

# De l'expression des gènes aux réseaux

Ecole-chercheur de mai à décembre 2017



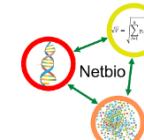
Le projet  
Formation



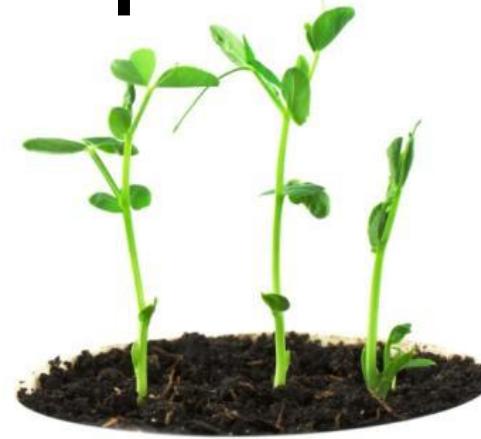
Paris-Saclay  
Computational Biology



Réseaux NETBIO  
du département MIA



# Premières impressions



Toxicogenomics   Morphomics   Epigenomics   Archaeomics  
Proteomics   Phosphoproteomics   Regulomics  
Transcriptomics   Orfeomics  
Metabolomics   Glycomics   Kinomics   Alternomics  
Genomics   Behavioromics   Lipoproteomics   Secretomics  
Lipidomics   Fluxomics  
Interomics



$$y = \sum_{k=0}^{\infty} \frac{x_k - y_k}{\rho(x)} \quad \sigma(u) = \prod_{k=1}^{\infty} (u - \frac{1}{x_k})$$

$$\pi k \leq p\theta - \alpha_0 \leq \pi/2 + 2\pi k, \quad p = 2\sqrt{\nu_0} + (1/2)[\text{sg } A_1 - \text{sg } (A_{n-1} A_n)]$$

$$= \sum_{j=0, j \neq p}^n A_j \rho^j \cos [(p-j)\theta - \alpha_j] + \rho^p.$$

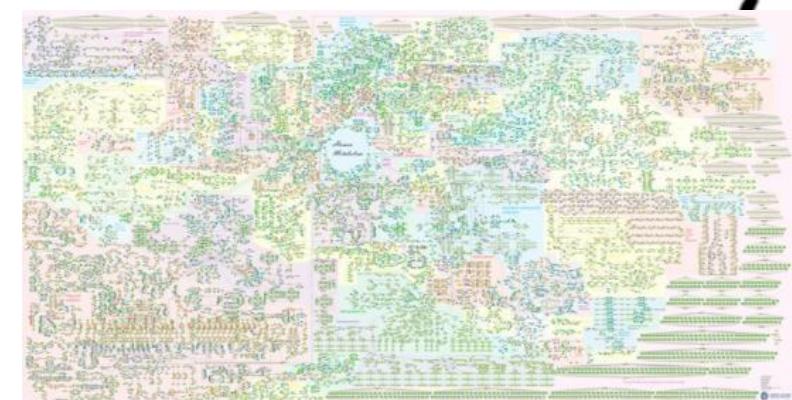
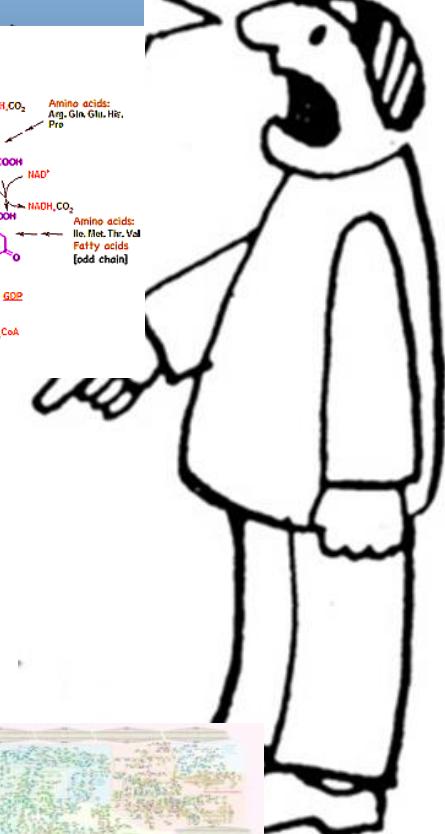
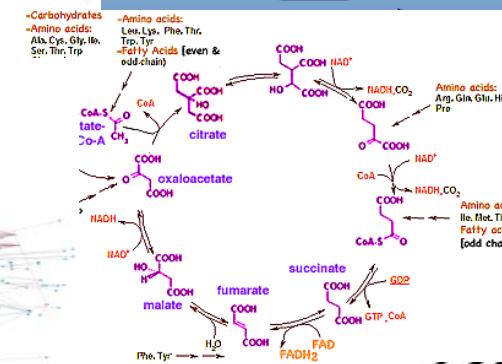
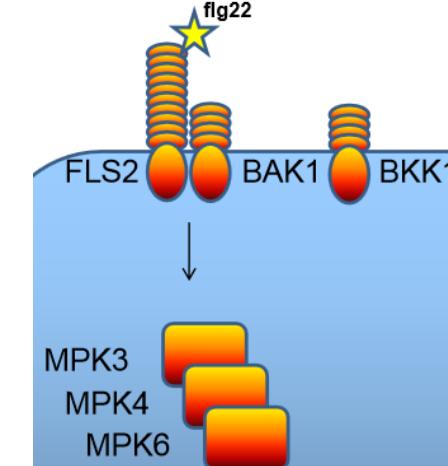
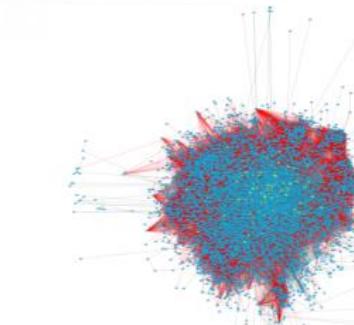
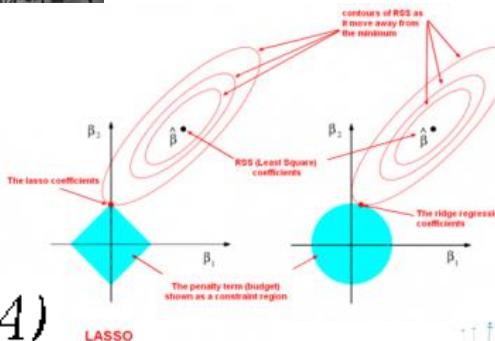
$$\mu \quad \rho^p > \sum_{j=0, j \neq p}^n A_j \rho^j, \quad \Delta_L \arg f(z) = (\pi/2)(S_1 + S_2)$$

$$G(u) = \prod_{k=1}^n (u + u_k) G_0(u), \quad \rho(x) = -G(-x^2)/[xH(-x^2)]$$

$$(A_{n-1} A_n)] \quad p = 2\sqrt{\nu_0} + (1/2)[\text{sg } A_1 - \text{sg } (A_{n-1} A_n)]$$

$$= 2\sqrt{\nu_0} - (1/2)[1 - \text{sg } A_1] \quad \rho^p > \sum_{j=0, j \neq p}^n A_j \rho^j, \quad \mu = -\pi/2 + 2\pi k \leq$$

$$f(z) = \frac{(\pi/2)(S_1 + S_2)}{\mu} \quad G(u) = \prod_{k=1}^n (u + u_k) G_0(u), \quad K^{(r)}(x, y) = K_r(x, y) +$$



