25- Hydroxyvitamin D is Inversely Associated with Serum MMP-9 Concentration in ESRD Patients H Wasse**, F Cardarelli**, C Hooper*, E Veledar**, I Guessous*

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Background

- Chronic inflammation increases the risk of cardiovascular disease and death in ESRD patients
- Studies suggest an inverse association between circulating 25(OH)D and inflammatory markers in general population
- ESRD patients have high prevalence of vitamin
 D deficiency
 - Vitamin D deficiency increases risk of and cardiovascular disease
 - Mechanisms are not known
- We tested whether circulating 25(OH)D concentration is associated with serum inflammatory biomarkers in cross-sectional pilot study of prevalent ESRD patients

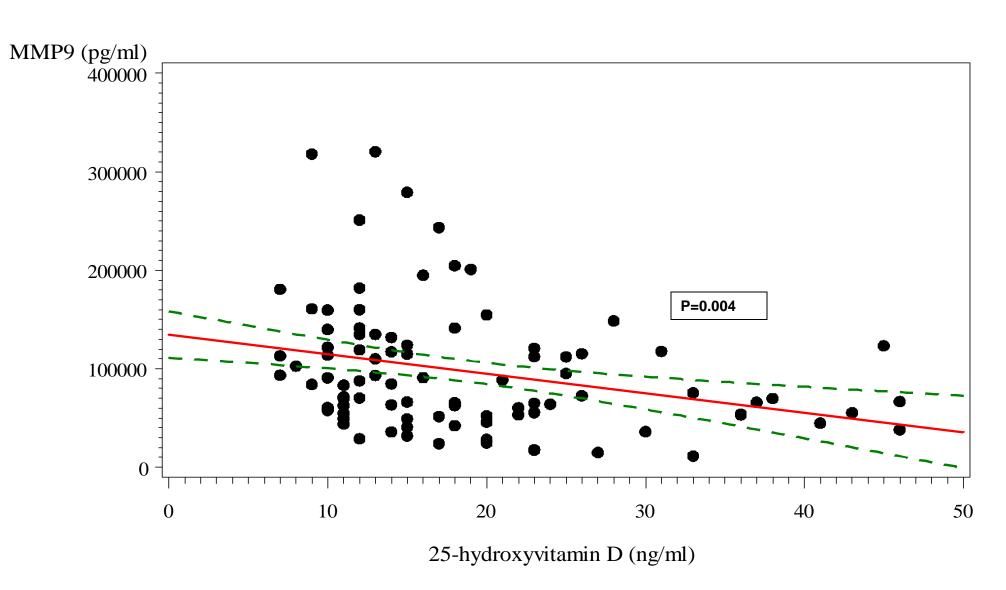
Methods

- Prevalent African-American ESRD patients in 6
 Emory University—affiliated Davita dialysis units,
 9/01/06 to 11/30/08
- Cross-sectional study
- Exclusion criteria
 - Known malignancy, active vasculitis
 - Evidence current infection, inflammation
 - Current steroid, calcineurin inhibitor, antimetabolite use
- Blood sample for vitamin D, inflammatory biomarkers
- Logistic regression models to examine associations between 25(OH)D concentrations and inflammation

Table 1. Characteristics of study participants by circulating 25(OH)D concentration

| Characteristic | All patients | Vit D< 15 ng/ml | Vit D≥ 15 ng/ml | |
|-------------------------|----------------------|----------------------|----------------------|---------|
| | N=91 | N=39 | N=52 | p value |
| Age (years) | 59.25 <u>+</u> 12.42 | 56.38 <u>+</u> 12.59 | 61.41 <u>+</u> 11.96 | 0.055 |
| Sex | | | | 0.275 |
| Female | 43 (47.25) | 21 (48.54) | 22 (51.16) | |
| Male | 48 (52.75) | 18 (37.50) | 30 62.50) | |
| Vitamin D level (ng/ml) | 18.78 <u>+</u> 9.60 | 11.10 <u>+</u> 1.95 | 24.53 <u>+</u> 8.99 | <0.0001 |
| | | | | |

Figure 1: Correlation between Serum 25 (OH) D and MMP-9 Concentration



Pearson correlation coefficient -0.29578

Continuous line= Regression Line; Dashed lines= 95% confidence intervals

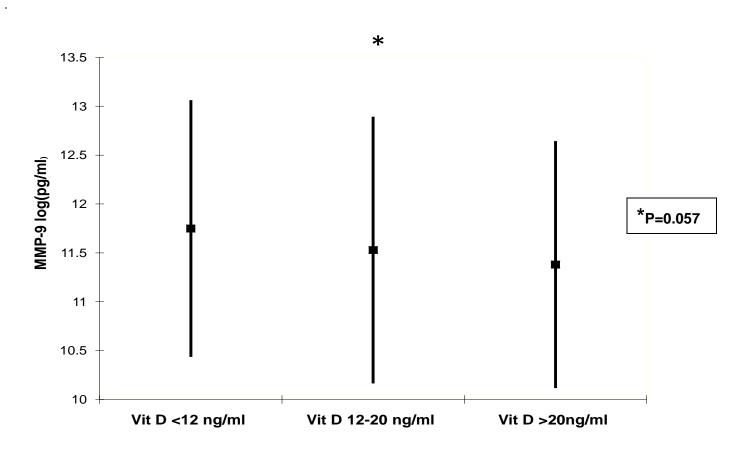
Results

Table 2 : Adjusted * associations between 25(OH)D and Inflammatory Markers

| Biomarker | Beta coefficient (SE) | P value | |
|-------------------|-----------------------|---------|--|
| | | | |
| Log MMP9 (pg/ml) | -0.017 (0.008) | 0.04 | |
| Log CRP (mg/ml) | -0.013 (0.015) | 0.39 | |
| Log IL-10 (pg/ml) | 0.022 (0.0103) | 0.04 | |
| | | | |

^{*}Adjusted for smoking, gender, BMI, age, diabetes

Figure 2: Difference in Log Concentration of MMP-9 by 25(OH) D Tertile



Conclusions

- 25 (OH) D concentration is inversely and significantly correlated with MMP-9 concentration (P value=0.004)
- MMP-9 expression is independently associated with 25(OH)D concentration in ESRD patients (P value=0.04)
- Results may help elucidate the mechanism by which 25(OH)D functions as a cardiovascular risk factor
- Future studies are needed to characterize the relationship between vitamin D therapy and inflammation

Acknowledgement: We thank the patients who participated in this study and DaVita Clinical Research® for providing partial grant funding of this research project.

