

# Decoding the polyploid wheat genome using gene networks



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# Outline

- 1) Why is polyploidy relevant?
- 2) Why now?
- 3) Can we use gene networks to escape polyploidy?



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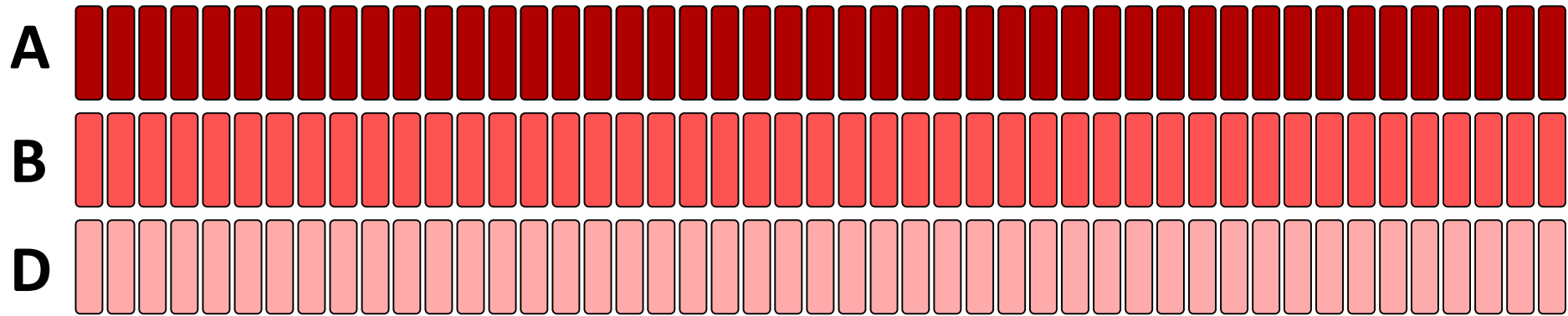


# Polyploid plants are abundant

- >70 % of flowering plants have a polyploid evolutionary history
- Many major crops are polyploid



# The Challenge of Wheat

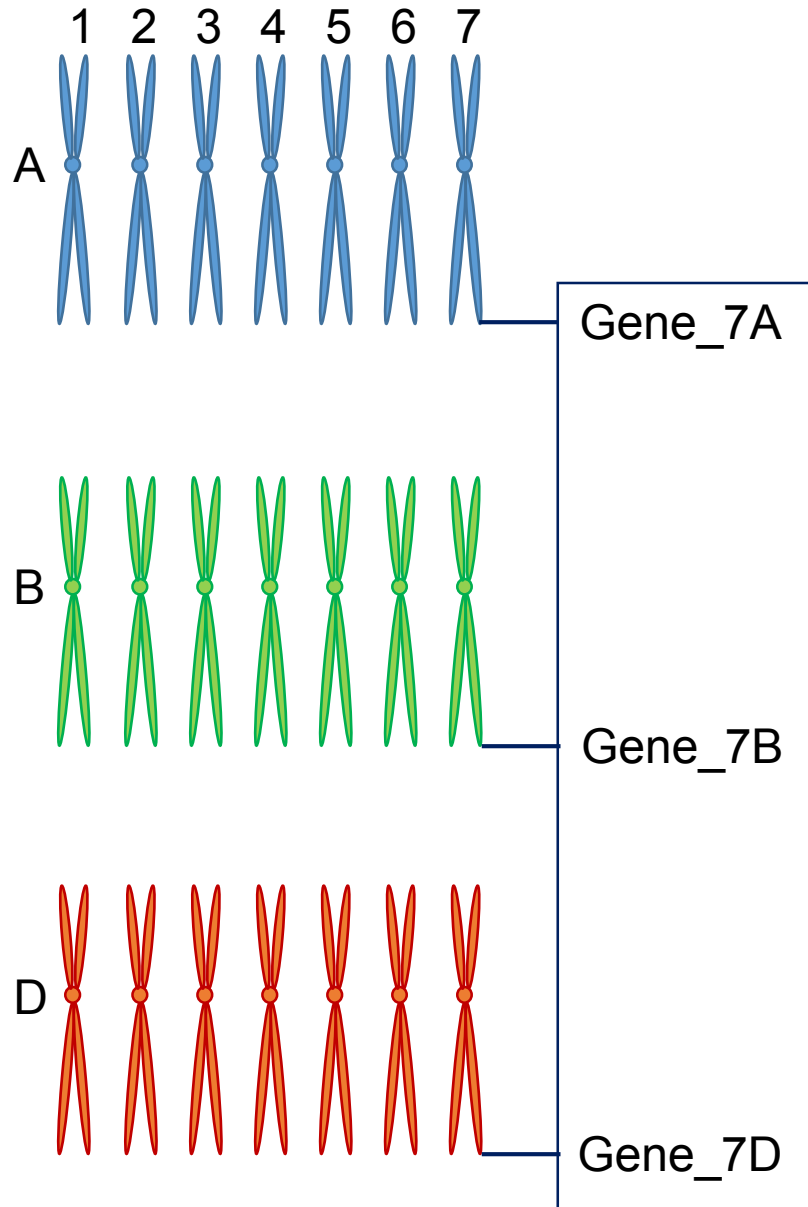


Arabidopsis

Rice

Human

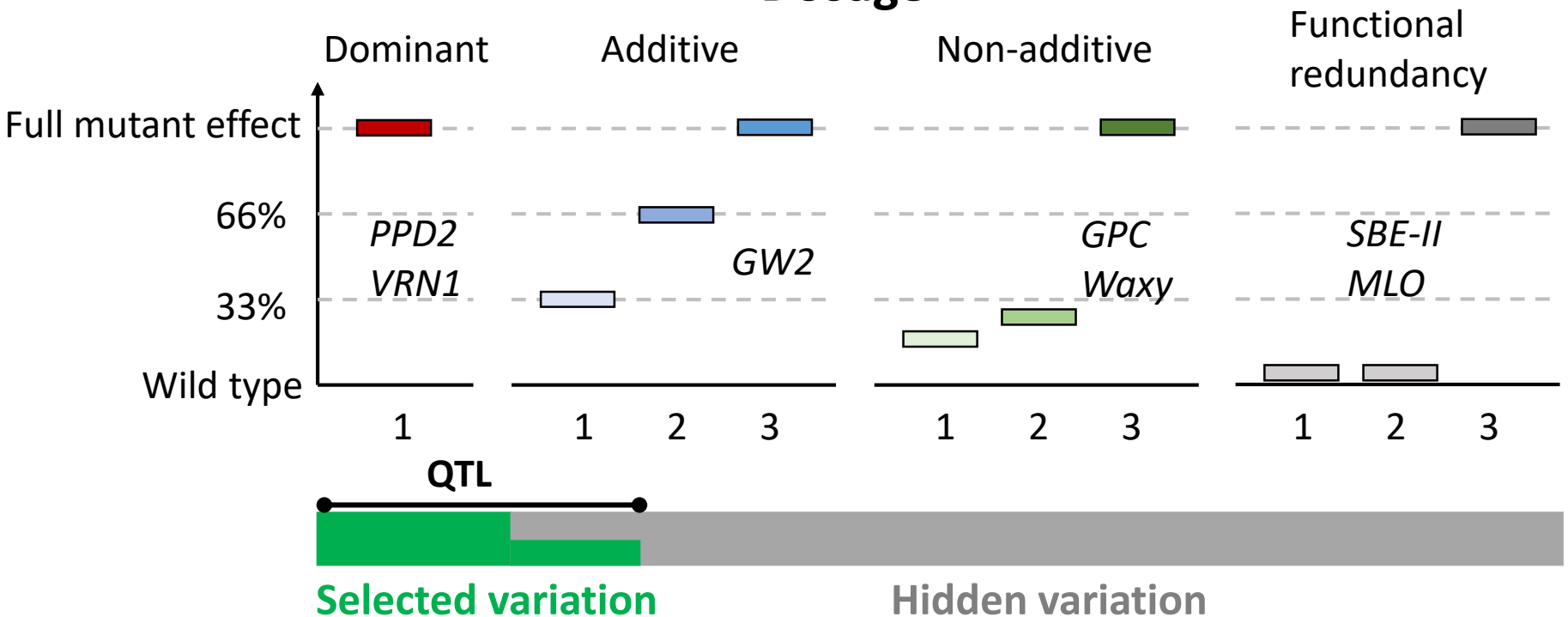
## Wheat chromosomes



**Homoeologues  
>95 % CDS similarity**

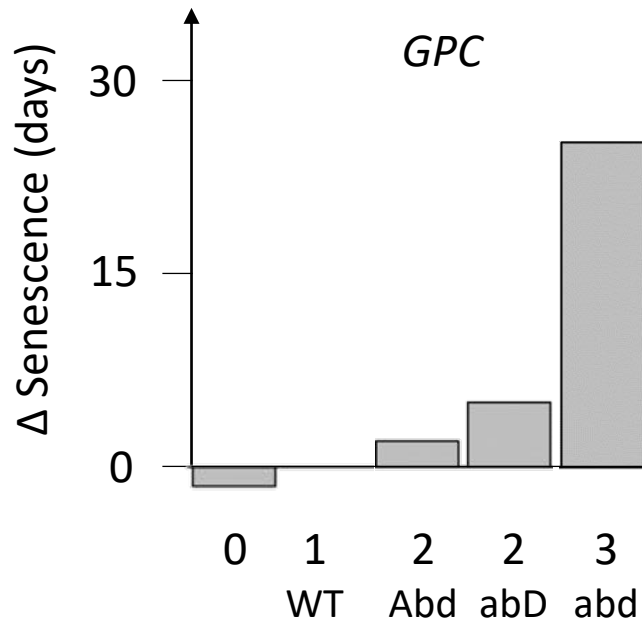
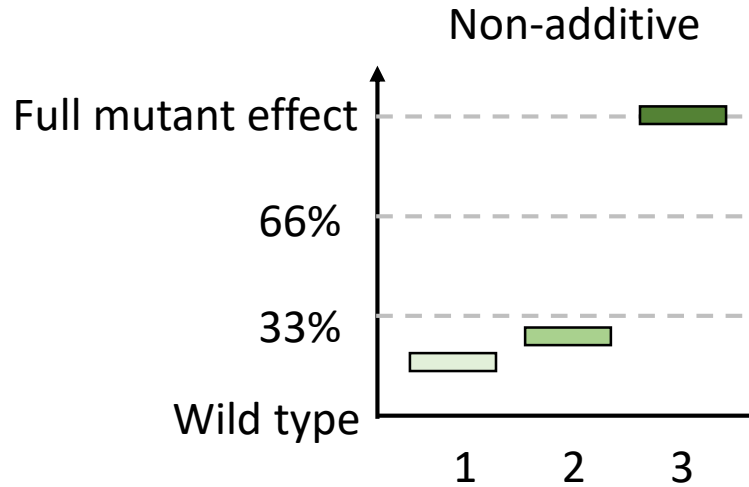
# Functional redundancy hides variation