# Objectifs de l'école chercheurs

- Comprendre les spécificités des données omiques, en particulier de transcriptomique
  - Comprendre les objectifs des analyses bio-informatiques et statistiques
  - Connaître les méthodes proposées
  - Savoir interpréter a minima les résultats
- 
- Se familiariser avec les concepts et partager un vocabulaire technique commun
  - Etre capable d'interagir avec les plateformes de omiques et les modélisateurs
  - Savoir faire des choix méthodologiques avisés

# 6 modules de formation indépendants

Matin > Cours en amphithéâtre

2 fois 90 minutes avec une pause de 15 minutes

Après-midi > 2 TP de 90 minutes pour un groupe  
de 20 personnes

# 6 modules de formation indépendants

Organisés selon les différentes étapes de l'analyse

Les technologies de séquençage actuelles et futures et leurs protocoles associés (30 mai)

La bioinformatique du RNAseq (29 juin)

Normalisation et analyse différentielle des données RNAseq  
(28 septembre)

Analyse de la coexpression de données RNAseq (12 octobre)

Les réseaux de régulation (30 novembre)

Les intégrations de données omiques (14 décembre)

# Gene co-expression analysis

## MORNING TALKS

- Co-expression analysis: what for? Examples of co-expression studies
- Co-expression analysis: How?

## AFTERNOON PRACTICAL SESSIONS

- coseq R package for RNA-seq co-expression analysis
- Using the Gem2Net database

# Co-expression analysis: what for?

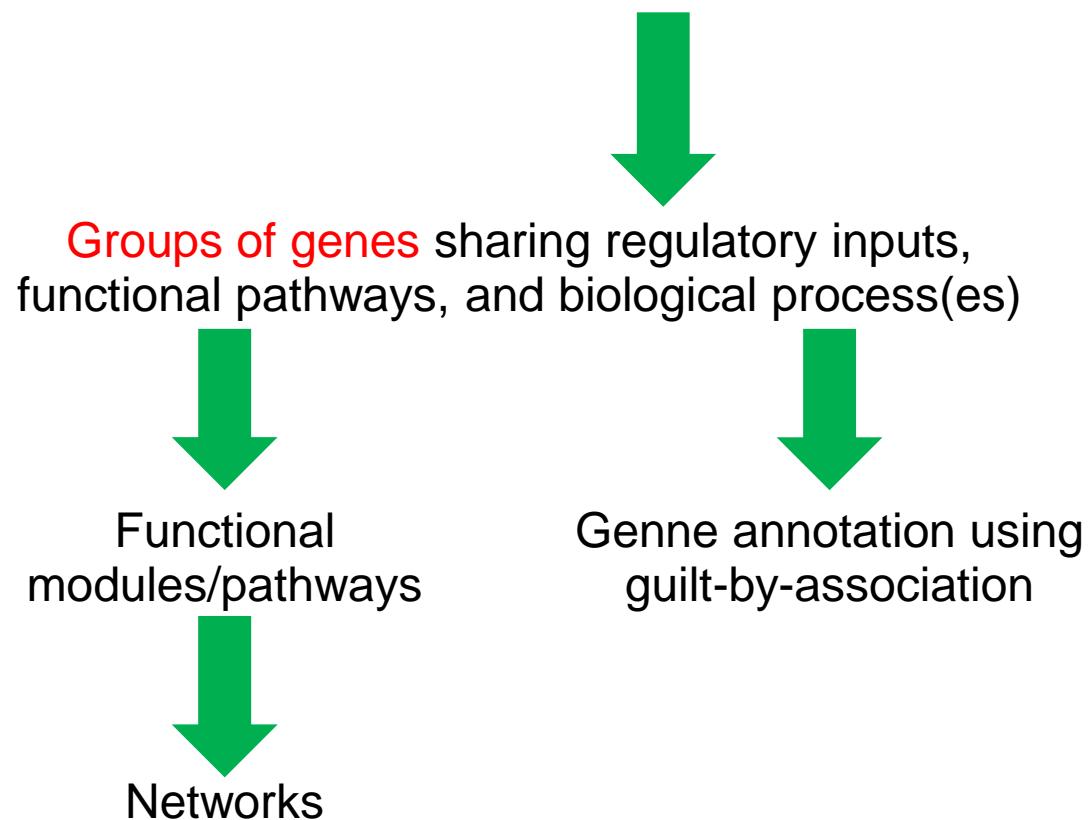
## Examples of co-expression studies

Co-expressed genes =  
Groups of genes that have **similar expression patterns** over a range of different experiments

# Co-expression analysis: what for?

## Examples of co-expression studies

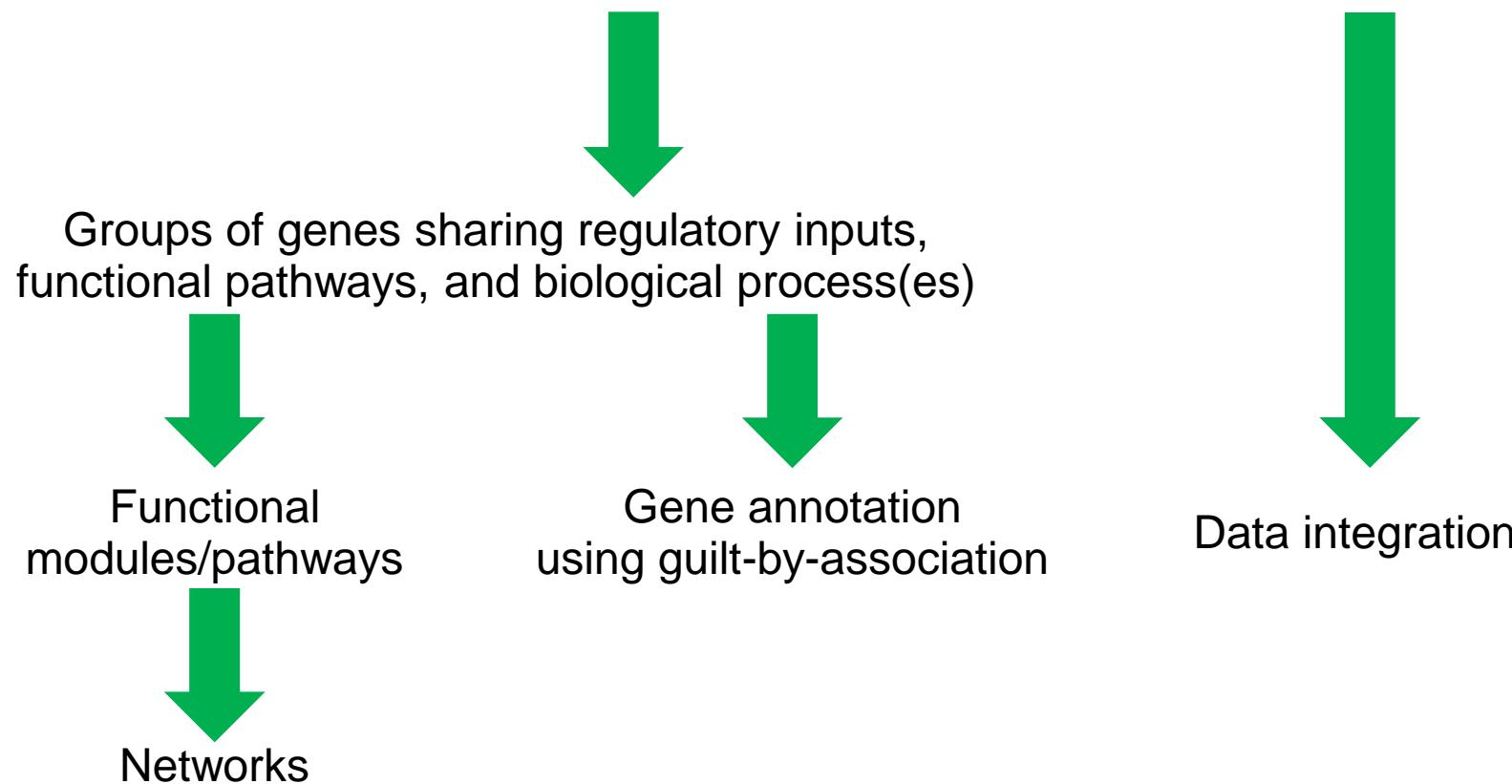
Co-expressed genes =  
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# Co-expression analysis: what for?

