

OA-016 **PREVALENCE AND RISK FACTORS FOR EFVIRENZ-BASED ANTIRETROVIRAL TREATMENT-ASSOCIATED SEVERE VITAMIN D DEFICIENCY: A PROSPECTIVE COHORT STUDY**

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**Background** Initiation of efavirenz-based combination antiretroviral therapy (cART) is associated with Vitamin D deficiency, but the risk factors for cART-induced severe vitamin D deficiency (SVDD) and the impact of anti-tuberculosis (TB) co-treatment is not explored well.

**Methods** Treatment-naïve HIV patients with (n=102) or without (n=89) tuberculosis co-infection were enrolled prospectively and received efavirenz-based cART. In TB-HIV co-infected patients, rifampicin-based TB treatment was initiated. Plasma 25-hydroxyvitamin D (25(OH)D), cholesterol and 4-beta hydroxycholesterol concentrations were measured at baseline, and weeks 4, 16 and 48 of cART. Plasma efavirenz concentrations were determined at week 4 and 16 of cART. Genotyping for CYP2B6, CYP3A5, ABCB1, SLCO1B1, and UGT2B7 were done.

**Results** TB-HIV patients had significantly lower plasma 25 (OH)D3 levels than HIV-only patients at baseline. TB co-infection, low Karnofsky score, high viral load and high CYP3A activity as measured by plasma 4-beta hydroxycholesterol/ cholesterol ratios were significant predictors of low 25 (OH) D3 levels at baseline. In HIV-only patients, initiation of efavirenz-based cART increased the prevalence of SVDD from 27% at baseline to 76%, 79% and 43% at weeks 4, 16 and 48 of cART, respectively. The median 25(OH)D3 levels declined from baseline by -40%, -50% and -14% at weeks 4, 16 and 48 of cART, respectively. In TB-HIV patients, prior TB therapy had no influence on 25(OH)D3 levels, but the initiation of efavirenz-based cART increased the prevalence of SVDD from 57% at baseline to 70% and 72% at weeks 4 and 16 of cART, respectively. Whereas the median plasma 25(OH)D3 declined from baseline by -17% and -21% at week 4 and 16 of cART, respectively. None of the genotypes were significantly associated with SVDD.

**Conclusions** Low plasma cholesterol, high CYP3A activity, and high plasma efavirenz concentrations are significant predictors of early efavirenz-based cART-induced SVDD. Low plasma 25 (OH)D3 level at baseline is associated with TB co-infection and HIV diseases progression.