

## 1.2 Chronological Versus Vascular Age: Predictive Value for Cardiovascular Events and Identification of Patients with SUPERNormal Vascular Aging (SUPERNOVA)

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### ABSTRACT

**Background:** Individuals whose arteries appears abnormally healthy in comparison to their CV risk factor burden (supernormal vascular aging, SUPERNOVA) represent an ideal model to discover novel pathways of cardiovascular protection. Purpose: 1) to introduce a novel definition of vascular age, as predicted age based on CV risk factors and PWV; 2) to test the hypothesis that individuals with the largest difference between chronological and vascular age (C-Vage) show a lower rate of CV events, and may thus be defined as SUPERNOVA.

**Methods:** The best fitting model for age prediction was investigated in the multicenter, European, cross-sectional Reference Values for Arterial stiffness Collaboration Database ( $n = 11406$ ). A survival analysis on the longitudinal cohort of the Malmo Diet and Cancer Study (MDCS,  $n = 2663$ ) was then performed. The main outcome was a composite endpoint of fatal and non-fatal major cardiac events and strokes.

**Results:** In the Reference Values Cohort, the model explained 59.8% of age variance, with PWV being the most relevant predictor ( $r^2: 0.105$ ). The model (derived on population  $>61$  years) was used to calculate vascular age in the MDCS Cohort. In the Cox survival analysis (follow-up 6.6 years, 286 events), C-Vage was significantly and inversely associated with CV events (Figure 1). Individuals in the highest C-Vage decile ( $>8.8$  years) were defined as SUPERNOVA. In the competing risk analysis, SUPERNOVA had a reduced age-adjusted rate of CV events [HR 0.51 (0.34–0.76)].

**Conclusion:** This proof-of-concept study demonstrated that SUPERNOVA individuals have a disproportionately lower rate of CV events. Future studies are needed to explore genetic/molecular mechanisms for this phenotype.

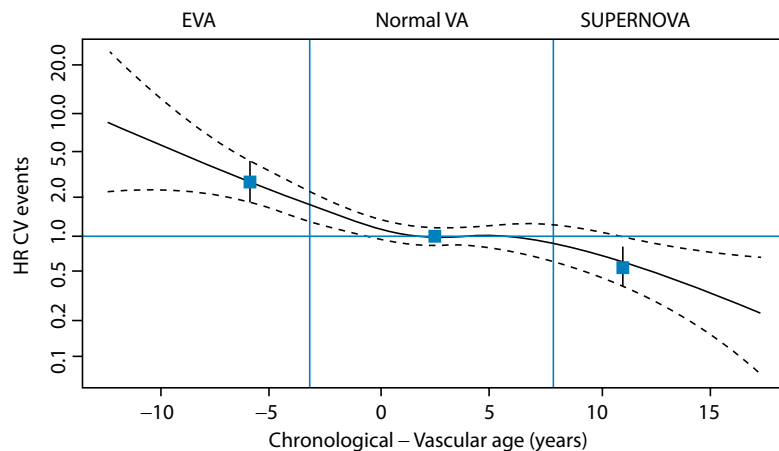


Figure 1

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