

Supplementary Figure 1. Estimating genome-wide distribution of label co-occurrences between gene modules. (A) Ellipses represent gene modules, while green edges depict significantly similar gene modules. Number of label co-occurrences between the modules are indicated by edge styles. Overlapping ellipses indicate which modules are sharing genes, i.e. overlapping. (B) Module-pair collection is sorted according to the number of label co-occurrences, with more similar module-pairs being collected first. (C) Here, the heuristic is determining overlapping modules, with modules having more label co-occurrences having higher precedence over modules with less number of label co-occurrences. For example, both modules in module-pair 2, and one module in module pair 6 are overlapping with modules that have higher precedence. If at least one of the modules is not overlapping with modules of higher precedence, the label co-occurrence value is collected. (D) In this example, out of six module pairs, 5 label co-occurrence values are collected. Note that the label co-occurrence value from pair 2 is disregarded, as both modules are overlapping with pair 1.



Supplementary Figure 2. An example of large gene modules involved in chromatin remodeling in rice. (A) Two gene modules from rice with loc_os10g31970 and loc_os02g46450 used as module centers (large nodes). The nodes represent label co-occurrences, while node labels represent genes assigned to the label co-occurrences. Gray edges represent associations of the label co-occurrences to the module centers. The two modules are overlapping to some degree and consequently share genes, shown by red dashed edges. The number of dashed edges is equal to the number of genes shared between the label co-occurrences. (B) Labels found in the label co-occurrences. For simplicity, only pfam labels are shown. The two modules show enrichment in ontologies representing transcription factors, chromatin remodeling/structure factors, signaling and cell division. The ontology analysis for both modules can be viewed at http://aranet.mpimpgolm.mpg.de/responder.py?name=gene!osa!13835 and http://aranet.mpimpgolm.mpg.de/responder.py?name=gene!osa!8427

dsRNA_bi ubiquitin zf-TAZ



Supplementary Figure 10. An example of large gene modules involved in ribosome biosynthesis in tobacco. (A) To make comparisons of gene module content easier, the co-expression networks are simplified by collecting all genes belonging to one label co-occurrence and representing it as one node. In this example, genes B and C belong to same label co-occurrence (green node) and are assigned to the same node in simplified network. (B) Two gene modules from tobacco with C1368 and C1349 used as module centers (large nodes). The nodes represent label co-occurrences, while node labels represent genes assigned to the label co-occurrences. Gray edges represent associations of the label co-occurrences to the module centers. The two modules are weakly overlapping and consequently sharing genes, which is shown by connecting the overlapping label co-occurrences by red dashed edges. (C) Labels found in the label co-occurrences. For simplicity, only pfam labels are shown. The two modules show enrichment in ontologies representing ribosome structural components.



Supplementary Figure 4. Estimating the distribution of representative module degrees.

(A) Nodes represent modules, and edges indicate similar modules. Numbers adjacent to a module indicate the degree (d) of a module. (B) Module collection is determined by module degree, with modules with higher degree having higher precedence. (C) The first module, with highest degree is collected (module D), together with its neighbors (modules B, C,E and F). Modules can only be collected once. In this example, out of six modules, two module degrees were collected (d=2, d=4 for modules A and D).