## Dosage





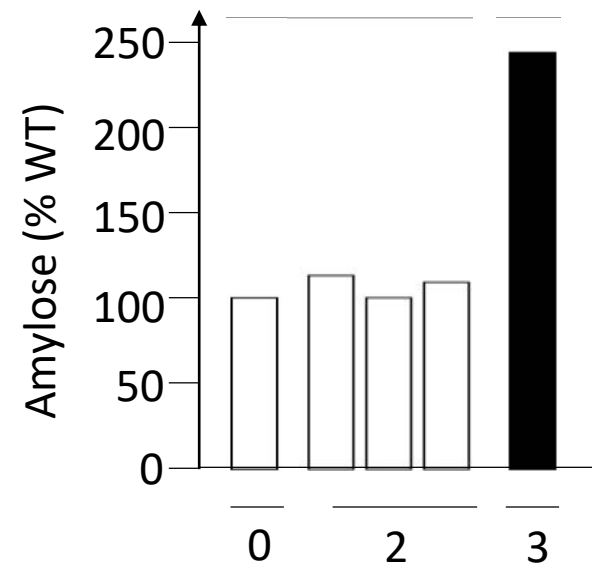
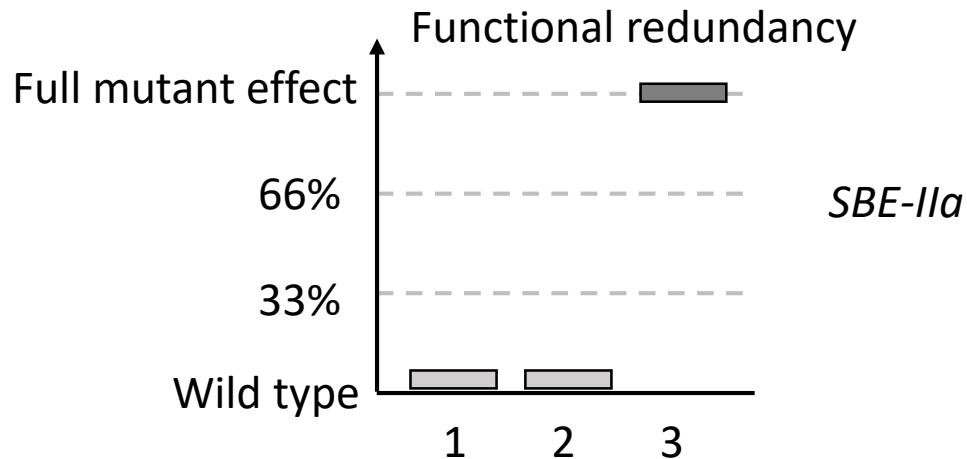
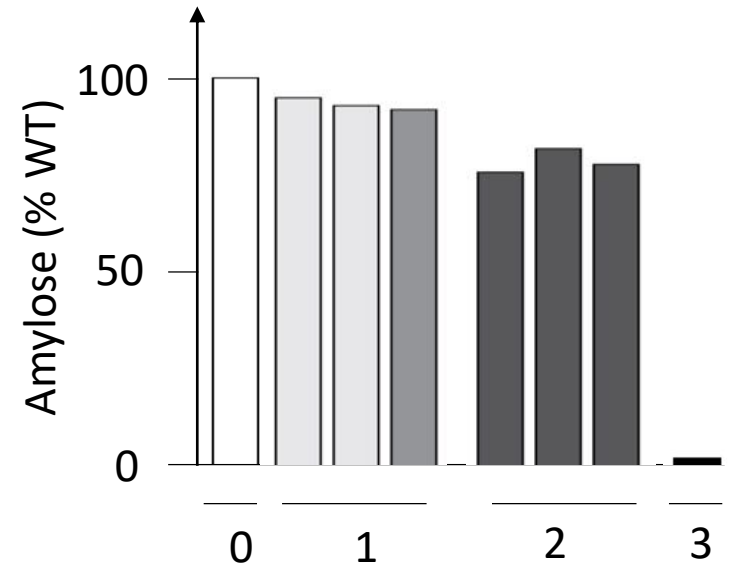
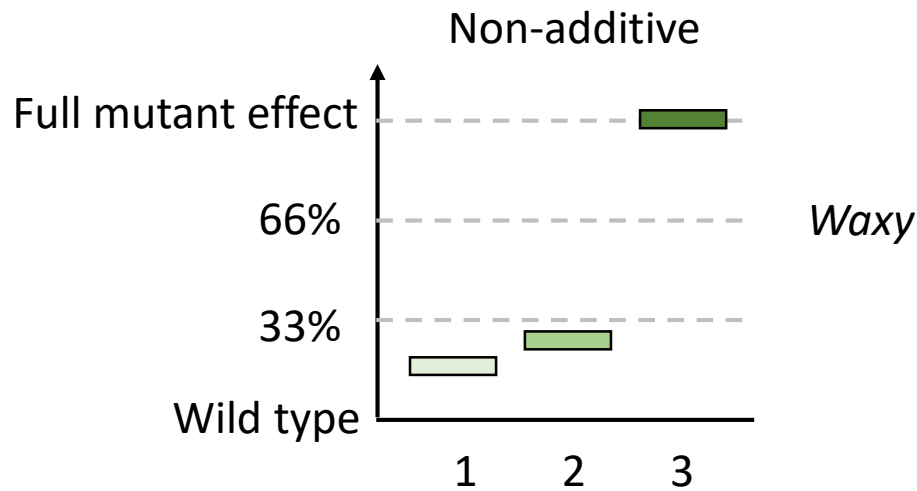
# In young polyploids recessive alleles are missed by breeders



Uauy et al 2006 *Science*; Avni et al 2014 *Planta*;  
Borrill et al 2015 *Plos One*.  
Borrill, Adamski & Uauy 2015 *New Phytologist*

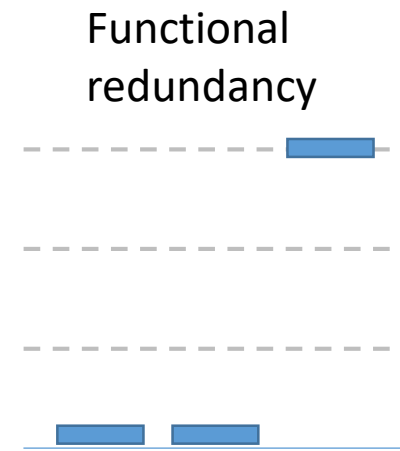


# Incredible potential hiding in the wheat genome



# 1) Why is polyploidy relevant?

- Polyploidy is common in major crop species
- Hides variation





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# Fully sequenced wheat genome



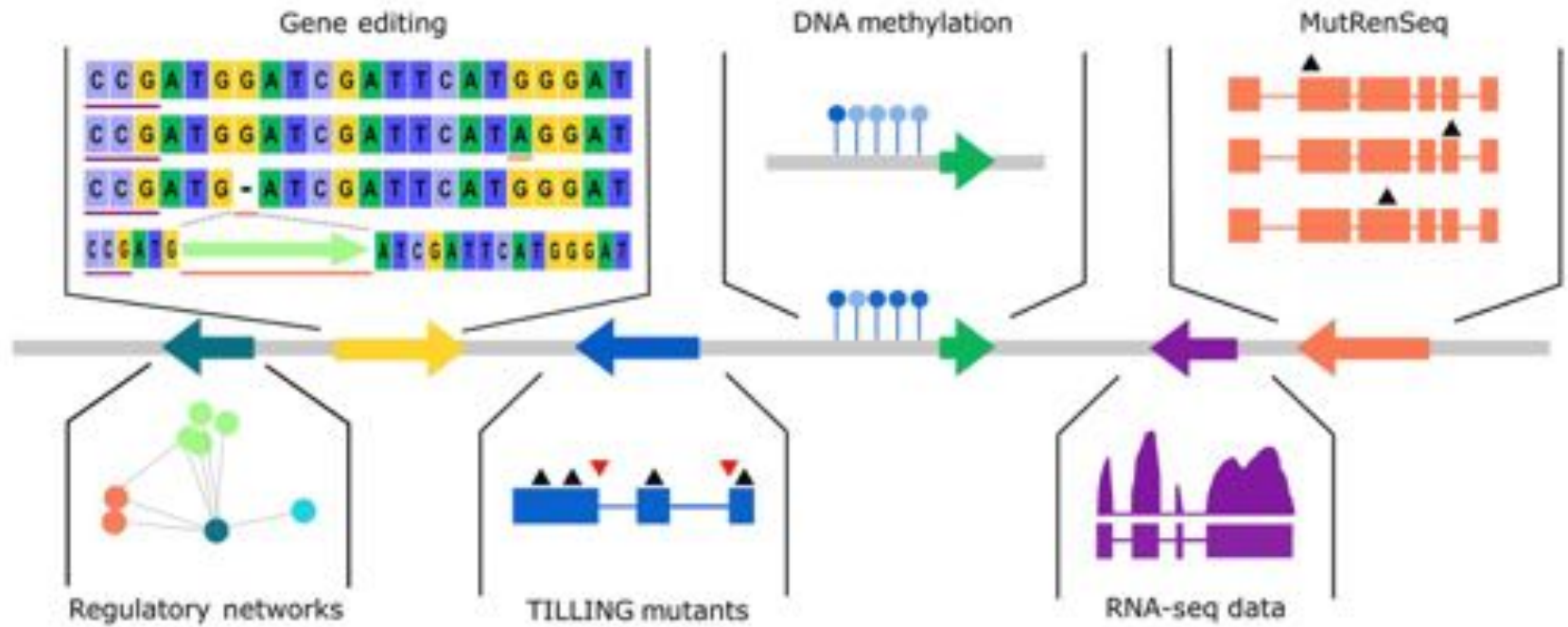
14.5 Gb assembled (genome size ~15.5 Gb -> **94%**)

