |
| Mayeux (1998) | Clinical diagnosis by NINCDS-ADRDA, DSM-III, or Cummings and Benson criteria. | NADD | Autopsy | |
| Pucci (1998) | MRI of MTA | ND | Clinical diagnosis | |
| Shoji (1998) | CSF Ttau, A β_{1-40} , A β_{1-42} | ND | NINCDS-ADRDA criteria | |
| Arai (1997) | CSF Ttau | ND, NADD | Clinical diagnosis | |
| Bonte (1997) | SPECT | ND, NADD | Mixed Clinical diagnosis and Autopsy | |
| Ishii (1998) | FDG-PET | ND, DLB | Clinical diagnosis | |
| Pasquier (1997) | CT of MTA | NADD | Clinical diagnosis | |
| O'Brien (1997) | MRI of MTA | ND | Clinical diagnosis | |
| Scheltens (1997) | MRI of MTA, SPECT | ND | Clinical diagnosis | |
| Burdette (1996) | FDG-PET | ND | Undetermined | |
| Frisoni, (1996) | MRI of MTA and rWTH | ND | Clinical diagnosis | |
| Greene (1996) | HMPAO SPECT | ND | NINCDS-ADRDA criteria | |
| Van Dyke (1996) | ECD SPECT and HMPAO SPECT | ND | Clinical diagnosis | |
| Van Gool (1995) | HMPAO SPECT | ND | McKhann criteria | |
| Blacker (1994) | NINCDS-ADRDA criteria | NADD | Autopsy | Post-consensus SN and SP |
| Claus (1994) | HMPAO SPECT | ND | NINCDS-ADRDA criteria | |
| Desmond (1994) | MRI of MTA | ND | NINCDS-ADRDA criteria | |
| Messa (1994) | FDG-PET | ND | Clinical diagnosis | |
| McMurdo (1994) | HMPAO SPECT | ND | NINCDS-ADRDA criteria | |
| Szelies (1994) | FDG-PET | ND, VD | Clinical diagnosis | |
| Azari (1993) | FDG-PET | ND | Undetermined | |
| Bonte (1993) | HMPAO SPECT | NADD | Mixed Clinical diagnosis and Autopsy | |
| Hanyu (1993) | SPECT | NADD | Undetermined | |
| Herholz (1993) | FDG-PET | ND | NINCDS-ADRDA criteria | |
| Johnson (1993) | SPECT | ND | NINCDS-ADRDA criteria | |
| Erkinjuntti (1993) | MRI of MTA | ND | NINCDS-ADRDA criteria | Results for hippocampal formation |
| Kippenhan (1992) | FDG-PET | ND | NINCDS-ADRDA criteria | |
| Powers (1992) | PET | ND | Undetermined | |
| Scheltens (1992) | MRI of MTA | ND | Clinical diagnosis | |
| Grady (1990) | FDG-PET | ND | Undetermined | |
| Herholz (1990) | FDG-PET | ND | NINCDS-ADRDA criteria | Abstract Only |
| Johnson (1990) | IMP SPECT | ND | Undetermined | Abstract Only |