34th European **NEUROLOGY CONGRESS**

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DEMENTIA AND VASCULAR DEMENTIA

26th International Conference on **NEUROSURGERY AND NEUROSCIENCE**

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June 24-25, 2020

Ahmed M. Alsehli et al., J Spine Neurosurg 2020, Volume 09

The cognitive effects of statins and the role of age

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Ctatins are the first-choice treatment against hypercholesterolemia and associated cardiovascular disease, • the main global cause of morbidity and mortality according to WHO. Statins are among the most prescribed drugs worldwide with an estimated 25% of the world population older than 65 years currently under statin treatment and the numbers increasing. Currently, there is a large controversy about whether or not statins affect cognitive function. Studies have provided indications for both sides, as well as reported beneficial and detrimental effects. Altogether, findings in the literature are highly inconsistent. Thus, to reveal new insights into statin cognitive effects, we performed an observational study on a population-based sample of 245,731 control and 55,114 statin-taking individuals from the UK Biobank (Alsehli et al., Sci Rep. 2020 10, 6187). Cognitive performance in terms of reaction time, working memory and fluid intelligence was analysed at baseline and two follow-ups (within 5-10 years). Subjects were classified depending on age (up to 65 and over 65 years) and treatment duration (1-4 years, 5-10 years and over 10 years). Data were adjusted for healthand cognition-related covariates. Subjects generally improved in test performance with repeated assessment and middle-aged persons performed better than older persons. The effect of statin use differed considerably between the two age groups with a beneficial effect on reaction time in older persons and fluid intelligence in both age groups, and a negative effect on working memory in younger subjects. Our analysis suggests a modulatory impact of age on the cognitive side effects of statins, revealing a possible reason for profoundly inconsistent findings on statin-related cognitive effects in the literature. The study highlights the importance of characterising modifiers of statin effects to improve knowledge and shape guidelines for clinicians when prescribing statins and evaluating their side effects in patients.