14.1 Gb in 21 pseudomolecules (97%)  
1601 superscaffolds (avg 76 per chr.)

110,790 genes



# Integrating genomic resources



REVIEW ARTICLE



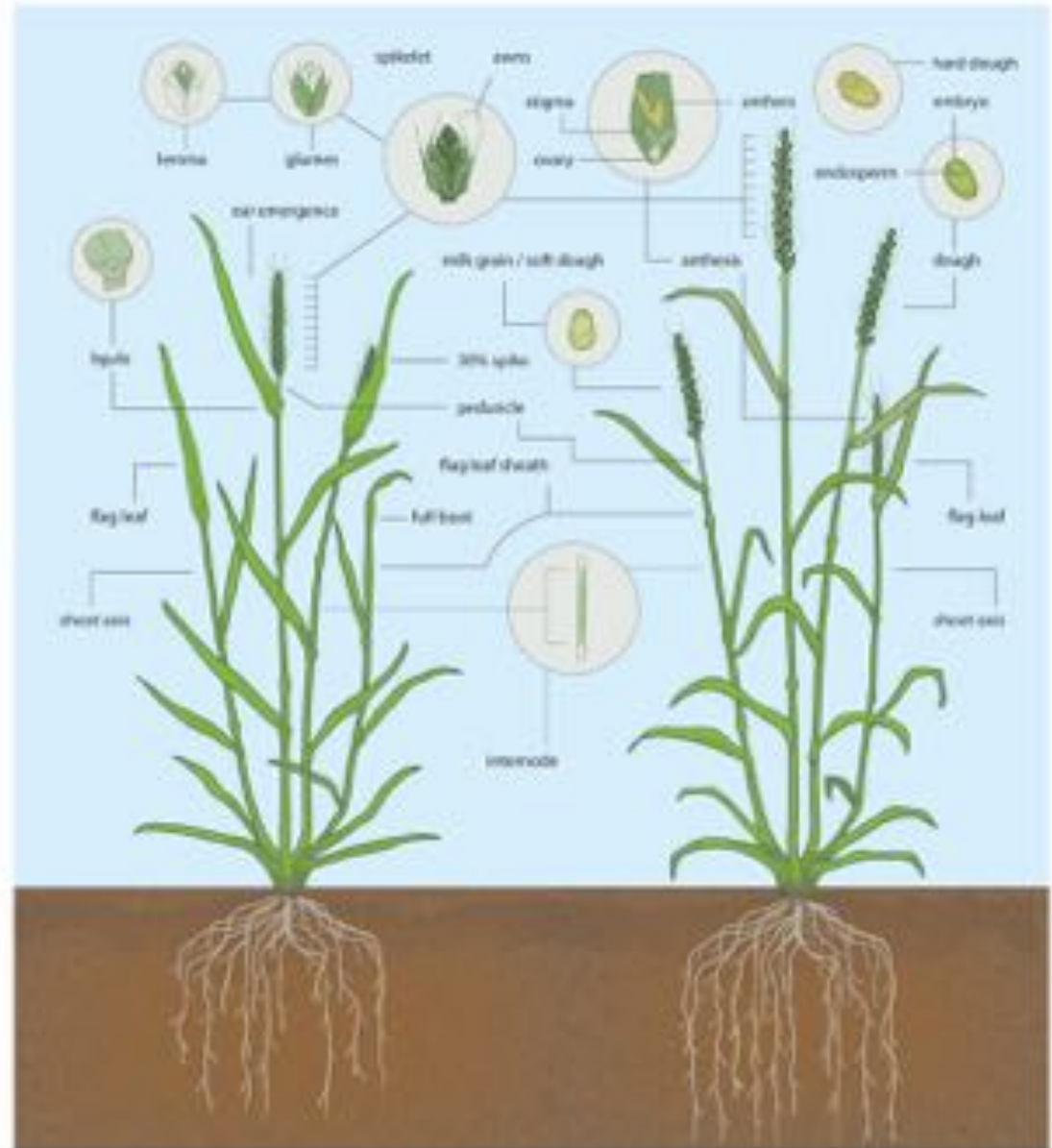
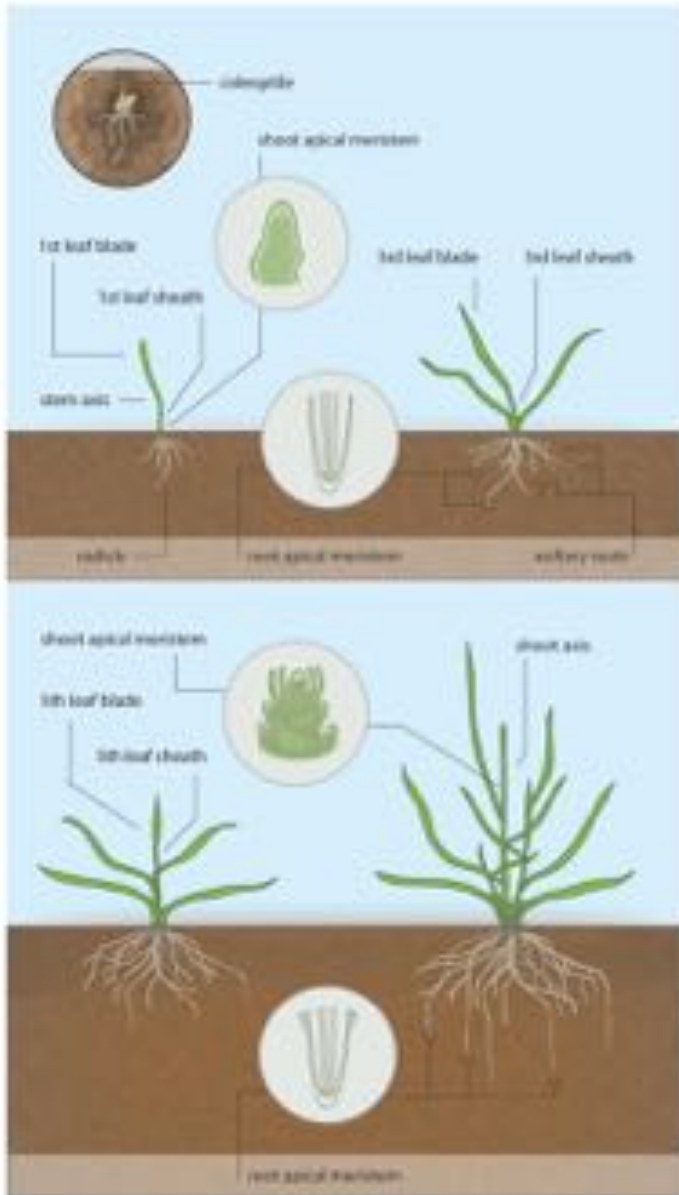
## A roadmap for gene functional characterisation in crops with large genomes: Lessons from polyploid wheat

Nikolai M Adamski<sup>1†</sup>, Philippa Borrill<sup>2†</sup>, Jemima Brinton<sup>1†</sup>, Sophie A Harrington<sup>1†</sup>, Clémence Marchal<sup>1†</sup>, Alison R Bentley<sup>2</sup>, William D Bovill<sup>4</sup>, Luigi Cattivelli<sup>3</sup>, James Cockram<sup>2</sup>, Bruno Contreras-Moreira<sup>4</sup>, Brett Ford<sup>4</sup>, Sreya Ghosh<sup>4</sup>, Wendy Harwood<sup>1</sup>, Keywan Hassani-Pak<sup>2</sup>, Sadiye Hayta<sup>1</sup>, Lee T Hickey<sup>4</sup>, Kostya Kanyuka<sup>2</sup>, Julie King<sup>2</sup>, Marco Maccaferri<sup>1,3</sup>, Guy Naamati<sup>4</sup>, Curtis J Pozniak<sup>1†</sup>, Ricardo H Ramirez-Gonzalez<sup>2</sup>, Carolina Sansaloni<sup>1,2</sup>, Ben Trevaskis<sup>4</sup>, Luzie U Wingen<sup>1</sup>, Brande BH Wulff<sup>1</sup>, Cristobal Uauy<sup>1\*</sup>



Borrill *et al.*, 2019, *Plant J*  
Adamski *et al.*, 2020, *eLIFE*

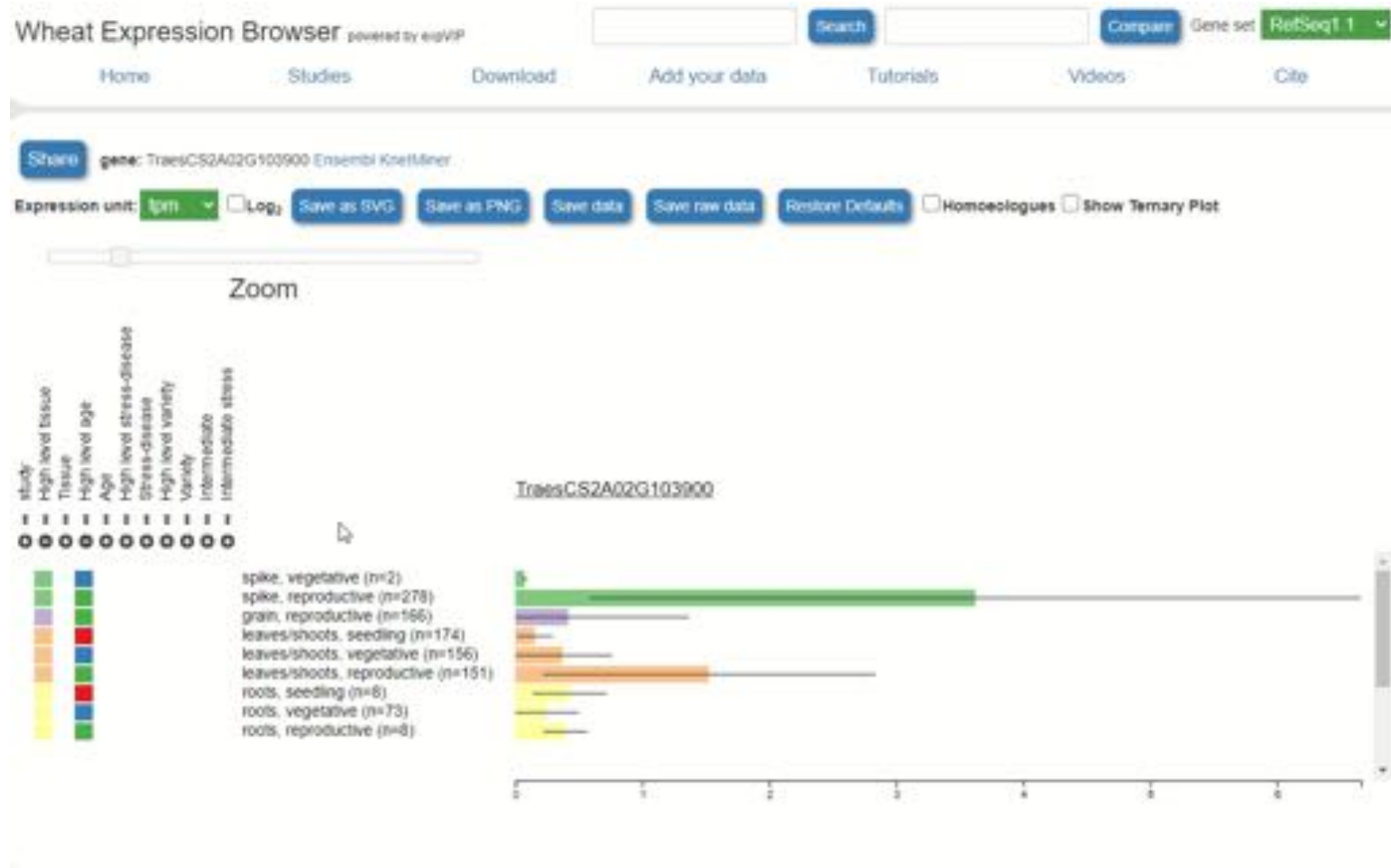
Over 850 samples; developmental time course, etc