## Examples of co-expression studies

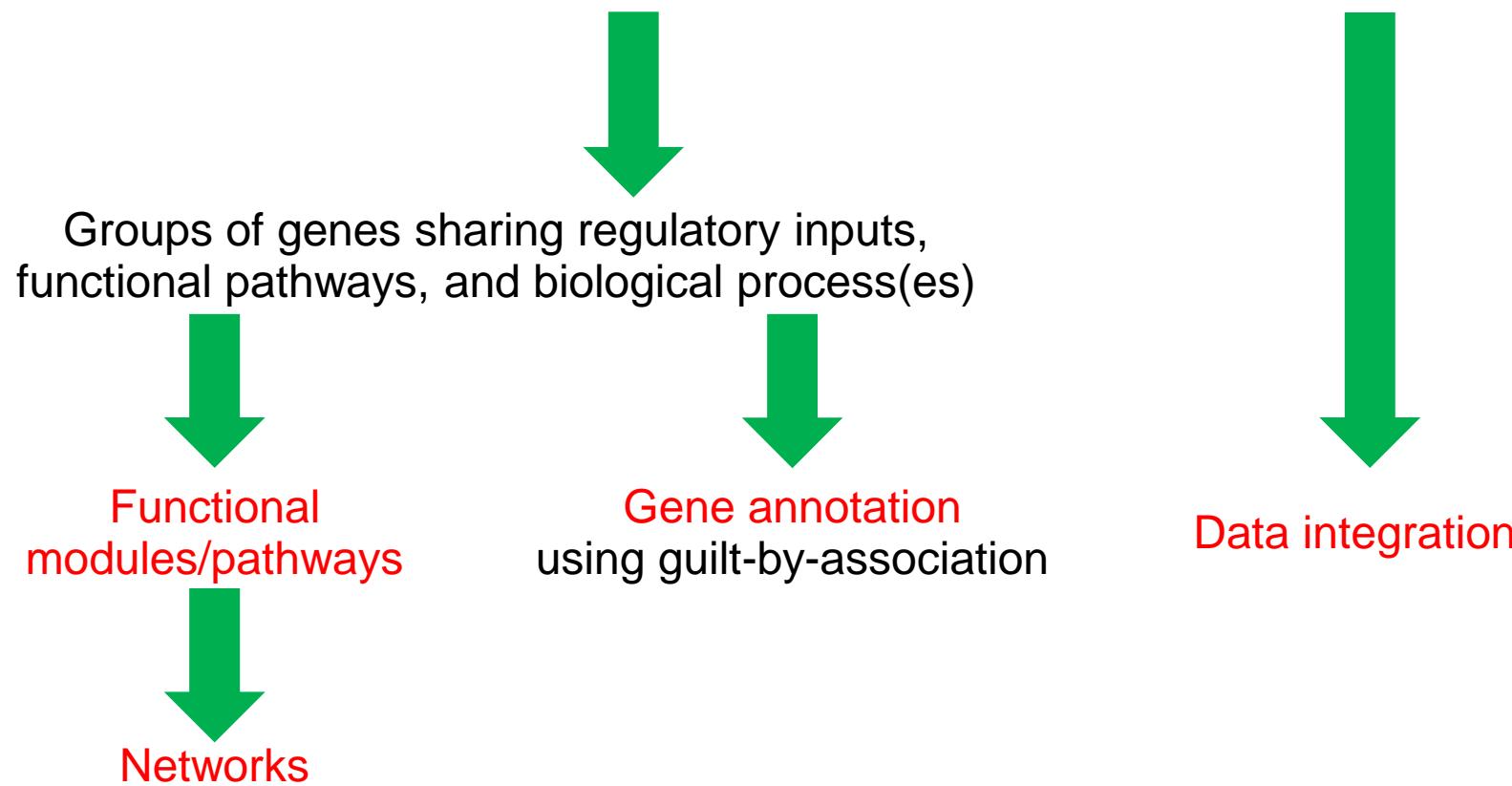
Co-expressed genes =  
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# Co-expression analysis: what for?

## Examples of co-expression studies

**Co-expressed genes =**  
Groups of genes that have similar expression patterns over a range of different experiments



# Co-expression analysis: what for?

## Examples of co-expression studies



RESEARCH ARTICLE



### Temporal network analysis identifies early physiological and transcriptomic indicators of mild drought in *Brassica rapa*

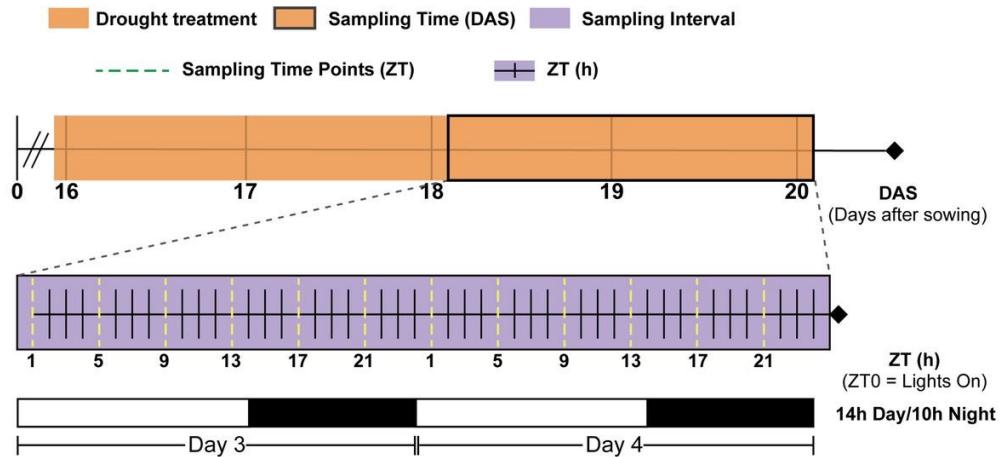
Kathleen Greenham<sup>1†</sup>, Carmela Rosaria Guadagno<sup>2†</sup>, Malia A Gehan<sup>3</sup>,  
Todd C Mockler<sup>3</sup>, Cynthia Weinig<sup>2,4,5</sup>, Brent E Ewers<sup>2,5</sup>, C Robertson McClung<sup>1\*</sup>

