## Supplementary Material

**Claim:** Correlation between  $B(i)$ ,  $B(j)$ , the i<sup>th</sup> and the j<sup>th</sup> components of B is negative when i,j are alternative features, which implies the assumption that  $f(X) = f(X_{swap(i,j)})$ , and X and  $X_{swap(i,j)}$  follows the same distribution.  $X_{swap(i,j)}$  is the matrix from exchanging the i<sup>th</sup> column and j<sup>th</sup> column of X.

## Proof:

Denote X as the feature vector, a D dimensional random vector, and Y is the responding variable. First we assume the ground true model is represented by the following formula.

$$
Y = f(X)
$$

When two genetic variants  $X_i$ ,  $X_i$  are a pair of alternative genetic variants for a responding variable, we assume that exchanging their values will not change the responding variable. Formally, it is defined as

$$
f(X_{\text{swap}(i,j)}) = f(X) \tag{2}
$$

We also assume  $X_{swap(i,j)}$  and X follows the same distribution. The optimal solution of a sparse model can be written as B.

$$
B = \operatorname{argmin}_{B} [f(X) - X^T B]^2 + \lambda |B|_1.
$$
\n(3)

So the solution will satisfy the following Karush–Kuhn–Tucker condition.  $\nabla_B[(Y - XB)^2 + \lambda |B|_1] = 0$  which is

$$
X[X^T B - f(X)] = \nabla_B[\lambda |B|_1].
$$
\n(4)

Since  $X^T B = X^T_{\text{swap}(i,j)} B_{\text{swap}(i,j)}$  and  $f(X_{\text{swap}(i,j)}) = f(X)$ , we have the following,

$$
X\left[X_{\text{swap}(i,j)}^T B_{\text{swap}(i,j)} - f(X_{\text{swap}(i,j)})\right] = \nabla_B[\lambda|B|_1].
$$

We can also swap the  $i^{\text{th}}$  and  $j^{\text{th}}$  column of each side of the equation.  $X_{\text{swap}(i,j)} [X_{\text{swap}(i,j)}^T B_{\text{swap}(i,j)} - f(X_{\text{swap}(i,j)})] = \nabla_{B_{\text{swap}(i,j)}} [\lambda | B_{\text{swap}(i,j)}|_1],$  which is

$$
X\left[X^T B_{\text{swap}(i,j)} - X_{\text{swap}(i,j)}f(X_{\text{swap}(i,j)})\right] = \nabla_{B_{\text{swap}(i,j)}}[\lambda|B_{\text{swap}(i,j)}|_1].
$$

Then we compute the expectation of the left side over all  $X$  and get the following equation.

$$
E[XX^T]B_{swap(i,j)} - E[X_{swap(i,j)}f(X_{swap(i,j)})] = E[XX^T]B_{swap(i,j)} - E[Xf(X)] = \nabla_{B_{swap(i,j)}}[ \lambda | B_{swap(i,j,)}|_1].
$$
 (5)

Therefore,  $B_{swap(i,j)}$  and B are solutions of the equation

$$
E[XXT]B - E[ Xf(X)] = \nabla_B[\lambda|B|_1].
$$
\n(6)

Then from a linear convex combination of Equations (5) and (6), we can know  $(1-a)B + aB_{swap(i,j)},$  $0 < a < 1$ , is also an optimal solution of the problem in Equation (3).

Because  $B(i) = B_{swap(i,j)}(j)$  and  $B(j) = B_{swap(i,j)}(i)$ , for all the model  $(1-a)B + a B_{swap(i,j)}$ , the i<sup>th</sup> component is going from  $B(i)$  to  $B(j)$ , while the j<sup>th</sup> component is going from  $B(j)$  to  $B(i)$ . Thus on the condition that there is a solution with  $B(i) \neq B(j)$ , the correlation coefficient of the i<sup>th</sup> and the j<sup>th</sup> component among all those models is  $-1$ . Q.E.D.