# [www.wheat-expression.com](http://www.wheat-expression.com)

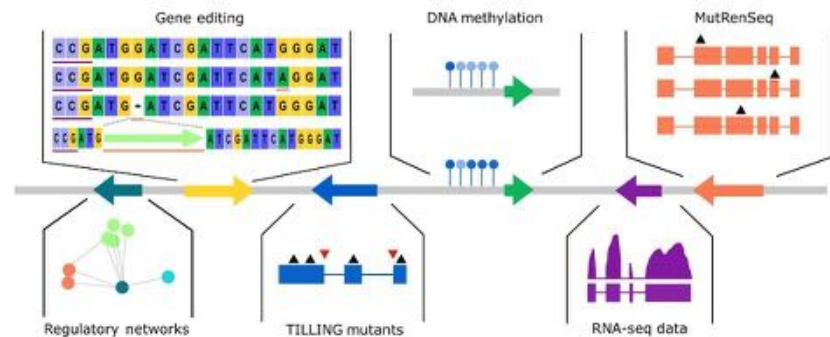
## >1,000 wheat RNA-Seq samples





## 2) Why now?

- Wheat genome sequence
- Functional genomic resources





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Gene editing



Natural/  
induced  
variation



Need to predict gene function

Grain yield



Grain quality

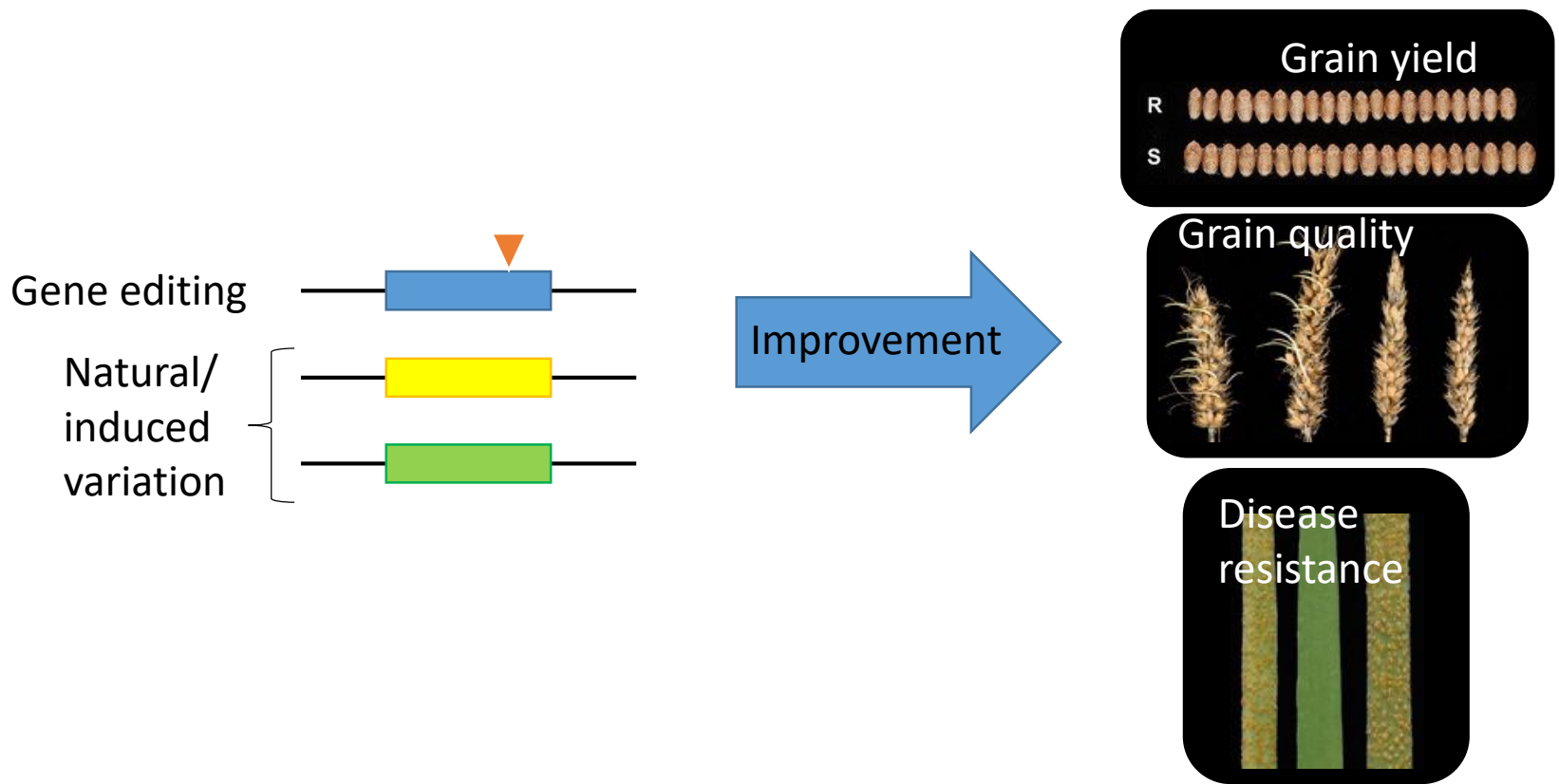
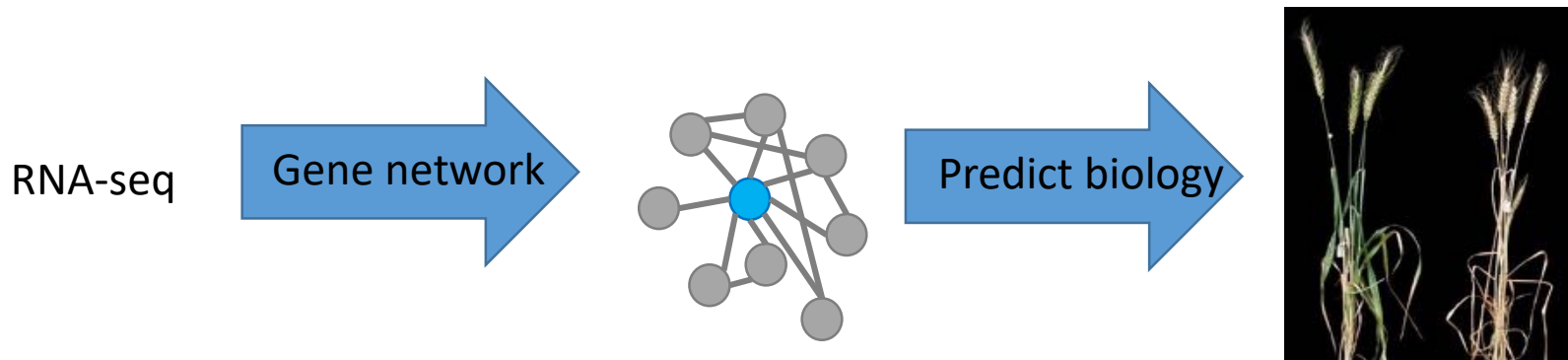


Disease resistance



## Genomics

- Pangenomics
- >7,000 RNA-Seq samples
- Epigenetics







28<sup>th</sup> June



15<sup>th</sup> July



22<sup>nd</sup> July



29<sup>th</sup> July

# Senescence



28<sup>th</sup> June



15<sup>th</sup> July

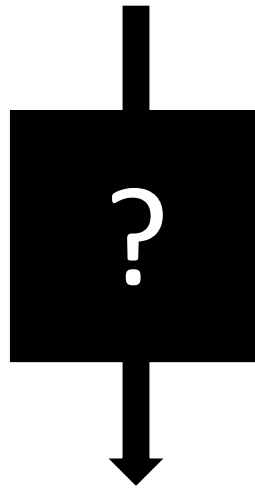


22<sup>nd</sup> July

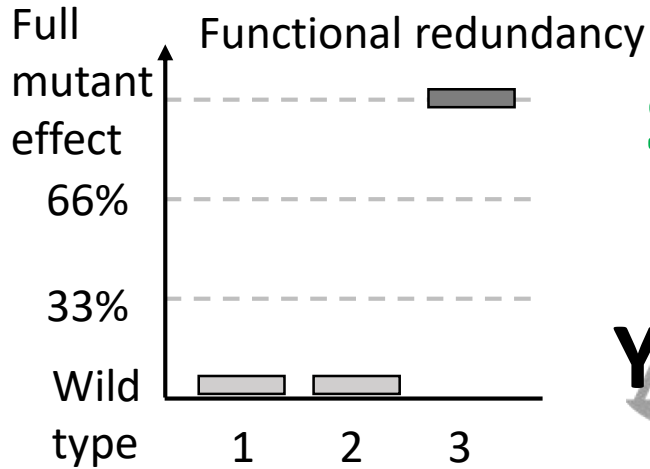
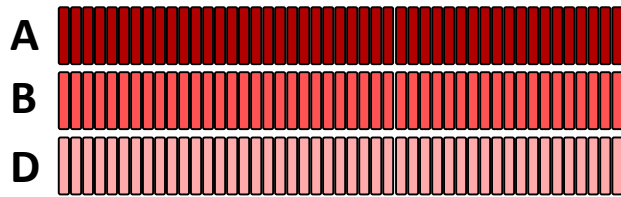
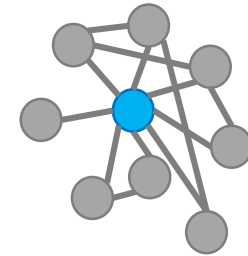


