

World Trade Center neurotoxic exposures are associated with elevated plasma amyloid, total-tau and neurofilament light in responders

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Abstract

Background: The collapse of the World Trade Center towers on September 11th 2001 resulted in a 16-acre environmental toxic disaster. More than 1.2 million tons of construction material and carcinogens including polycyclic aromatic hydrocarbons, gypsum and metals coalesced, resulting in a highly alkaline dust cloud. Police and Law enforcement were among the most highly exposed group.

Method: In this retrospective cohort study, we included a subset of high exposure Responders (n=424) with cross sectional plasma samples of amyloid β_{40} , amyloid β_{42} , total-tau, neurofilament light and a baseline evaluation of cognitive functioning assessed with the Montreal Cognitive Assessment (MoCA) to examine long-term associations between WTC neurotoxic exposures (e.g. diesel exhaust, chemicals) with levels of proteins associated with neuropathological characteristics of Alzheimer's disease and neurodegeneration. Spearman rho p values adjusted for multiple comparisons using the false discovery rate (FDR=0.05) examined associations with participant characteristics and plasma concentrations. Multivariate regressions ascertained independent effects of WTC neurotoxic exposures in predicting plasma biomarker concentrations.

Result: Responders were on average 54.3 years at blood draw. Worse performance on the baseline MoCA was associated with higher levels of $A\beta_{40}$. Plasma $A\beta_{40}$ and NfL were inversely correlated with dust exposure, $A\beta_{42}$ and ratio $A\beta_{42-40}$ were inversely correlated with total hours on site during 9/11-9/14 and working in enclosed work areas was associated with higher concentrations of $A\beta_{40}$ and lower concentrations of ratio $A\beta_{42-40}$. Diesel exhaust exposure predicted levels of $A\beta_{40}$, total tau and NfL whereas early exposure predicted $A\beta_{42}$ concentrations and dust exposure predicted ratio $A\beta_{42-40}$.

Conclusion: Differences across inhaled neurotoxins and time of arrival may have differential long-term effects on blood-based protein biomarkers of neuropathology and brain health.