October, 4th 2017

# Co-expression analysis: what for?

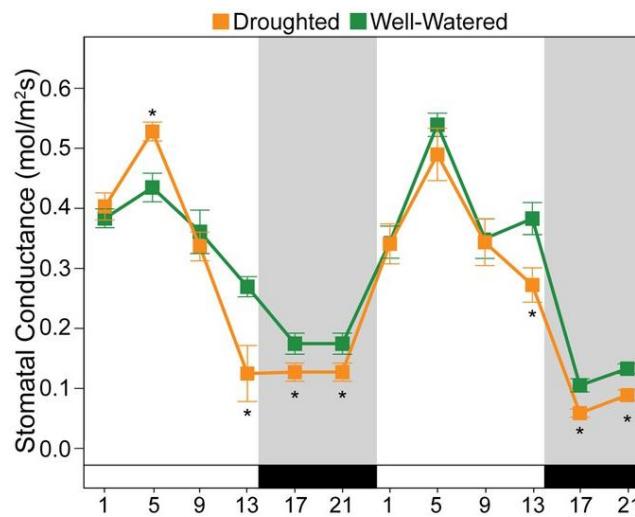
## Examples of co-expression studies

A



Mild drought, sampling every 4h for 48h

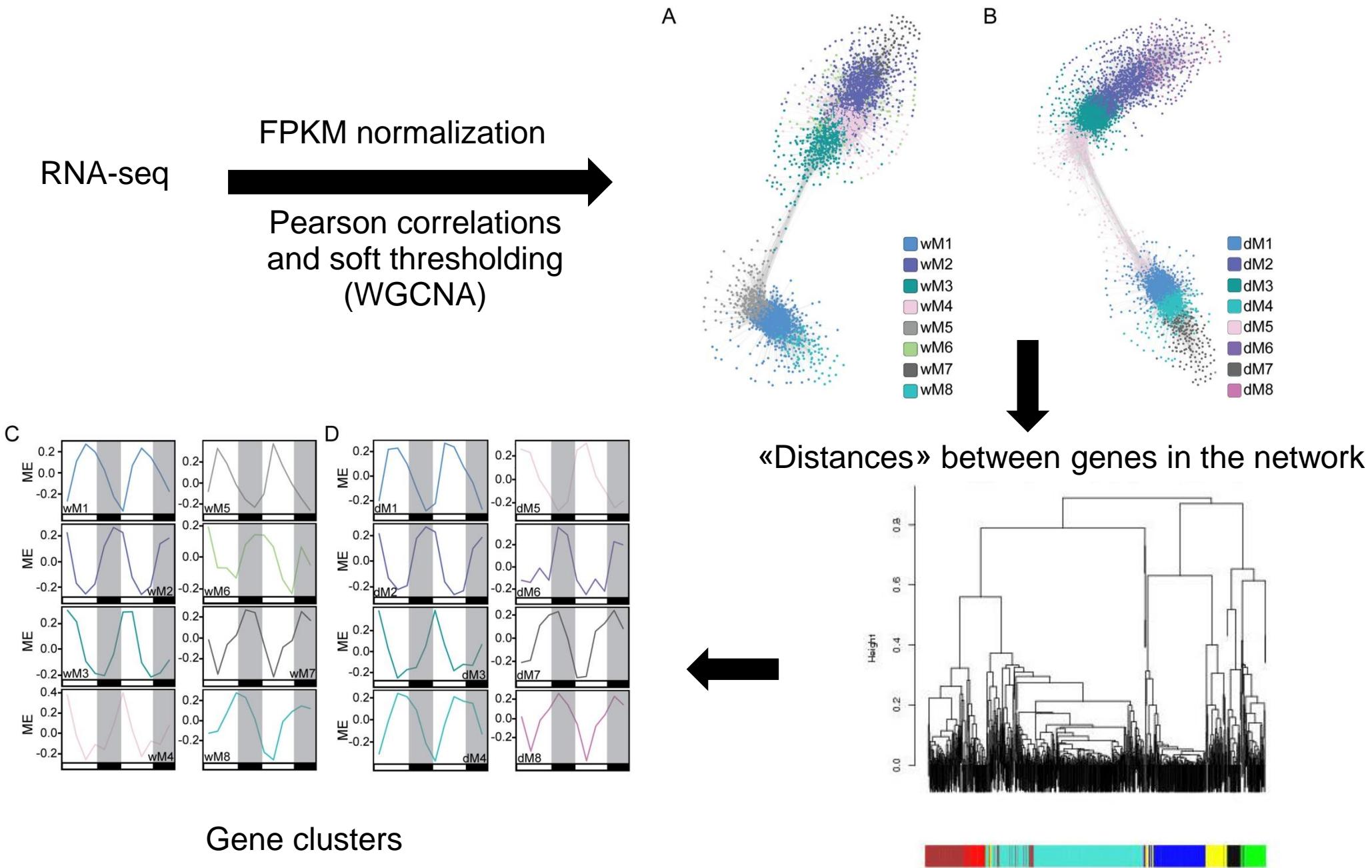
### Physiological measures



$F_v'/F_m'$ , gs and NSC are impacted by drought

# Co-expression analysis: what for?

## Examples of co-expression studies

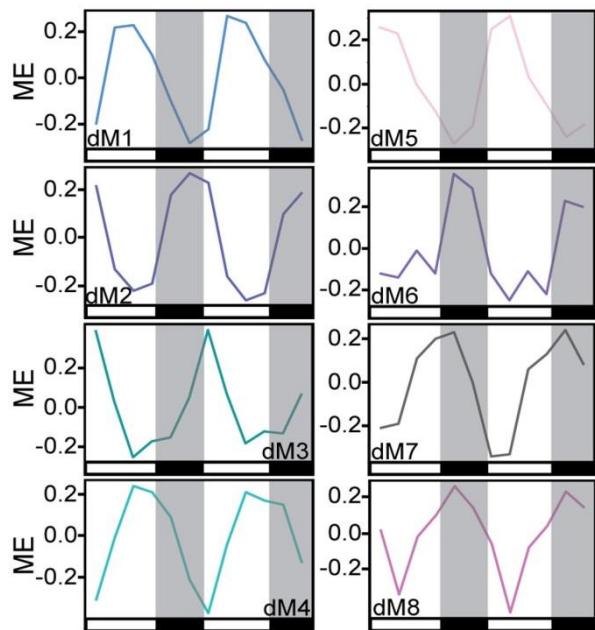


# Co-expression analysis: what for?

## Examples of co-expression studies

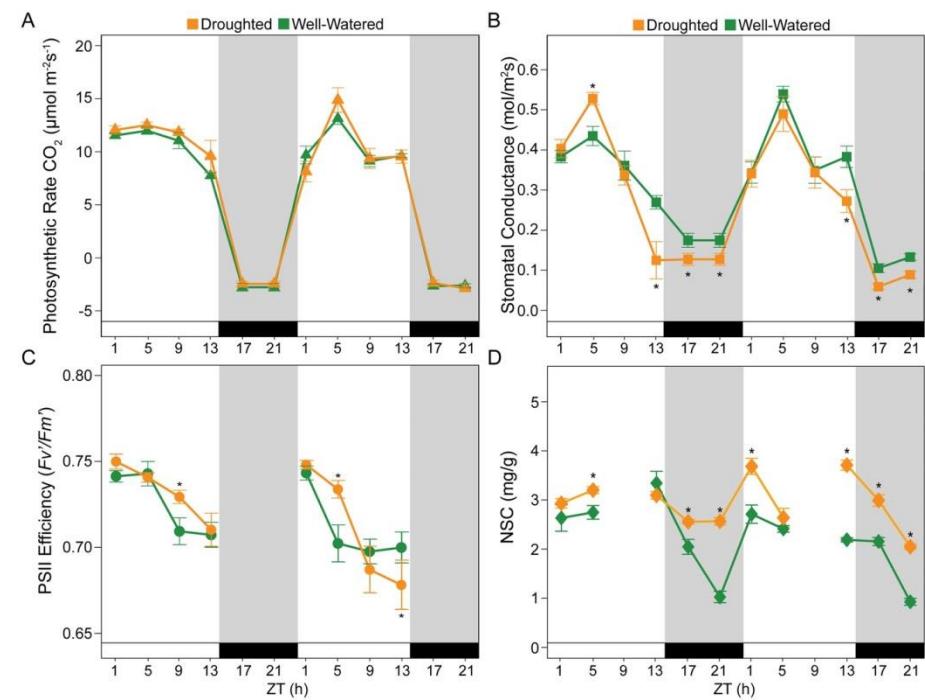
### Selection of the clusters of interest

clusters



Pearson correlation

physiology



+

changes in expression rythm between drought and control

