Estimate a least-squares regression.

$$Y = X_1^T \tilde{\beta}_1 + \tilde{\mu}$$

Then regress the residuals on another set of regressors.

$$\tilde{\mu} = X_2^T \tilde{\beta}_2 + \tilde{e}$$

Then regress on both set of regressors.

$$Y = X_1^T \hat{\beta}_1 + X_2^T \hat{\beta}_2 + \hat{e}$$

Is it true that $\tilde{\beta}_2 = \hat{\beta}_2$?

Proof.
$$\begin{split} &\tilde{\beta}_2=\hat{\beta}_2 \text{ if and only if } X_1 \text{ and } X_2 \text{ are uncorrelated, i.e. } X_1^TX_2=0 \\ &\text{Define } M_i=I-X_i(X_i^TX_i)^{-1}X_i^T \\ &\text{We obtain } \tilde{\mu}=M_1Y \\ &\tilde{\beta}_2=(X_2^TX_2)^{-1}X_2^T\tilde{\mu}=(X_2^TX_2)^{-1}X_2^TM_1Y \\ &\hat{\beta}_2=(X_2^TM_1X_2)^{-1}X_2^TM_1Y \\ &\text{Suppose } \tilde{\beta}_2=\hat{\beta}_2 \text{, then} \end{split}$$

$$\beta_2 = (X_2^T X_2)^{-1} X_2^T \tilde{\mu} = (X_2^T X_2)^{-1} X_2^T M_1 Y_2^T \tilde{\mu}$$

$$(X_2^T X_2)^{-1} X_2^T M_1 Y = (X_2^T M_1 X_2)^{-1} X_2^T M_1 Y$$

By algebra, we obtain $X_2^T X_2 = X_2^T M_1 X_2$

$$X_2^T X_2 = X_2^T (I - X_1 (X_1^T X_1)^{-1} X_1^T) X_2 = X_2^T X_2 - X_2^T X_1 (X_1^T X_1)^{-1} X_1^T X_2$$

Thus $X_1^T X_2 = 0$ since $(X_1^T X_1)^{-1}$ is invertible.