29<sup>th</sup> July

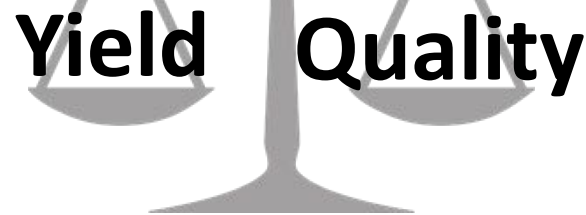
# Genes



Gene regulatory network modelling

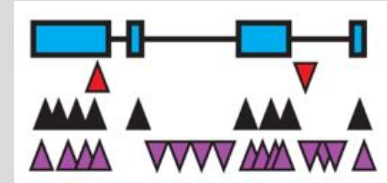
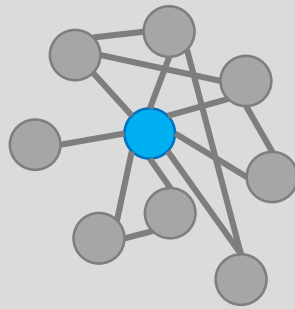
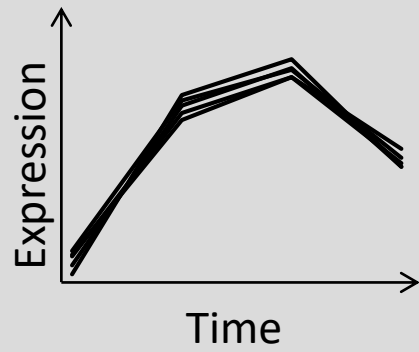


## Senescence





# Transcriptional networks



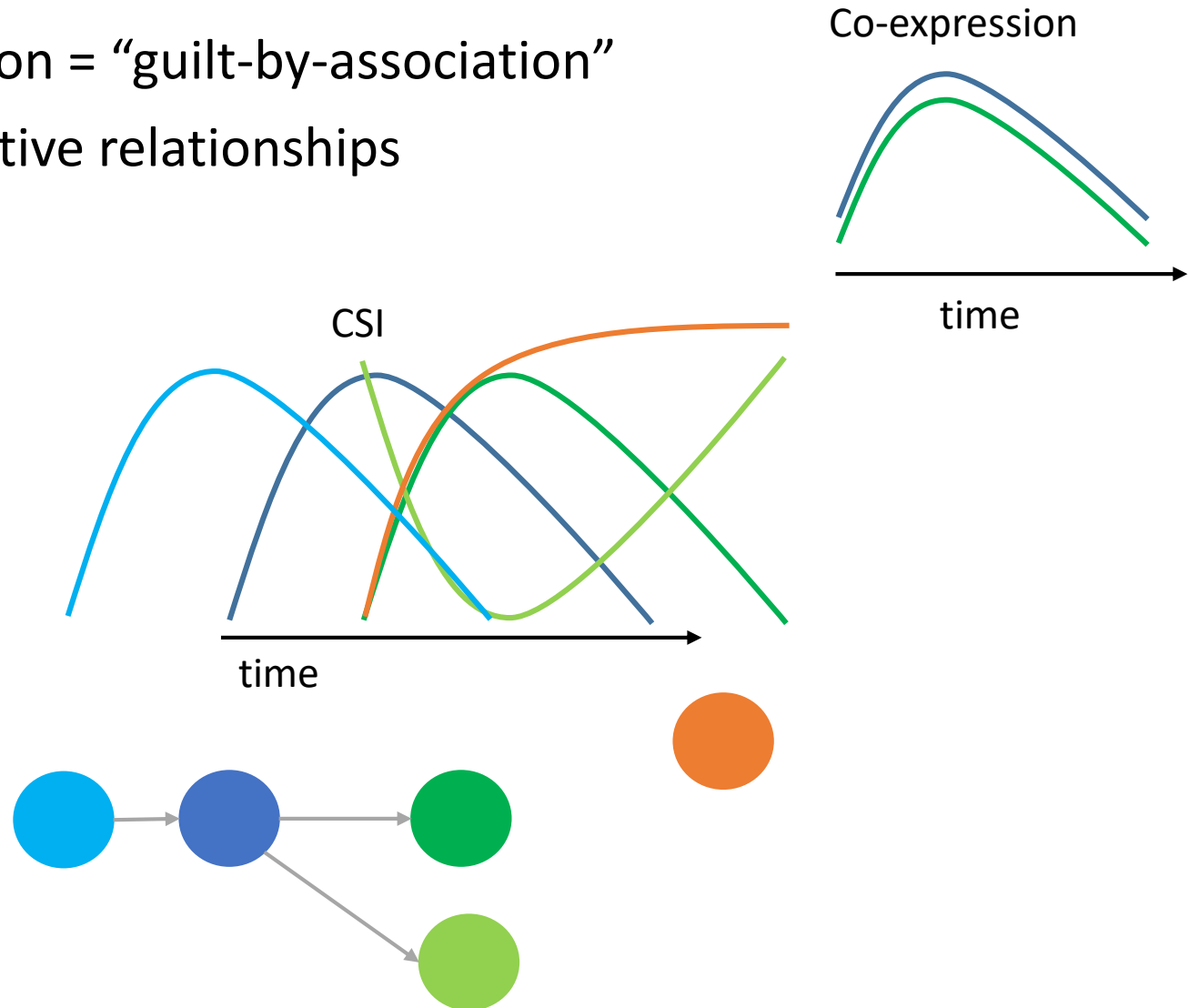
3    7    10    13    15    17    19    21    23    26