# Co-expression analysis: what for?

## Examples of co-expression studies

Correlated

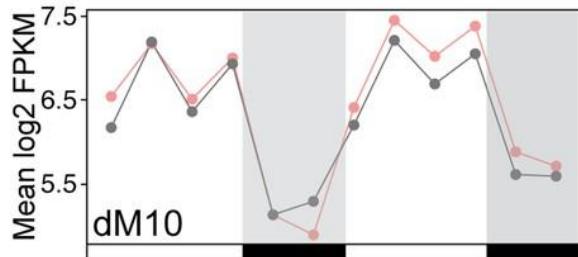
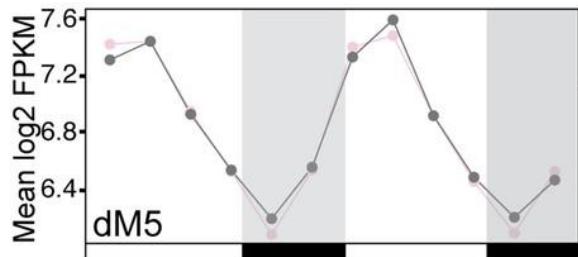
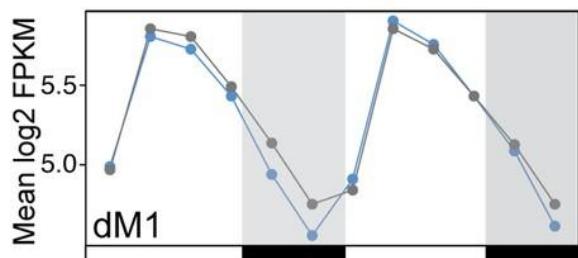
to

gs

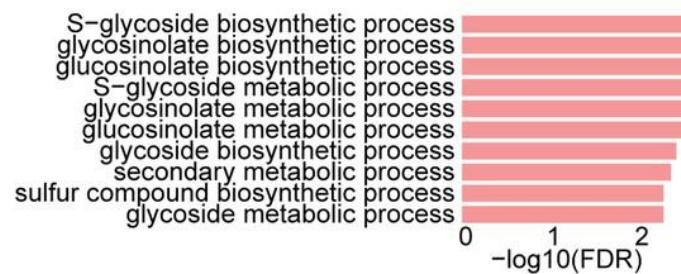
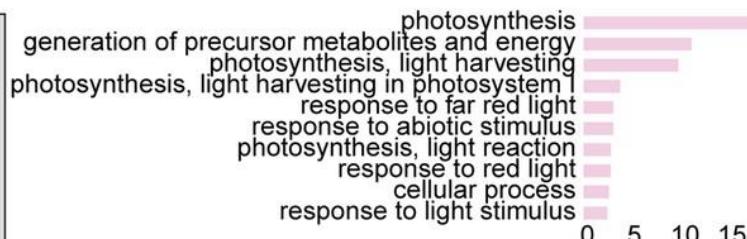
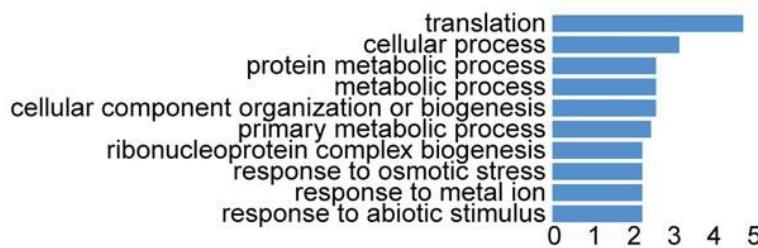
gs

Fv'/Fm'

Cluster expression profile



GO enrichments



Identification of functional modules (pathways) associated  
with the early onset of drought

# Co-expression analysis: what for?

## Examples of co-expression studies

The Plant Cell, Vol. 29: 2086–2105, September 2017, www.plantcell.org © 2017 ASPB.

