Effects considered	Variables correlated	Variables controlled	Spearman's ρ	<i>p</i> -value
positive	TI, CV		-0.712	<1e-300
positive	TI, mean net $ ES $		-0.604	<1e-300
positive	TI, CV	mean net ES	-0.610	7.1e-22
positive	TI, mean net $ ES $	CV	-0.434	1.2e-10
negative	TI, CV		-0.569	<1e-300
negative	TI, mean net ES		-0.485	<1e-300
negative	TI, CV	mean net ES	-0.392	7.9e-09
negative	TI, mean net $ ES $	CV	-0.209	2.9e-03

Table S1. Rank correlation between trait importance (*TI*) and mean net effect size (|ES|) or phenotypic variation (*CV*) of the wild-type when only positive effects or negative effects are considered.

Table S2. Spearman's rank correlation between the importance of a gene expression trait to fitness and the mean effect size of gene deletion $(|ES_G|)$ or environmental perturbation $(|ES_{\rm E}|)$ after expression levels are controlled.

Variables correlated	Variables controlled	Spearman's ρ	<i>p</i> -value
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³	-0.196	2.2e-16
Fitness effect ¹ , $ ES_G $	Expression level ³	-0.185	2.6e-25
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.145	3.5e-16
Fitness effect ¹ , $ ES_G $	Expression level ³ and $ ES_{\rm E} $	-0.129	4.5e-13
Essentiality ² , $ ES_{\rm E} $	Expression level ³	-0.117	5.7e-11
Essentiality ² , $ ES_G $	Expression level ³	-0.127	1.3e-12
Essentiality ² , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.080	7.7e-06
Essentiality ² , $ ES_G $	Expression level ³ and $ ES_{\rm E} $	-0.094	1.6e-07

¹ Fitness defect caused by deleting the gene. ² Essentiality = 0 for nonessential traits and 1 for essential traits. ³ Expression level of the gene in the wild-type.

Variables correlated	Variables controlled	Spearman's ρ	<i>p</i> -value
Fitness effect ¹ , $ ES_{\rm E} $		-0.191	6.6e-27
Fitness effect ¹ , $ ES_G $		-0.180	3.7e-24
Fitness effect ¹ , $ ES_{\rm E} $	$ ES_{\rm G} $	-0.138	9.4e-15
Fitness effect ¹ , $ ES_G $	$ ES_{\rm E} $	-0.123	5.5e-12
Essentiality ² , $ ES_{\rm E} $		-0.117	6.4e-11
Essentiality ² , $ ES_G $		-0.125	2.2e-12
Essentiality ² , $ ES_{\rm E} $	$ ES_{\rm G} $	-0.078	1.3e-05
Essentiality ² , $ ES_G $	$ ES_{\rm E} $	-0.091	4.1e-07
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³	-0.234	5.8e-40
Fitness effect ¹ , $ ES_G $	Expression level ³	-0.185	2.6e-25
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.183	7.8e-25
Fitness effect ¹ , $ ES_G $	Expression level ³ and $ ES_{\rm E} $	-0.111	4.5e-10
Essentiality ² , $ ES_{\rm E} $	Expression level ³	-0.141	2.4e-15
Essentiality ² , $ ES_G $	Expression level ³	-0.127	1.3e-12
Essentiality ² , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.104	6.9e-09
Essentiality ² , $ ES_G $	Expression level ³ and $ ES_{\rm E} $	-0.083	4.0e-06

Table S3. Spearman's rank correlation between the importance of a gene expression trait to fitness and the mean effect size of gene deletion $(|ES_G|)$ or environmental perturbation $(|ES_{\rm E}|)$ when $|ES_{\rm E}|$ are measured in all environmental changes.

¹ Fitness defect caused by deleting the gene. ² Essentiality = 0 for nonessential traits and 1 for essential traits. ³ Expression level of the gene in the wild-type.