Days after anthesis



# Predictive network modelling with CSI

- Co-expression = “guilt-by-association”
- CSI = predictive relationships



# Computationally expensive therefore need to prioritise genes to include in CSI

110,790 genes

↓ Expressed

52,905

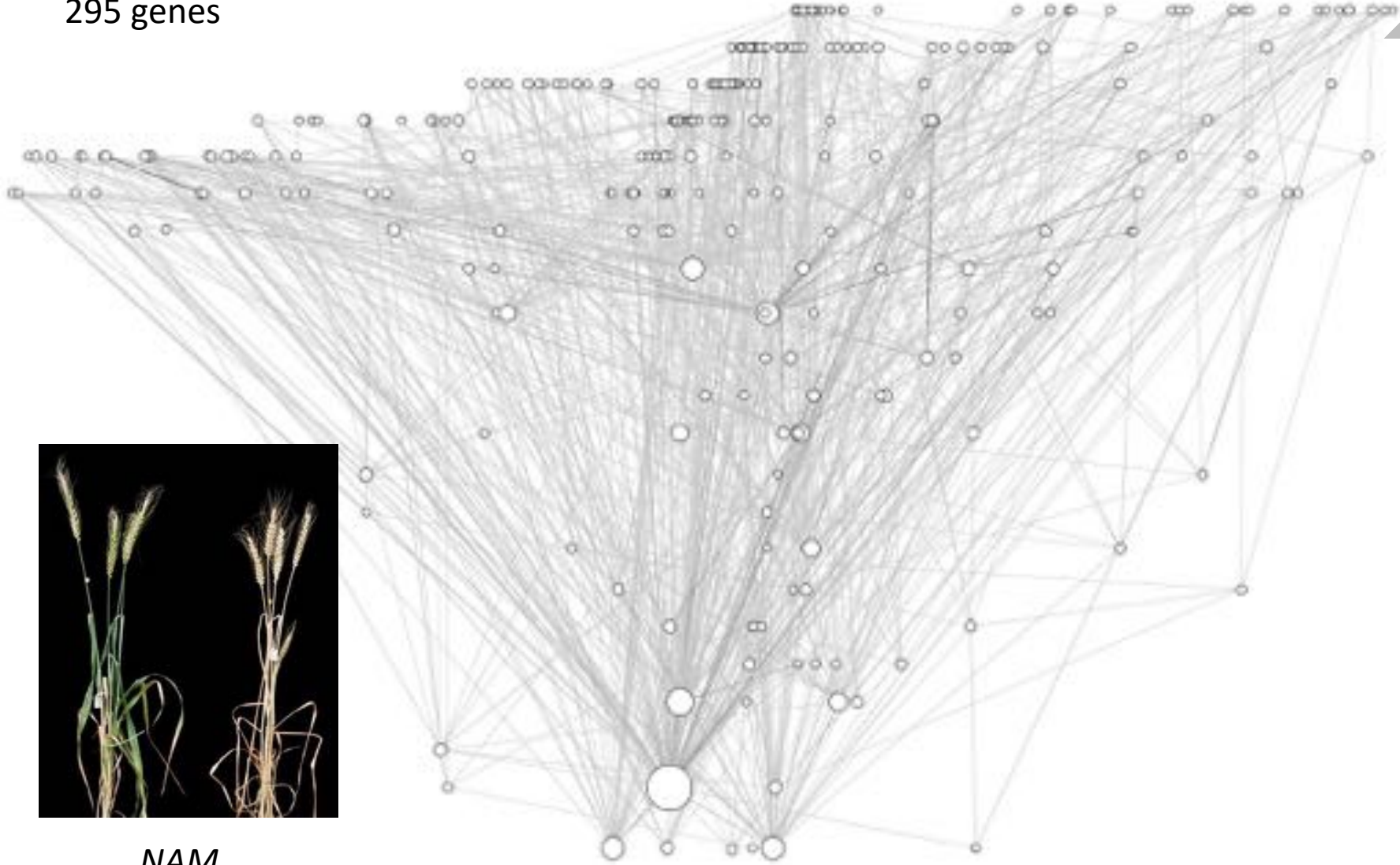
↓ Differentially expressed

9,533

↓ TFs

341 genes

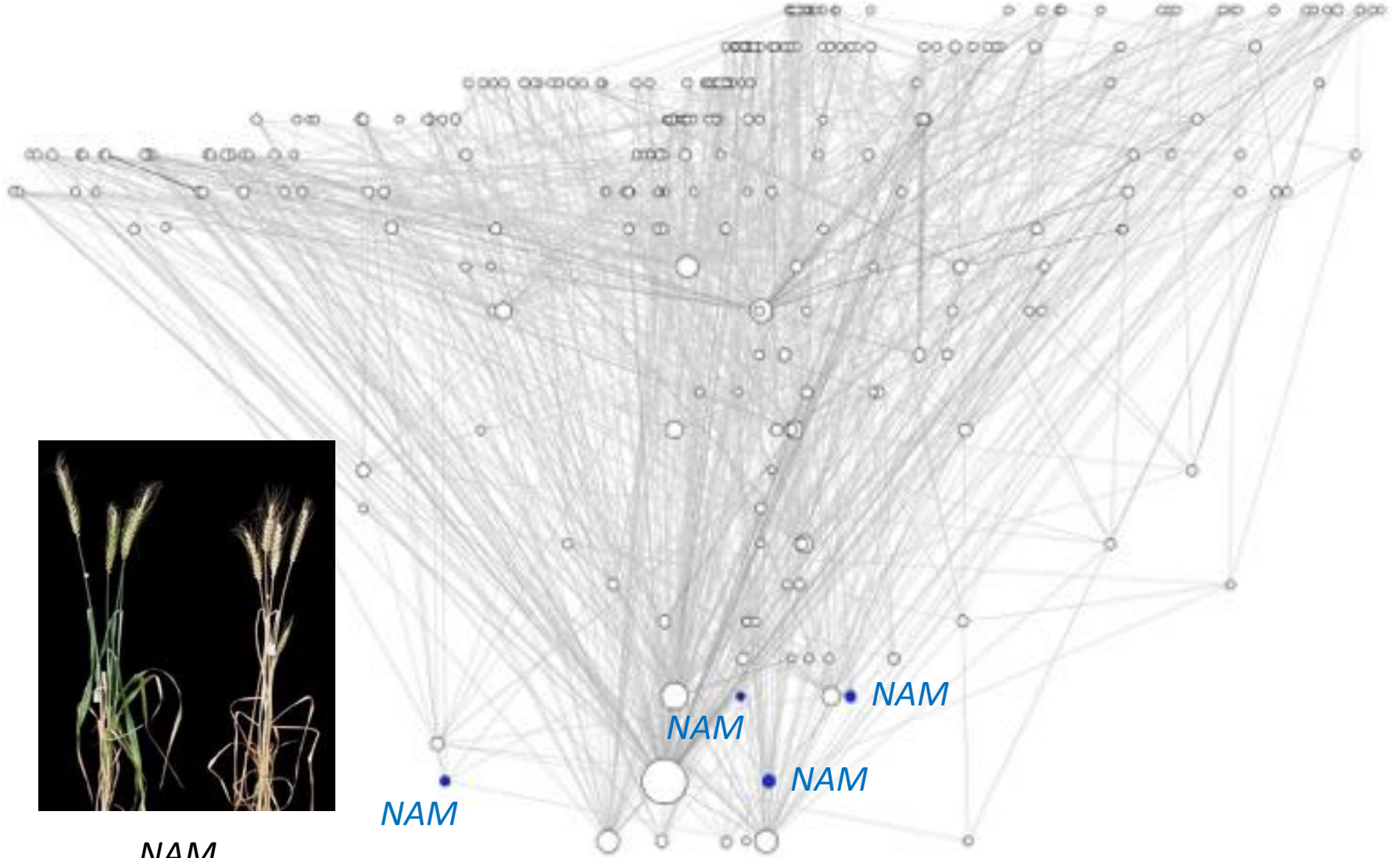
295 genes



*NAM*

Uauy *et al.*, 2006, *Science*

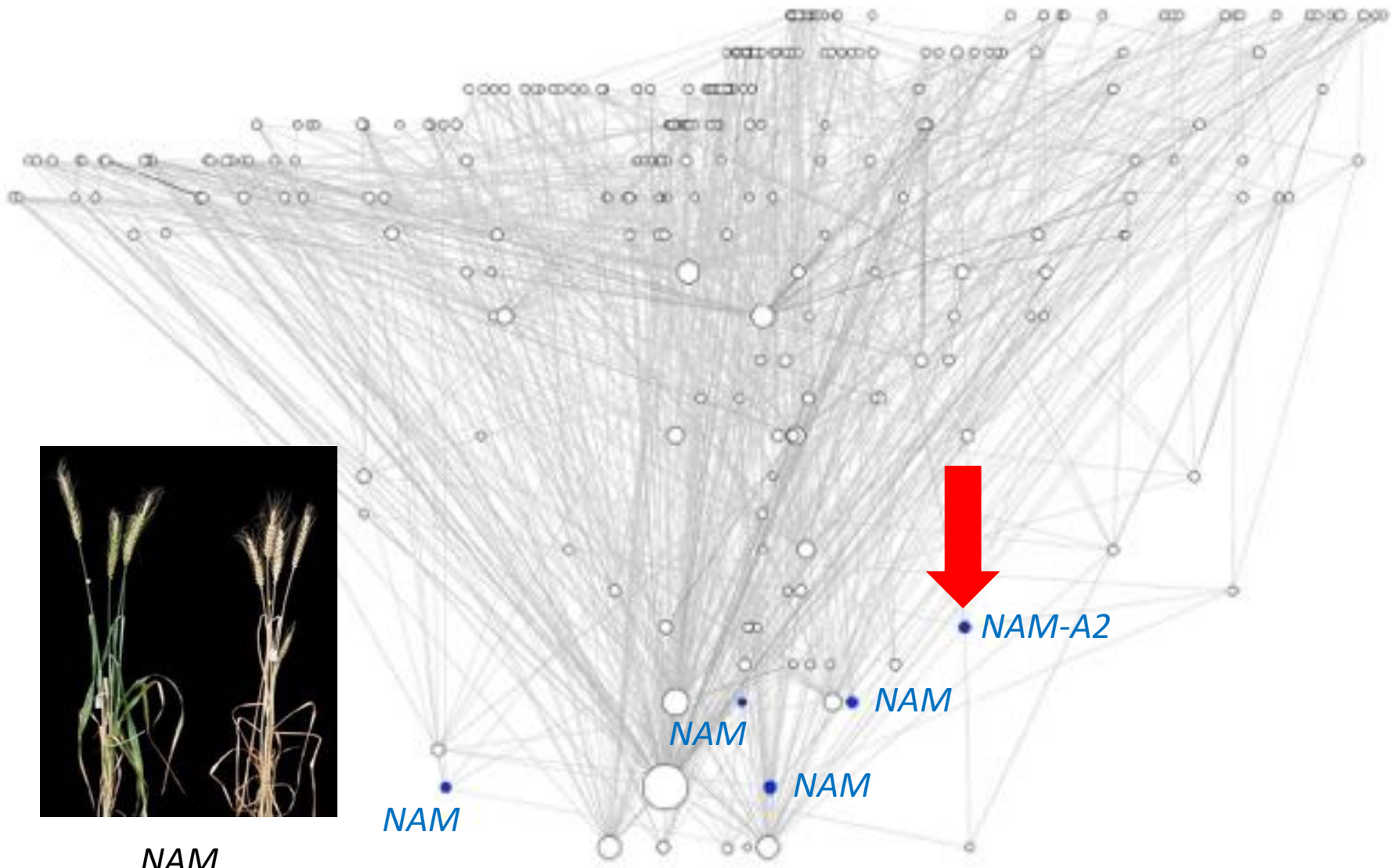
Borrill *et al.*, 2019, *Plant Physiol.*



