

Figure S1. Plot of smallest β -weight for simulation study 1. (a) In absence of outliers. (b) In presence of one outlier in each of 10% genes. Where the smallest β -weight represents the minimum value of 20 β -weights for 20 samples for each gene. The outlier genes are indicated in red color. The gray line indicate the maximum value of cutoff, λ =0.13 for outlying genes.



Figure S2. Performance evaluation using ROC curve produced by di erent methods with large-sample case $(n_1 = n_2 = 25)$. (a) In absence of outliers. (b) In presence of one outlier in each of 10% genes (c) In presence of one outlier in each of 20% genes. (d) In presence of one outlier in each of 50% genes.



Figure S3. Performance evaluation using boxplot of MER values estimated by five methods for small-sample case $(n_1=n_2=n_3=n_4=3)$. Boxplot of MER values in absence and presence of one outlier in each of 10%, 20% and 50% genes for small-sample case $(n_1=n_2=n_3=n_4=3)$. The MER values were calculated by five methods (ANOVA, KW, SAM, LIMMA and Proposed) based on top 200 genes.



Figure S4. Four different patterns of DE genes for small-sample case $(n_1=n_2=n_3=n_4=3)$.



Figure S5. Comparison of the top 1944 selected genes by five methods with 1944 valid DE gene set for Platinum Spike dataset for small-sample case $(n_1=n_2=3)$. In absence of outliers, Venn diagram of top 1944 genes detected by (a) the SAM, LIMMA and Proposed method or by (b) the ANOVA, KW and Proposed method with 1944 valid DE gene set. In presence of one outlier in 20% of 1944 valid DE genes, Venn diagram of top 1944 genes detected by (c) the SAM, LIMMA and Proposed method or by (d) the ANOVA, KW and Proposed method with 1944 valid DE gene set.



Figure S6. Gene ontology (GO) categories of three (3) genes for BRCA dataset. The directed acyclic graph (DAG) shows the GO categories of three (3) genes, detected by the Proposed method only for breast cancer (BRCA) dataset. In the DAG tree; each box in the tree lists the name of the GO category, the number of genes in the category, and adjusted *p*-value. The box with red categories indicates that they are enriched with adj. *p*-value<0.05.

Methods	In Absence of Outliers				
	TPR	TNR	FPR	FNR	AUC
ANOVA	0.818	0.978	0.021	0.181	0.816
KW	0.809	0.977	0.022	0.190	0.805
SAM	0.833	0.980	0.019	0.166	0.832
LIMMA	0.829	0.980	0.019	0.170	0.828
EB	0.751	0.971	0.028	0.249	0.749
GAGA	0.770	0.973	0.026	0.229	0.768
BRIDGE	0.716	0.967	0.032	0.284	0.713
Proposed	0.833	0.980	0.019	0.167	0.833
Methods	In Pre	sence of Single	Outlier in 20%	of valid 1944 D	E genes
Methods	In Pre TPR	sence of Single	Outlier in 20%	of valid 1944 D	E genes
Methods ANOVA	In Pre TPR 0.677	TNR 0.962	Outlier in 20% FPR 0.037	of valid 1944 D FNR 0.323	E genes AUC 0.674
Methods ANOVA KW	In Pre TPR 0.677 0.673	sence of Single TNR 0.962 0.962	FPR 0.037 0.037	of valid 1944 D FNR 0.323 0.326	E genes AUC 0.674 0.669
Methods ANOVA KW SAM	In Pre TPR 0.677 0.673 0.736	sence of Single TNR 0.962 0.962 0.971	FPR 0.037 0.037 0.031	of valid 1944 D FNR 0.323 0.326 0.261	E genes AUC 0.674 0.669 0.733
Methods ANOVA KW SAM LIMMA	In Pre TPR 0.677 0.673 0.736 0.680	sence of Single TNR 0.962 0.962 0.971 0.963	FPR 0.037 0.037 0.031 0.037	of valid 1944 D FNR 0.323 0.326 0.261 0.320	E genes AUC 0.674 0.669 0.733 0.678
Methods ANOVA KW SAM LIMMA EB	In Pre TPR 0.677 0.673 0.736 0.680 0.641	sence of Single TNR 0.962 0.962 0.971 0.963 0.958	FPR 0.037 0.037 0.031 0.037	of valid 1944 D FNR 0.323 0.326 0.261 0.320 0.358	E genes AUC 0.674 0.669 0.733 0.678 0.638
Methods ANOVA KW SAM LIMMA EB GAGA	In Pre TPR 0.677 0.673 0.736 0.680 0.641 0.665	sence of Single TNR 0.962 0.962 0.971 0.963 0.958 0.961	FPR 0.037 0.037 0.031 0.037 0.031 0.037	of valid 1944 D FNR 0.323 0.326 0.261 0.320 0.358 0.335	E genes AUC 0.674 0.669 0.733 0.678 0.638 0.638 0.662
Methods ANOVA KW SAM LIMMA EB GAGA BRIDGE	In Pre TPR 0.677 0.673 0.736 0.680 0.641 0.665 0.832	sence of Single TNR 0.962 0.962 0.963 0.963 0.958 0.961 0.982	FPR 0.037 0.037 0.031 0.037 0.031 0.037 0.031 0.037 0.031 0.037	of valid 1944 DI FNR 0.323 0.326 0.261 0.320 0.358 0.335 0.158	E genes AUC 0.674 0.669 0.733 0.678 0.638 0.638 0.662 0.832

Table A1. Performance evaluation of different methods based on Spike gene expression dataset.

In this table the summary statistics (TPR, TNR, FPR, FNR and AUC) are estimated by different methods (ANOVA, KW, SAM, LIMMA, EB, GAGA, BRIDGE and Proposed) based on valid 1944 DE genes for both in absence and presence of one outlier in 20% of valid 1944 genes.