Variables correlated	Variables controlled	Spearman's ρ	<i>p</i> -value
Fitness effect ¹ , $ ES_{\rm E} $		-0.156	6.2e-10
Fitness effect ¹ , $ ES_G $		-0.181	5.1e-13
Fitness effect ¹ , $ ES_{\rm E} $	$ ES_{G} $	-0.099	8.9e-05
Fitness effect ¹ , $ ES_G $	$ ES_{\rm E} $	-0.136	6.3e-08
Essentiality ² , $ ES_{\rm E} $		-0.104	4.0e-05
Essentiality ² , $ ES_G $		-0.131	2.1e-07
Essentiality ² , $ ES_{\rm E} $	$ ES_{\mathbf{G}} $	-0.062	1.5e-02
Essentiality ² , $ ES_G $	$ ES_{\rm E} $	-0.101	6.5e-05
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³	-0.196	4.7e-15
Fitness effect ¹ , $ ES_G $	Expression level ³	-0.185	1.7e-13
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.141	2.1e-08
Fitness effect ¹ , $ ES_G $	Expression level ³ and $ ES_{\rm E} $	-0.124	8.2e-07
Essentiality ² , $ ES_{\rm E} $	Expression level ³	-0.126	5.3e-07
Essentiality ² , $ ES_G $	Expression level ³	-0.132	1.7e-07
Essentiality ² , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.085	7.7e-04
Essentiality ² , $ ES_{G} $	Expression level ³ and $ ES_{\rm E} $	-0.093	2.3e-04

Table S4. Spearman's rank correlation between the importance of a gene expression trait to fitness and the mean effect size of gene deletion $(|ES_G|)$ or environmental perturbation $(|ES_{\rm E}|)$ after the removal of highly correlated expressional traits.

¹ Fitness defect caused by deleting the gene.
 ² Essentiality = 0 for nonessential traits and 1 for essential traits.
 ³ Expression level of the gene in the wild-type.

Variables	Variables controlled	TATA (<i>n</i> =563)		Non-TATA (<i>n</i> =2726)	
correlated		Spearman's	o p-value	Spearman's μ	<i>p</i> -value
Fitness effect ¹ , $ ES_{\rm E} $		-0.185	2.4E-04	-0.178	6.7E-21
Fitness effect ¹ , $ ES_G $		-0.267	8.5E-08	-0.155	4.0E-16
Fitness effect ¹ , $ ES_{\rm E} $	$ ES_{\rm G} $	-0.062	2.2E-01	-0.139	3.4E-13
Fitness effect ¹ , $ ES_G $	$ ES_{\rm E} $	-0.206	4.4E-05	-0.107	2.3E-08
Essentiality ² , $ ES_{\rm E} $		-0.168	8.4E-04	-0.105	3.7E-08
Essentiality ² , $ ES_G $		-0.201	6.2E-05	-0.109	1.3E-08
Essentiality ² , $ ES_{\rm E} $	$ ES_{\rm G} $	-0.081	1.1E-01	-0.076	7.4E-05
Essentiality ² , $ ES_G $	$ ES_{\rm E} $	-0.138	6.6E-03	-0.080	2.6E-05
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³	-0.232	3.6E-06	-0.214	1.2E-29
Fitness effect ¹ , $ ES_G $	Expression level ³	-0.294	3.4E-09	-0.147	1.5E-14
Fitness effect ¹ , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.112	2.8E-02	-0.179	5.6E-21
Fitness effect ¹ , $ ES_G $	Expression level ³ and $ ES_E $	-0.215	1.9E-05	-0.085	9.3E-06
Essentiality ² , $ ES_{\rm E} $	Expression level ³	-0.205	4.6E-05	-0.124	7.9E-11
Essentiality ² , $ ES_{\rm G} $	Expression level ³	-0.221	1.1E-05	-0.102	9.0E-08
Essentiality ² , $ ES_{\rm E} $	Expression level ³ and $ ES_G $	-0.118	2.0E-02	-0.097	3.8E-07
Essentiality ² , $ ES_{\rm G} $	Expression level ³ and $ ES_E $	-0.144	4.5E-03	-0.067	4.9E-04

Table S5. Spearman's rank correlation between the importance of a gene expression trait
 to fitness and the mean effect size of gene deletion $(|ES_G|)$ or environmental perturbation $(|ES_{\rm E}|)$ for genes with and without a canonical TATA box in the promoter, respectively.

¹ Fitness defect caused by deleting the gene. ² Essentiality = 0 for nonessential traits and 1 for essential traits. ³ Expression level of the gene in the wild-type.