NAM

Uauy *et al.*, 2006, *Science*

Borrill *et al.*, 2019, *Plant Physiol.*



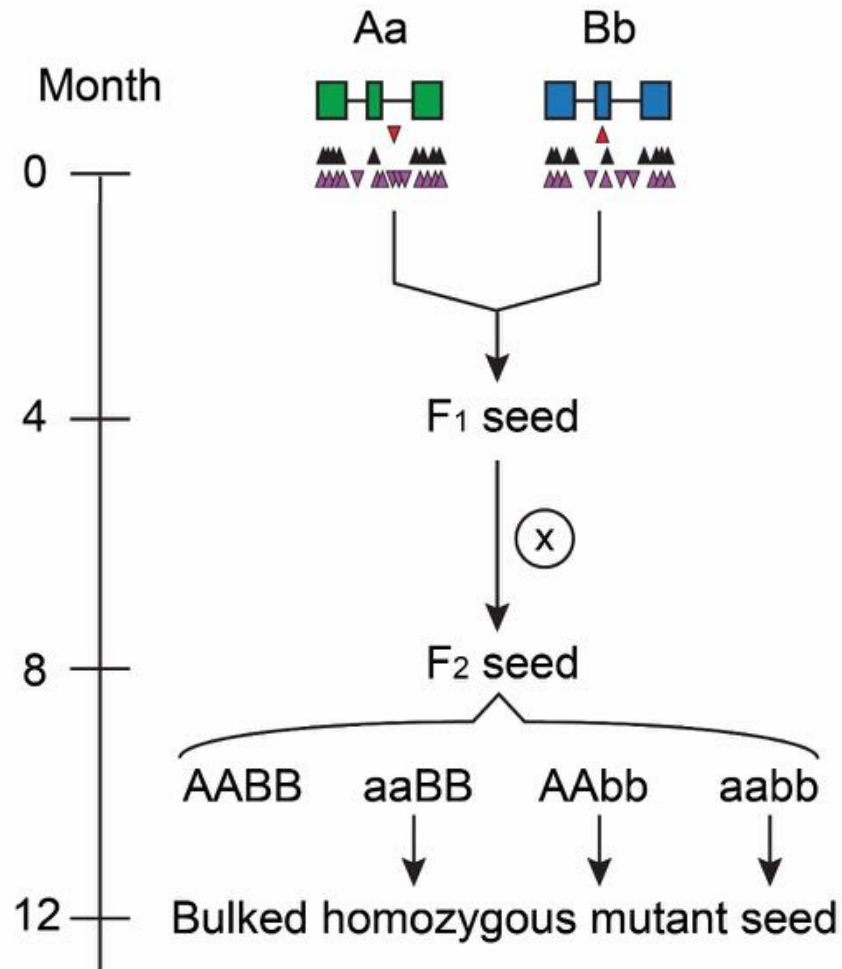
NAM

Uauy et al., 2006, *Science*

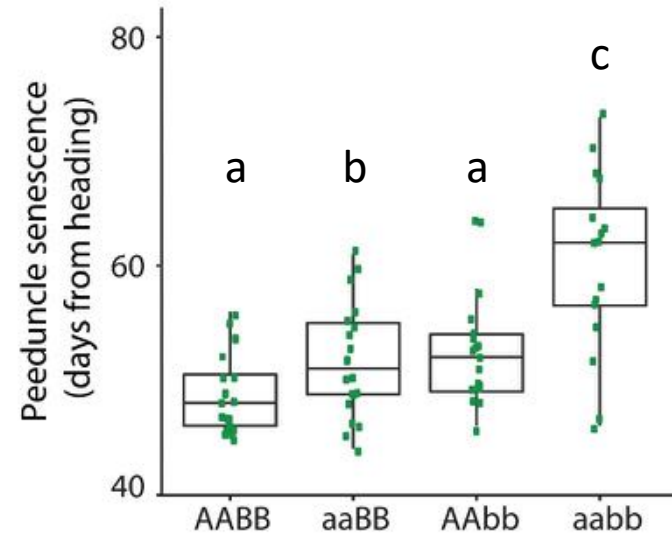
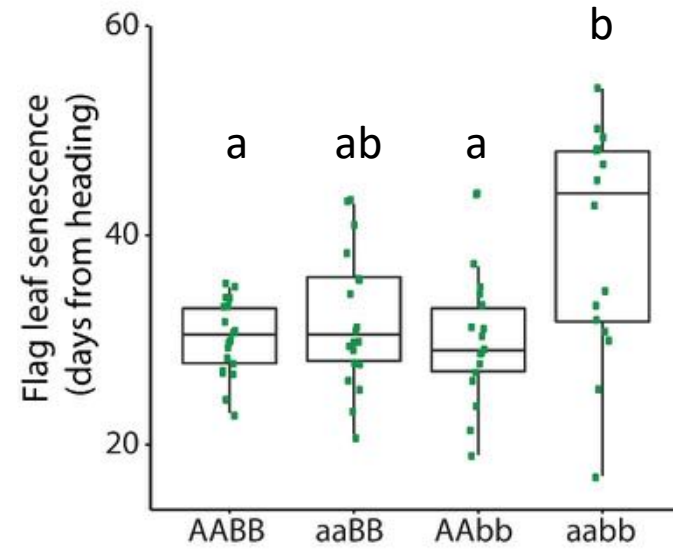
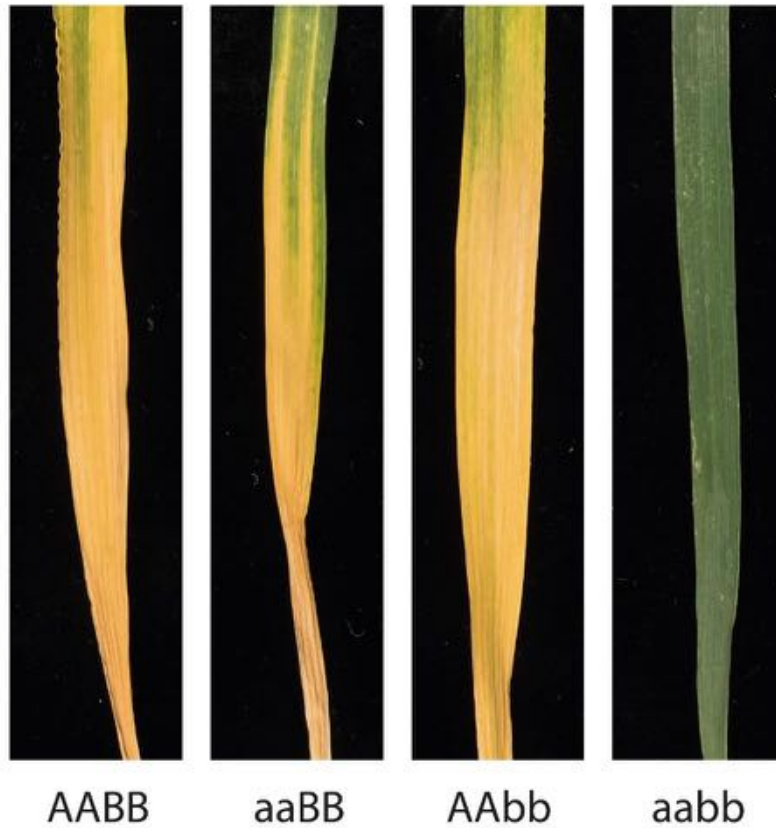
Borrill et al., 2019, *Plant Physiol.*

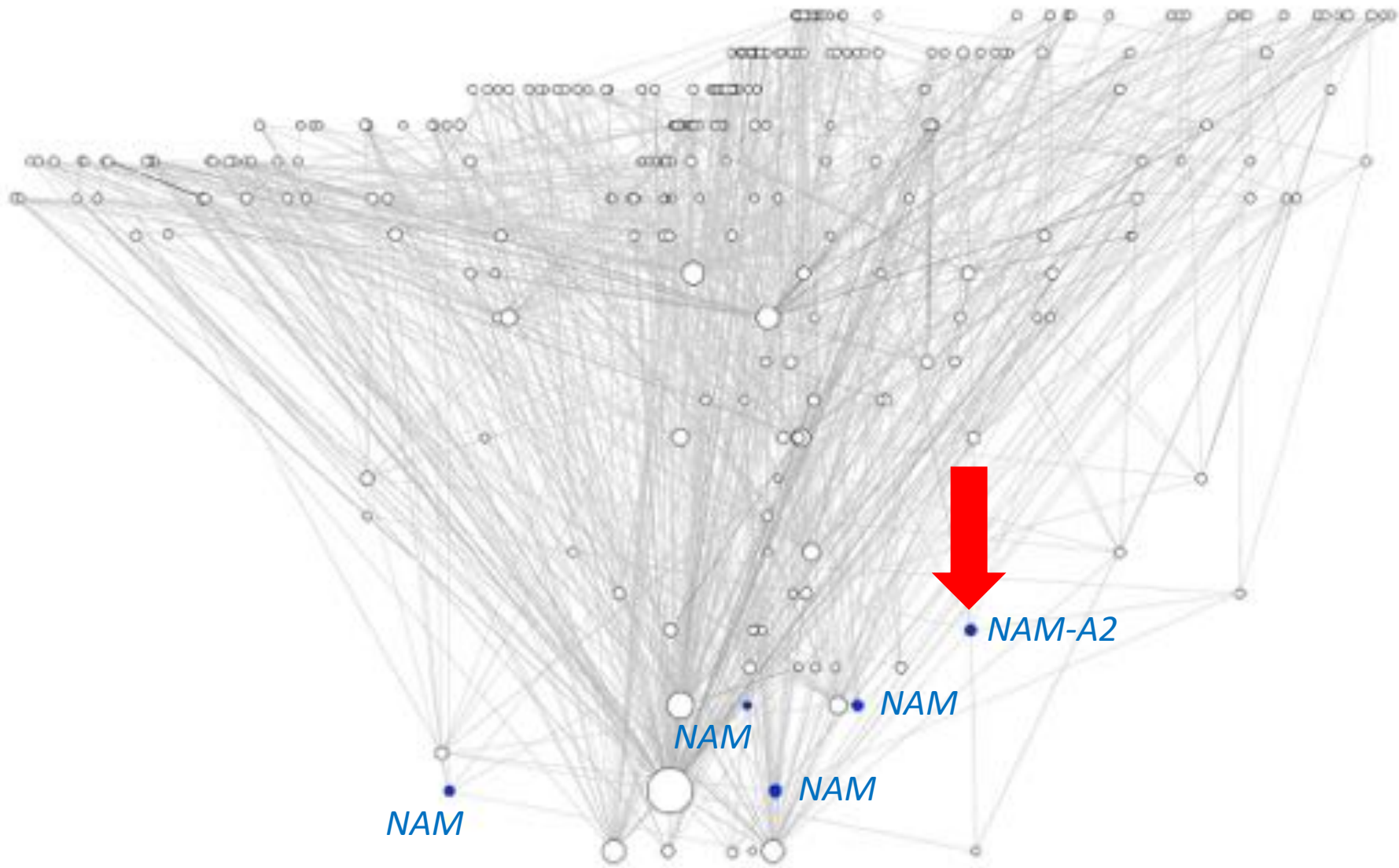


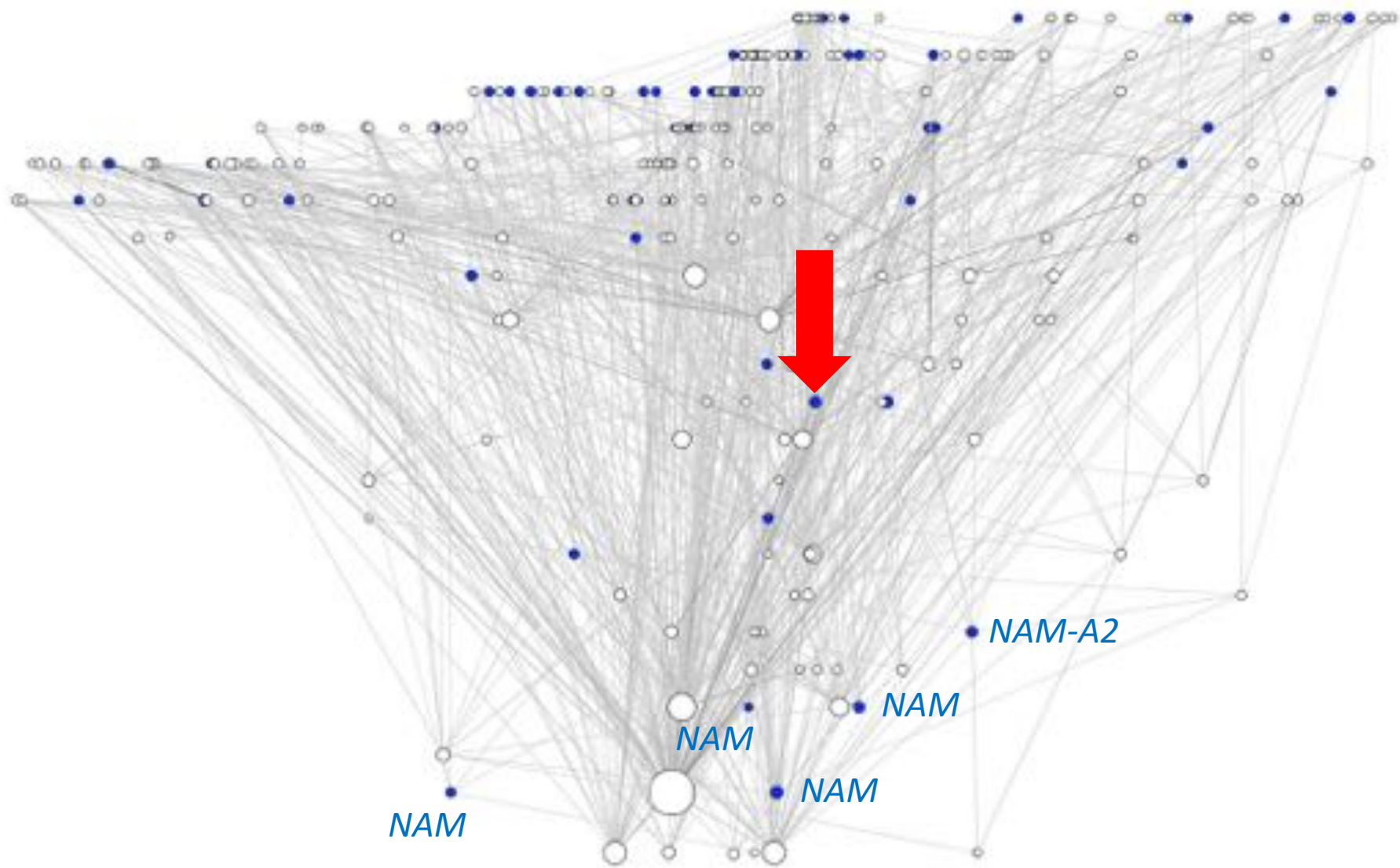
# Develop double mutants for candidate genes

