

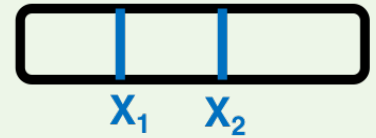
❖ Simulation of phenotypic values (**100** iterations)

$$\mathbf{y} = \mathbf{X}_1\beta_1 + \mathbf{X}_2\beta_2 + \mathbf{X}_3\beta_3 + \mathbf{Z}\mathbf{u} + \mathbf{e}, (\text{Eq. 1})$$

$\mathbf{y}$: phenotypic values $\mathbf{X}_1\beta_1, \mathbf{X}_2\beta_2, \mathbf{X}_3\beta_3$: fixed effects (marker effects)
 $\mathbf{Z}\mathbf{u}$: random effects (polygene effects) $\mathbf{e}$: residuals

- $\mathbf{X}_1$ and $\mathbf{X}_2$ belong to the same haplotype block.

one haplotype block



❖ We assumed 2 scenarios.

- ① **Coupling** : $\text{cor}(\mathbf{X}_1, \mathbf{X}_2) > 0 \rightarrow \beta_1 = \beta_2$, $\text{cor}(\mathbf{X}_1, \mathbf{X}_2) < 0 \rightarrow \beta_1 = -\beta_2$
- ② **Repulsion** : $\text{cor}(\mathbf{X}_1, \mathbf{X}_2) > 0 \rightarrow \beta_1 = -\beta_2$, $\text{cor}(\mathbf{X}_1, \mathbf{X}_2) < 0 \rightarrow \beta_1 = \beta_2$



❖ Perform GWAS with 4 methods.

- ① Our proposed method, **RAINBOW** (R package RAINBOWR)
- ② The **single-SNP** method (R package rrBLUP)
- ③ **Haplotype-based** method introduced by Yano *et al.*, 2016
- ④ **SNP-set** method, SKAT (R package SKAT)



❖ Evaluate the results with the following summary statistics

- ① $-\log_{10}(p)$ and $-\log_{10}(p_a)$, correspond to the **detection power**
- ② **Recall, Precision, and F-measure**
- ③ **AUC** (area under the curve) for **regions around causals**