

NEUROPSYCHIATRY AND BEHAVIORAL NEUROLOGY

Predictors of Cognitive Decline in Individuals with Subjective Cognitive Decline of the DELCODE Study Cohort

Lena Sannemann¹ | Katharina Buerger^{2,3} | Julian Hellmann-Regen^{4,5,6} |
 Luca Kleineidam^{7,8} | Christoph Laske^{9,10} | Robert Perneczky^{3,11,12,13} |
 Oliver Peters^{4,5,6} | Josef Priller^{4,14,15,16} | Alfredo Ramirez^{7,8,17,18,19} |
 Anja Schneider^{7,8} | Annika Spottke^{8,20} | Matthias Synofzik^{21,22} | Stefan J. Teipel^{23,24} |
 Michael Wagner^{8,25} | Jens Wiltfang^{26,27,28} | Renat Yakupov^{29,30} | Emrah Düzel^{29,31} |
 Frank Jessen^{1,8,19} | the DELCODE Study Group

¹University of Cologne, Faculty of Medicine and University Hospital Cologne, Department of Psychiatry, Cologne, Germany

²Institute for Stroke and Dementia Research (ISD), University Hospital, LMU, Munich, Germany

³German Center for Neurodegenerative Diseases (DZNE), Munich, Germany

⁴German Center for Neurodegenerative Diseases (DZNE), Berlin, Germany

⁵Charité – Universitätsmedizin Berlin, corporate member of Freie Universität Berlin and Humboldt-Universität zu Berlin – Institute of Psychiatry and Psychotherapy, Berlin, Germany

⁶ECRC Experimental and Clinical Research Center, Charité Universitätsmedizin Berlin, Berlin, Germany

⁷Department of Old Age Psychiatry and Cognitive Disorders, University Hospital Bonn and University of Bonn, Bonn, Germany

⁸German Center for Neurodegenerative Diseases (DZNE), Bonn, Germany

⁹German Center for Neurodegenerative Diseases (DZNE), Tuebingen, Germany

¹⁰Section for Dementia Research, Hertie Institute for Clinical Brain Research and Department of Psychiatry and Psychotherapy, University of Tuebingen, Tuebingen, Germany

¹¹Ageing Epidemiology (AGE) Research Unit, School of Public Health, Imperial College London, London, United Kingdom

¹²Munich Cluster for Systems Neurology (SyNergy), Munich, Germany

¹³Department of Psychiatry and Psychotherapy, University Hospital, LMU Munich, Munich, Germany

¹⁴Department of Psychiatry and Psychotherapy, School of Medicine and Health, Technical University of Munich, and German Center for Mental Health (DZPG), Munich, Germany

¹⁵University of Edinburgh and UK DRI, Edinburgh, United Kingdom

¹⁶Department of Psychiatry and Psychotherapy, Charité, Charitéplatz 1, Berlin, Germany

¹⁷Division of Neurogenetics and Molecular Psychiatry, Department of Psychiatry and Psychotherapy, Faculty of Medicine and University Hospital Cologne, University of Cologne, Cologne, Germany

¹⁸Department of Psychiatry and Glenn Biggs Institute for Alzheimer's and Neurodegenerative Diseases, San Antonio, TX, USA

¹⁹Excellence Cluster on Cellular Stress Responses in Aging-Associated Diseases (CECAD), University of Cologne, Cologne, Germany

²⁰Department of Neurology, University of Bonn, Bonn, Germany

²¹German Center for Neurodegenerative Diseases (DZNE), Tübingen, Germany

²²Division of Translational Genomics of Neurodegenerative Diseases, Hertie Institute for Clinical Brain Research and Center of Neurology, University of Tübingen, Tübingen, Germany

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²³Department of Psychosomatic Medicine, Rostock University Medical Center, Rostock, Germany

²⁴German Center for Neurodegenerative Diseases (DZNE), Rostock, Germany

²⁵Department for Cognitive Disorders and Old Age Psychiatry, University Hospital Bonn, Bonn, Germany

²⁶Department of Psychiatry and Psychotherapy, University Medical Center Goettingen, University of Goettingen, Goettingen, Germany

²⁷Neurosciences and Signaling Group, Institute of Biomedicine (iBiMED), Department of Medical Sciences, University of Aveiro, Aveiro, Portugal

²⁸German Center for Neurodegenerative Diseases (DZNE), Goettingen, Germany

²⁹German Center for Neurodegenerative Diseases (DZNE), Magdeburg, Germany

³⁰Institute of Cognitive Neurology and Dementia Research (IKND), Otto-von-Guericke University, Magdeburg, Germany

³¹Institute of Cognitive Neurology and Dementia Research (IKND), Otto-von-Guericke University, Magdeburg, Sachsen Anhalt, Germany

Correspondence

Lena Sannemann, University of Cologne,
Faculty of Medicine and University Hospital
Cologne, Department of Psychiatry, Cologne,
Germany.

Email: lena.sannemann@uk-koeln.de

Abstract

Background: Subjective cognitive decline (SCD) refers to a self-perceived, persistent cognitive decline compared to previous levels in individuals with objectively unimpaired cognition. Studies have repeatedly shown associations of SCD characteristics with amyloid pathology and increased risk of future cognitive decline, especially in memory-clinic settings. The aim of this project is to model individual differences of cognitive decline in individuals with SCD.

Method: Latent Growth Curve Model (LGCM) analysis was applied to individuals with SCD from the DZNE Longitudinal Cognitive Impairment and Dementia (DELCODE) study. We chose a sample of $n = 203$ participants who showed a decline on the Preclinical Alzheimer's Cognitive Composite (PACC5) score over five years. First, a two-factor linear growth model was fitted on the annualized PACC5 data. We then calculated and compared two models to which we added the following baseline predictors: 1) plasma A β 42/40, plasma ptau181, ApoE-4-carrier status and hippocampal volume (biological model), and 2) Geriatric Depression Scale (GDS), Geriatric Anxiety Inventory–Short Form (GAIS-SF) and Neuropsychiatric Inventory Questionnaire (NPI-Q) total scores (neuropsychiatric model).

Result: The LGCM of longitudinal PACC-5 scores yielded adequate model fit for a linear model ($X^2(16)=67.5$, $p < .001$, CFI = 0.93, SMRM=0.07, AIC=1437.10). The baseline PACC score was -0.03 ($SE = 0.05$, $p = .533$) and average cognition declined slightly over time by -0.13 ($SE = 0.01$, $p < .001$). The biological model showed an improvement in fit, with an AIC of 742.30. Here, we observed a positive relationship between plasma A β 42/40 and the intercept ($B = 7.60$, $SE = 3.01$, $p = .012$) and a negative relationship between plasma ptau181 and the intercept ($B = -0.22$, $SE = 0.09$, $p = .016$). Plasma A β 42/40 was the only significant predictor of the PACC5 slope in this model ($B = 1.35$, $SE = 0.62$, $p = .030$). In comparison, the AIC value for the neuropsychiatric model was 1341.17, with the GDS total score being negatively related to the PACC5 slope ($B = -0.03$, $SE = 0.01$, $p = .009$).

Conclusion: These results add to gaining a better understanding of SCD trajectories and specific predictors of cognitive decline, which is relevant to power future clinical trials in this population.