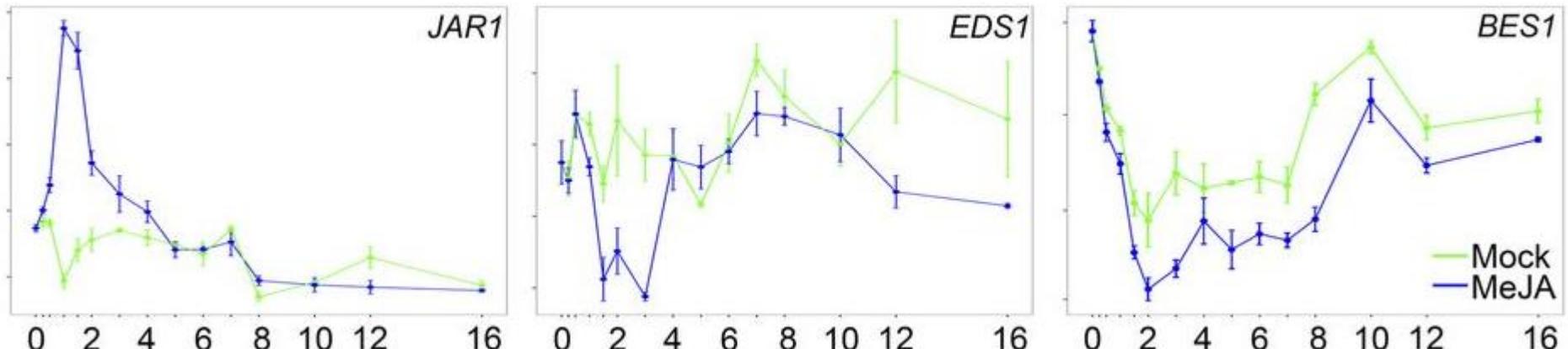


LARGE-SCALE BIOLOGY ARTICLE

## Architecture and Dynamics of the Jasmonic Acid Gene Regulatory Network<sup>OPEN</sup>

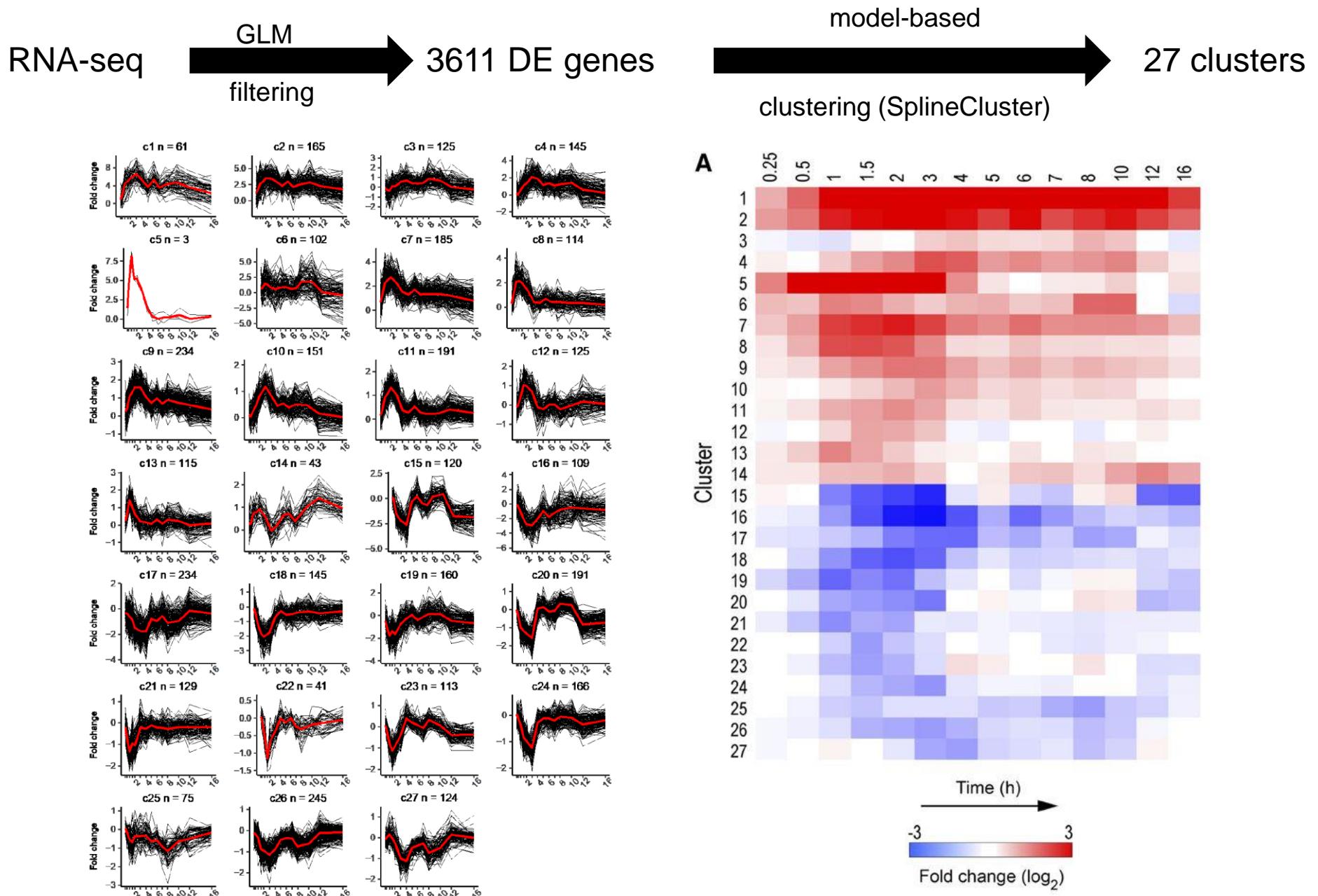
Richard Hickman,<sup>a,1</sup> Marcel C. Van Verk,<sup>a,b,1</sup> Anja J.H. Van Dijken,<sup>a</sup> Marciel Pereira Mendes,<sup>a</sup> Irene A. Vroegop-Vos,<sup>a</sup> Lotte Caarls,<sup>a</sup> Merel Steenbergen,<sup>a</sup> Ivo Van der Nagel,<sup>a</sup> Gert Jan Wesselink,<sup>a</sup> Aleksey Jironkin,<sup>c</sup> Adam Talbot,<sup>d,e</sup> Johanna Rhodes,<sup>c</sup> Michel De Vries,<sup>f</sup> Robert C. Schuurink,<sup>f</sup> Katherine Denby,<sup>c,d,e</sup> Corné M.J. Pieterse,<sup>a</sup> and Saskia C.M. Van Wees<sup>a,2</sup>

MeJA treatment of *Arabidopsis thaliana* leaf 6 (+mock).  
Kinetic over 16h with 15 time points followed by RNA-seq analysis



