



Apolipoprotein A1 (ApoA1) and Epidermal Growth Factor (EGF) as Plasma-based Biomarkers in Parkinson's Disease

Christine Swanson, PhD, and Alice Chen-Plotkin, MD.
University of Pennsylvania, Department of Neurology

Summary (or Abstract)

Suggested 200 words maximum

We assayed plasma from 100 normal controls as well as 154 PD individuals from the PPMI study at 3 timepoints (baseline, 6 mo, 12 mo) for epidermal growth factor (EGF) and Apolipoprotein A1 (ApoA1). EGF was measured using an enzyme-linked immunosorbent assay (ELISA) routinely used in our lab, while ApoA1 was measured using a clinical immunoturbidimetric assay. In the past, we have seen that low EGF levels correlate with poorer cognitive performance in PD patients (Chen-Plotkin et al., 2011) -- we sought to determine this also holds true in PPMI subjects. We have also previously observed that higher ApoA1 levels have a "protective effect" in PD -- correlating with increased age at disease onset and better levels of dopaminergic system integrity (Qiang et al., 2013). We evaluated if PPMI PD subjects with higher ApoA1 levels have a more benign course of disease.

Methods

Plasma levels of EGF were measured by enzyme-linked immunosorbent assay (ELISA; R&D Systems, Minneapolis, MN, Catalog #DEG00) as previously described (Chen-Plotkin et al., 2011) according to manufacturer instructions. Samples were run in duplicate and data used for this study met quality control measures for technical performance (coefficient of variation (CV) < 0.20 for replicates). In our hands, this EGF ELISA can be expected to yield values that are robust across plates/days/operators. To minimize variability, however, all samples from each subject were processed on one lot of ELISAs, with minimal time between runs, which all occurred within 2 weeks. Operators were blinded to disease status during ELISA measurement. A quality-control reference sample (generated by pooling plasma from multiple samples into a large batch that was then aliquoted into many identical samples) was run in duplicate on each plate, to evaluate plate-to-plate variation, which was minimal (CV=0.07 across 39 separate measurements – one plate had only one reference sample).

Plasma levels of ApoA1 and a basic lipid panel (comprised of HDL, LDL, total cholesterol, and triglycerides) were measured using a Roche Cobas c501 automated biochemical analyzer (Tina-quant assay, catalog number 03032566-122). Samples were run in five batches, by the same operator, who was blinded to disease status. Two quality-control pooled plasma samples were run in duplicate per batch, to test for batch-to-batch variation, which was minimal (CV = 0.03 across 10 measurements).





Samples with $CV > 0.2$ across duplicate spots are flagged as “high-CV” samples. These datapoints should be interpreted with caution. In addition, samples whose measures were lower than the lowest measurement on the standards used for each assay are flagged as “low-value” samples.

References

Chen-Plotkin, A.S., Hu, W.T., Siderowf, A., Weintraub, D., Goldmann Gross, R., Hurtig, H.I., Xie, S.X., Arnold, S.E., Grossmann, M., Clark, C.M., Shaw, L.M., McCluskey, L., Elman, L., Van Deerlin, V.M., Lee, V.M., Soares, H., Trojanowski, J.Q. (2011). *Plasma epidermal growth factor levels predict cognitive decline in Parkinson disease*. Ann Neurol. 69(4):655-63.

Qiang, J.K., Wong, Y.C., Siderowf, A., Hurtig, H.I., Xie, S.X., Lee, V.M., Trojanowski, J.Q., Yearout, D., Leverenz, J., Montine, T.J., Stern, M., Mendick, S., Jennings, D., Zabetian, C., Marek, K., Chen-Plotkin, A.S. (2013). *Plasma apolipoprotein A1 as a biomarker for parkinson's disease*. Ann Neurol. 74(1): 119-27.

About the Authors

This document was prepared by **Christine R. Swanson and Alice Chen-Plotkin, University of Pennsylvania, Department of Neurology**. For more information please contact **Alice Chen-Plotkin** at (215) 573-7193 or by email at chenplot@mail.med.upenn.edu.

Notice: This document is presented by the author(s) as a service to PPMI data users. However, users should be aware that no formal review process has vetted this document and that PPMI cannot guarantee the accuracy or utility of this document.