Α

HW09A20u_at	Contig18367_at	Contig9885_at C	Contig9764_at C	ontig7171_s_at	Contig14304_at rba	ah38o04_s_at C	ontig9632_at F	HW01L24u_at
Contig6333 at	Contig6238 s at	Contig2249 at C	ontig12071 at	rbags1c11 at	HV CEb0004015r2Co	ontig21968 at C	ontig23141 at C	Contig17921 at
Contig15264_at	Contig8958 at	Contig6546 at C	Contig6009 at	Contig5838 at	Contig2425 s at	_		
GST_ GST_	C A	DH_N DH_zinc_N	p450		DPGT	3Beta_HS Epimerase		hitin_bind_1 Jyco_hydro_19
WRK	Y \blacklozenge p	eroxidase	Bet_v_1	FM Fla	N_red vodoxin_1	Oxidored_		eptidase_C1
В								
Glyma09g067	3 Glyma09g362	29 Glyma06g1137	Glyma12g357	6 Glyma13g3	957 Glyma03g077	75 Glyma12g08	395 Glyma12g3	3568 Glyma20g1997
Glvma12q305	3 Glvma06q003	37 Glvma12q0759	Glvma11g117	2 Glvma15q0	390 Glvma07q182	7 Glvma14q00)67 Glvma13q1	1944 Glyma19q3719
Glyma14q1140) Glyma20q356	67 Glvma07q0023	Glvma15g179	7 Glvma03q3	390 Glvma11q101	5 Glvma11q00)78 Glvma07q1	827
RNA	_pol_Rpb1	RRM_1 NTF2		Frigida	FF WW	DE He DL	AD licase_C JF1605	SMC_N SMC_hinge
SNF2	2_N	Pre-SET SET YDG_SRA	Homeobo PHD DDT	DX BIA Coa TPF TPF	ND htomer_E ₹_1 ₹_2	WD40	PX	GYF

Supplementary Figure 5. Examples of frequently multiplied modules in plants. Genes/probesets that were used as module centers are indicated above the boxes. Colored shapes indicate label co-occurrences that were present in the respective modules. For simplicity, only pfam labels are shown. (A) Metabolism related modules in barley. (B) Transcription related modules in soybean.



Wild type

cobl10-4qrt1-2

pir1-1

pPIR1::GUS

Supplementary Figure 6. Mutants from the pollen cell wall module show normal pollen.

(A-D) Whole anthers and mature pollen (inset upper right) stained with Alexander stain and DAPI (insets lower left) indicate that pollen viability is not affected in the mutants. Note that cob/10-4 was crossed into the quartet (qrt)1-2 background, which displays tetrads of pollen grains after meiosis (Francis et al., 2006). (E) Pollination of wild type pistils with *pPIR::GUS* pollen shows pollen and pollen tube specific expression of *PIR1*. Scale bars: 50 µm (including insets).



Supplemental Figure 7. Hierarchical clustering analysis of LC-MS metabolite profile of tobacco tissues. Relative peak area was normalized by average value and shown with logarithmic scale (log2). Fold change is visualized by indicating color, red (high) and blue (low), respectively.



С



Supplementary Figure 8. EB427179-like gene modules in Arabidopsis. A) Gene module network of EB427179 with Arabidopsis modules shown. B) Expression profile of At5g53810. C) Gene module comparison of EB427179 and At5g53810 and At5g37170.



Supplementary Figure 9. Genes can be present in multiple modules and have multiple LSD relationships. Nodes represent genes, while black solid edges represent co-expression relationships. Node colors represent different gene labels. Dashed edges represent the three LSD relationships. In this example, genes 2 and 4 can be in the same module (module C), or in two similar modules (module A and E), depending on the investigated module.



Supplementary Figure 10. Counting and estimating the significance of large-scale duplicated genes (LSD) in modules. (A) Consider two similar modules, X and Y, containing four genes each. Nodes and node colors represent genes and labels, respectively. Red edges represent LSD pairs found across the two modules. Gray edges represent LSD pairs found within a module, while blue edges represent LSD pairs not found in two similar modules. In this example, 2 red edges, 1 gray edge and 2 blue edges (5 edges in total) were found. Note that for simplicity, only the violet label is analyzed in this example. (B) To estimate the significance of the edge distributions, the 5 LSD edges are distributed randomly among the members of the violet label. The criteria are: the number of edges must stay constant (i.e. 5) and the edges can be only distributed among the violet label. The LSD edges are permuted 1000 times, and the number of red, gray and blue edges is counted for each permutation. The analysis is done for each label, if any LSD gene-pairs are found for the label.