# Co-expression analysis: what for?

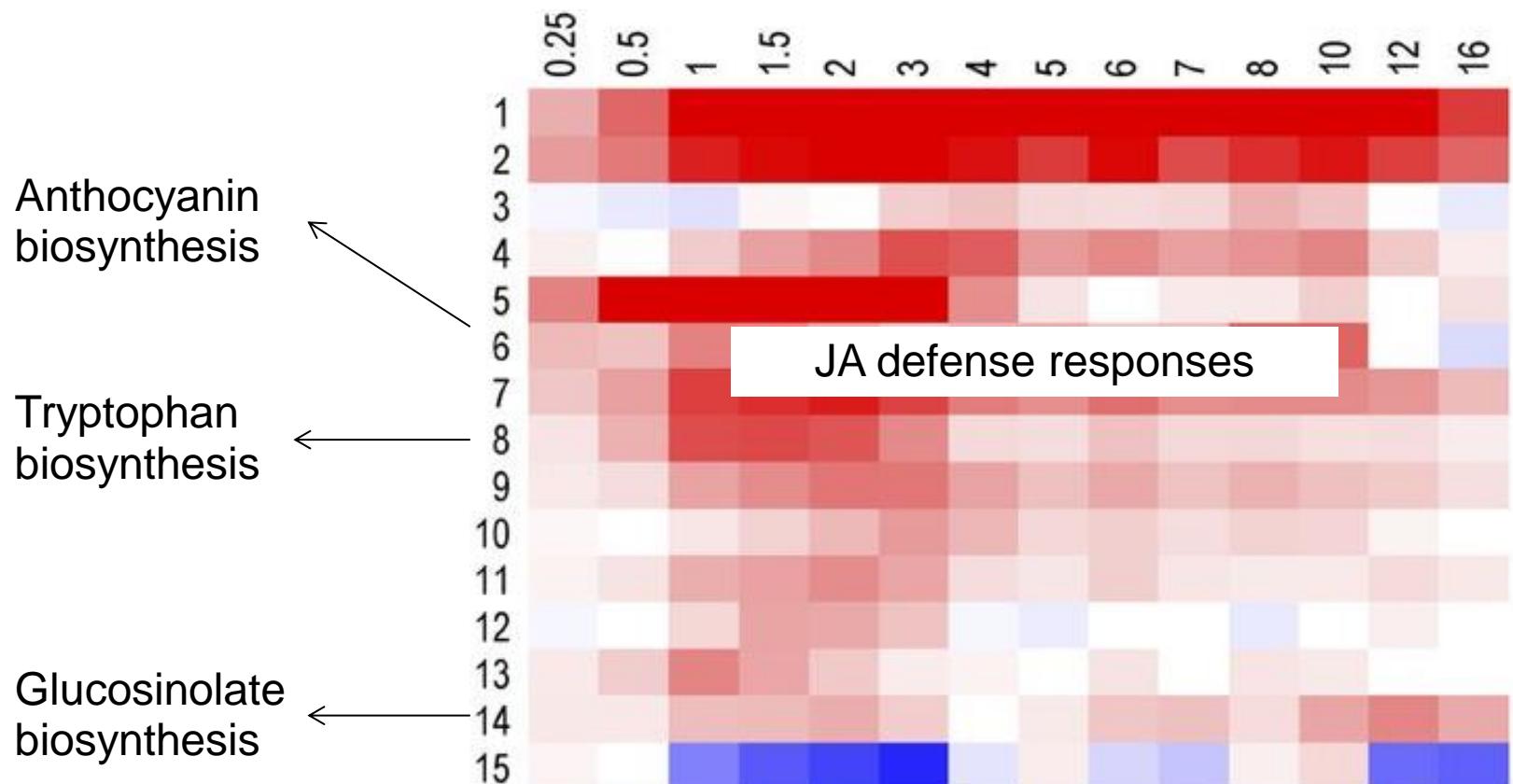
## Examples of co-expression studies



# Co-expression analysis: what for?

## Examples of co-expression studies

Functional modules: GO enrichments



Identification of discrete sectors of the JA-controlled gene network

# Co-expression analysis: what for?

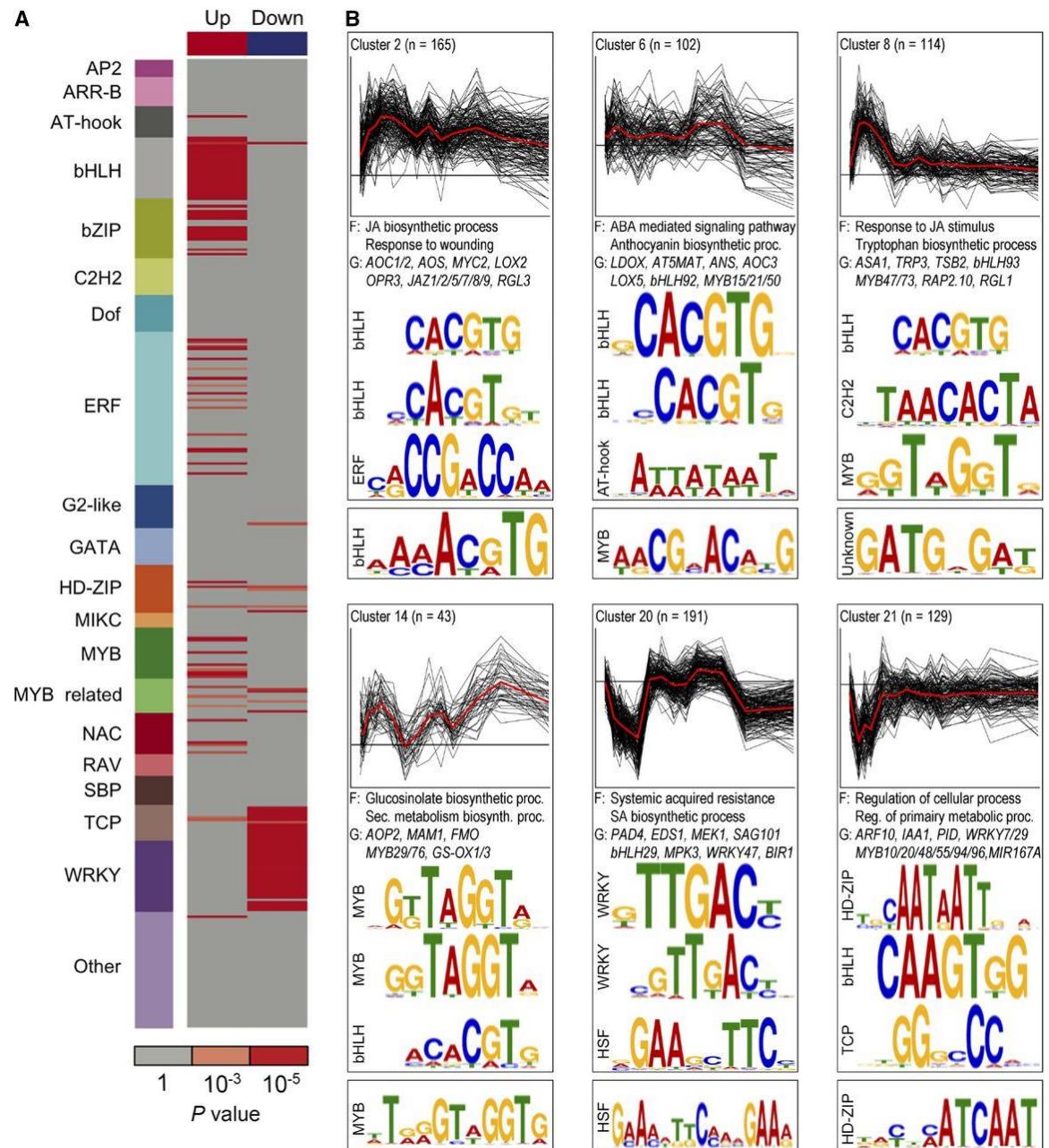
## Examples of co-expression studies

Gene regulation:  
DNA binding motif analysis

bHLH binding motifs enriched in most up-regulated clusters



bHLH TFs are essential components of MeJA induction



# Co-expression analysis: what for?

## Examples of co-expression studies

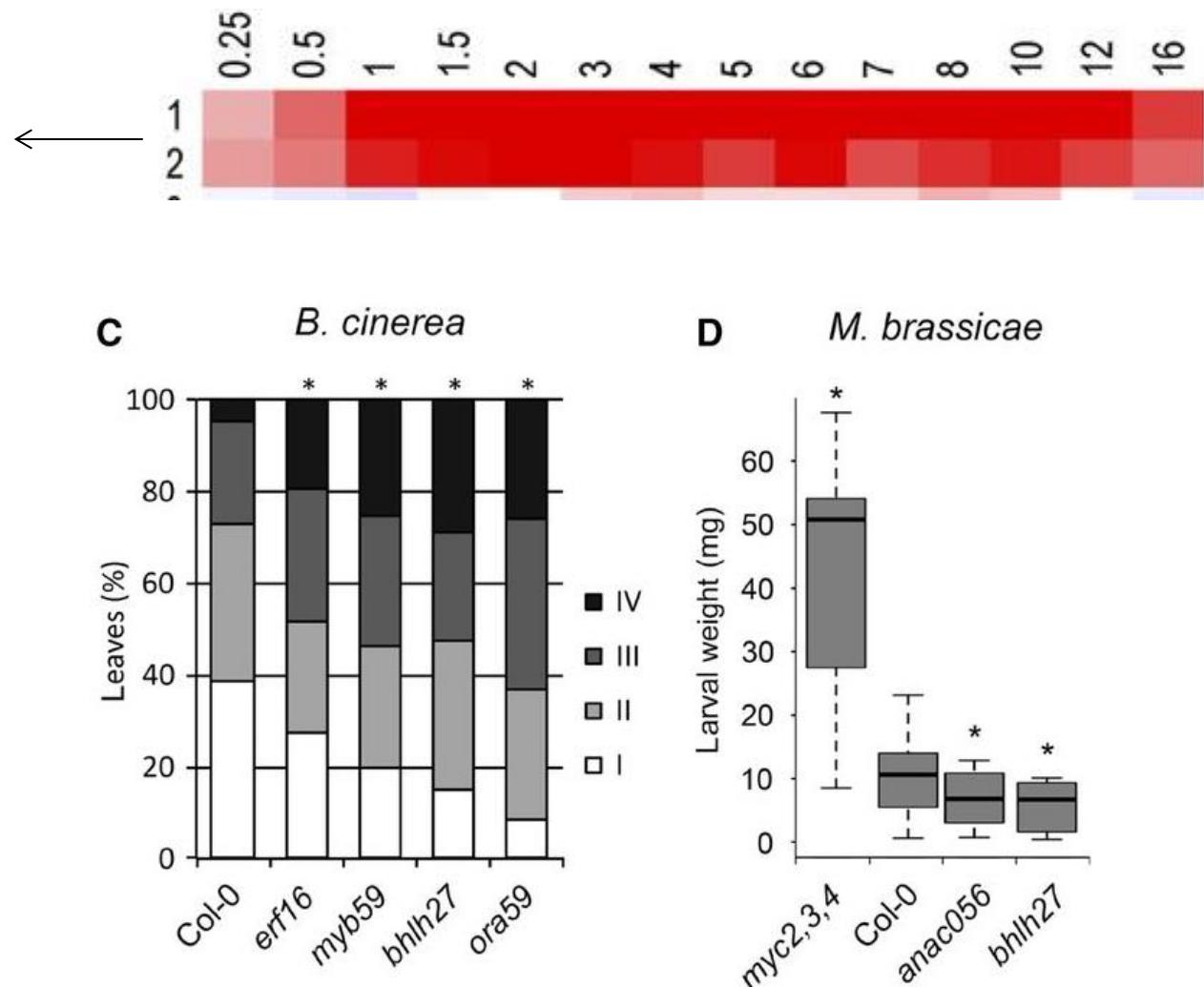
Gene annotation: New regulators of JA defense responses

Enriched in known JA-related genes including regulators



Testing of 12 uncharacterised TF from these clusters

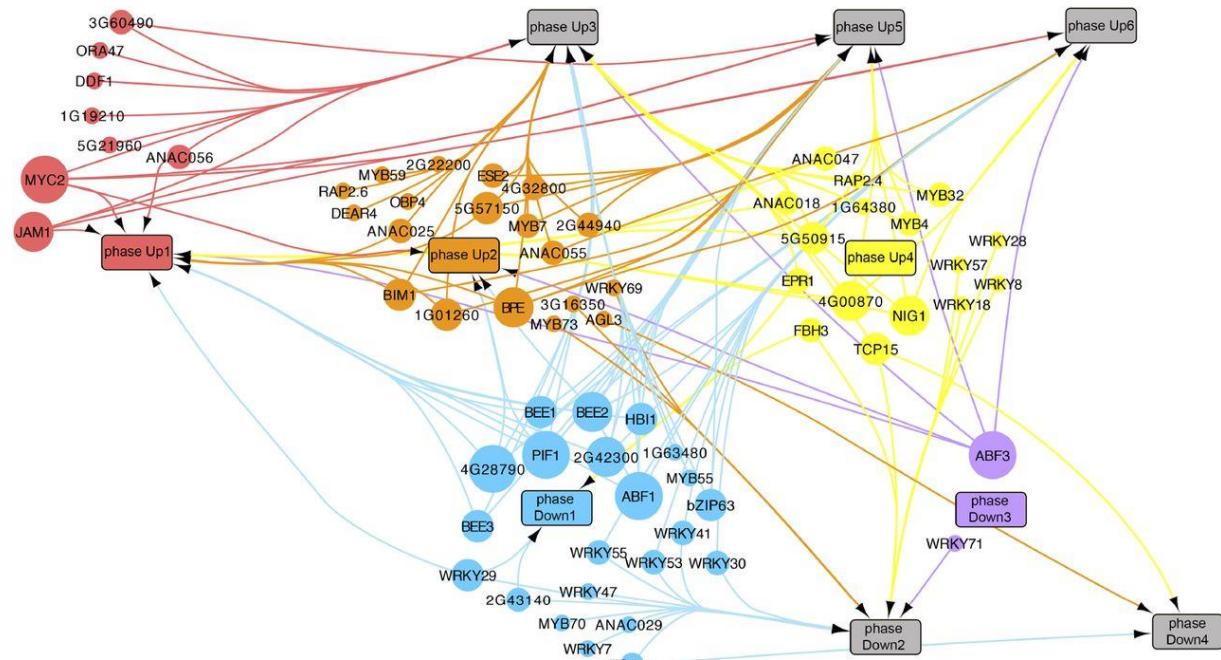
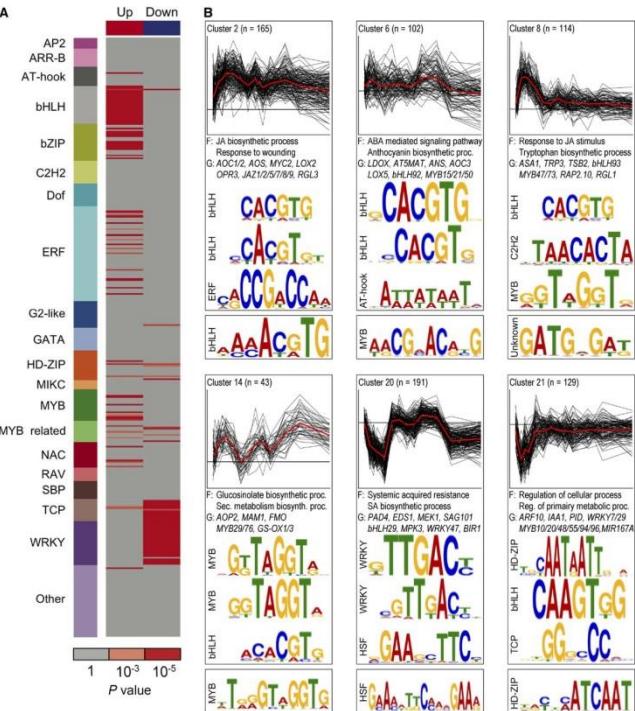
4/12 tested TF are involved in JA defense responses



# Co-expression analysis: what for?

## Examples of co-expression studies

Data integration and gene clustering for network inference



Gene clusters, gene annotation and promoter analysis

JA gene regulatory network

# Co-expression analysis: what for?

## Examples of co-expression studies

**Co-expressed genes =**  
Groups of genes that have similar expression patterns over a range of different experiments