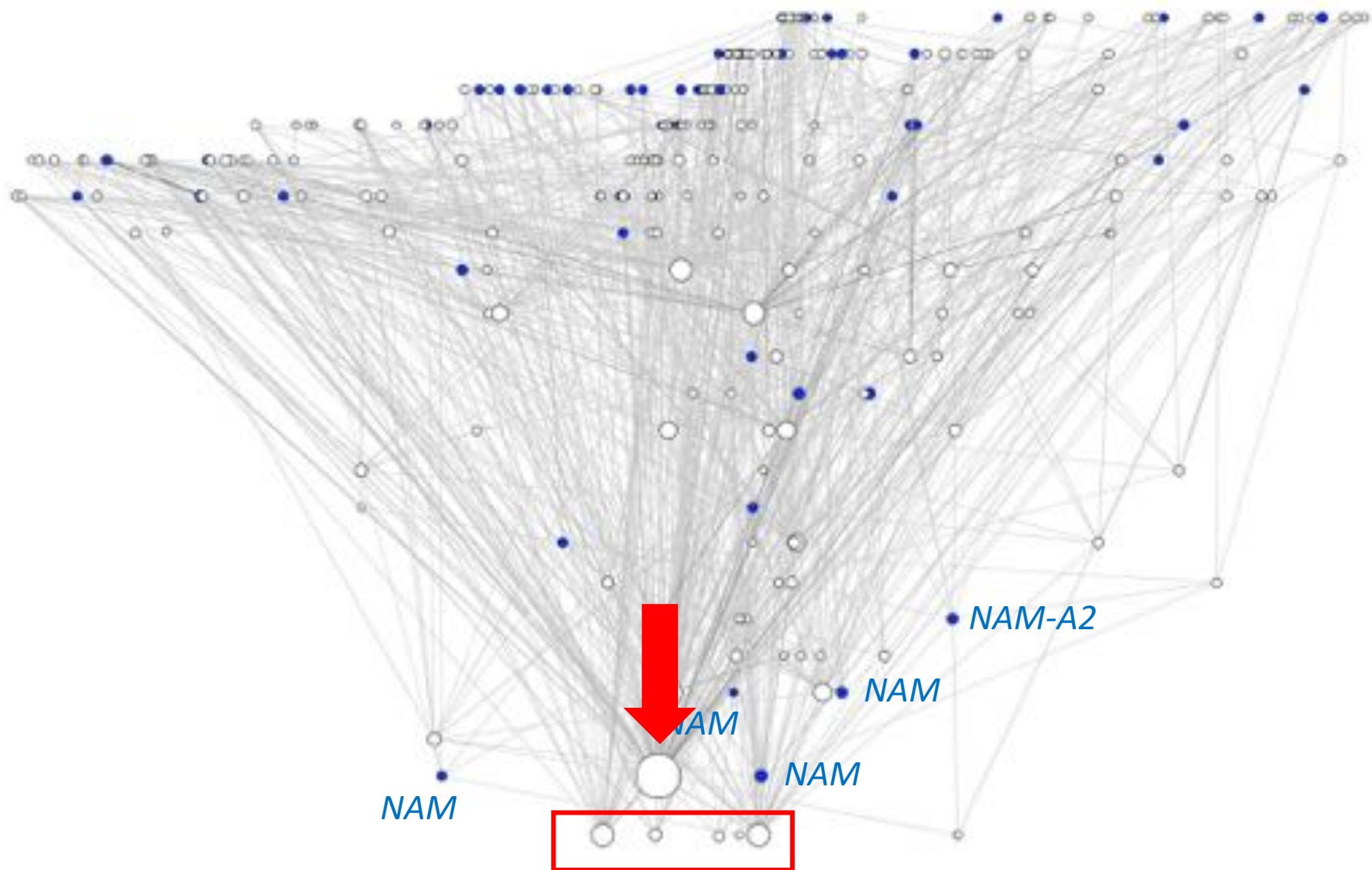


# NAM-2 phenotyping











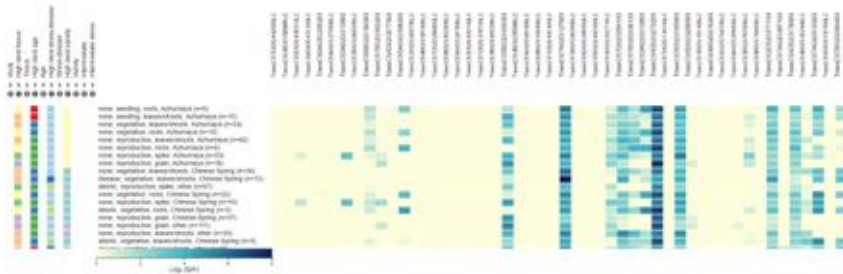
# Layering additional information

>200 gene candidates

How to choose which to prioritise?

Other expression data ([www.wheat-expression.com](http://www.wheat-expression.com))

>1,000 RNA-Seq samples



Ortholog information

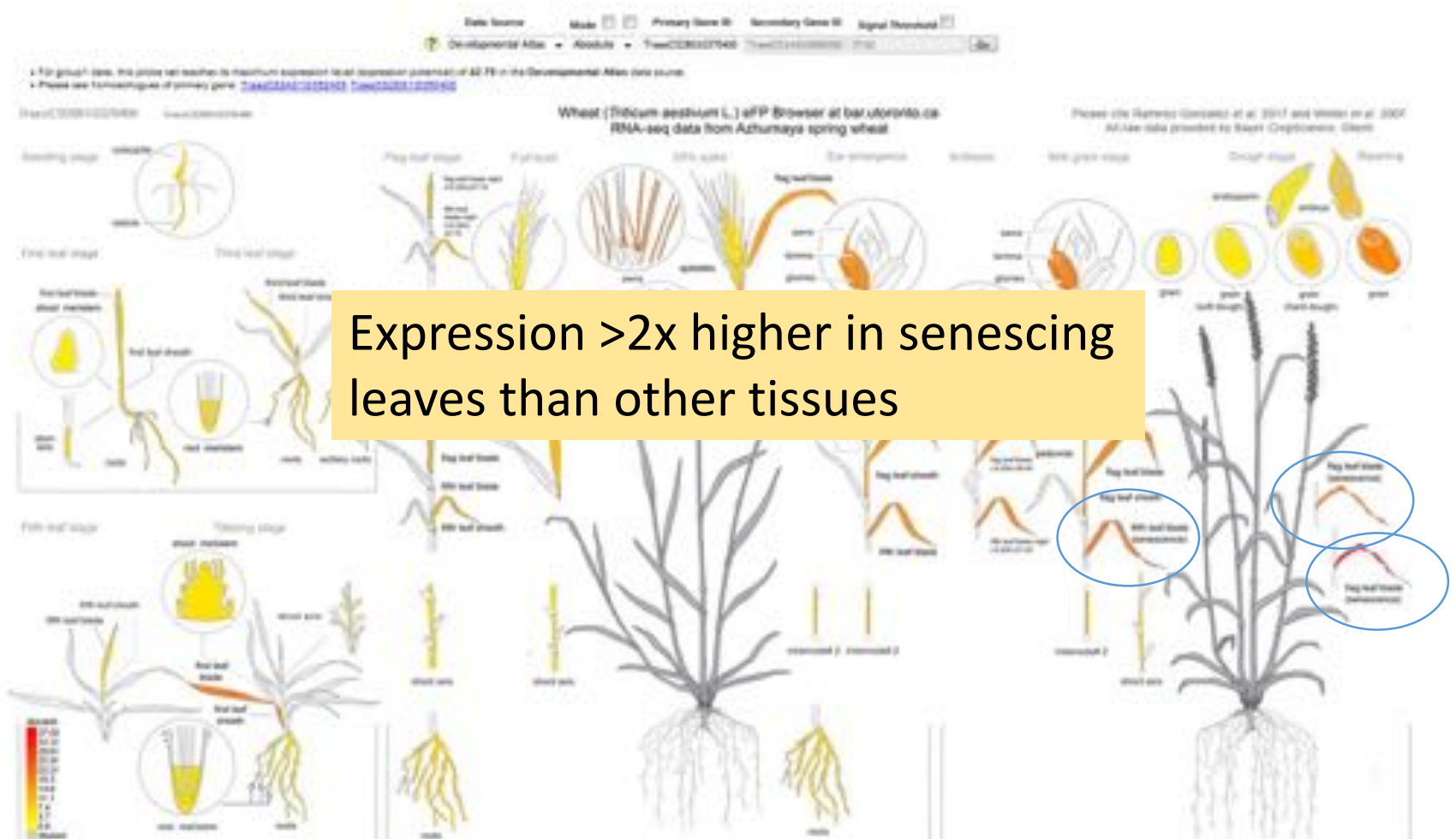


Gene network information

<http://knetminer.rothamsted.ac.uk/>

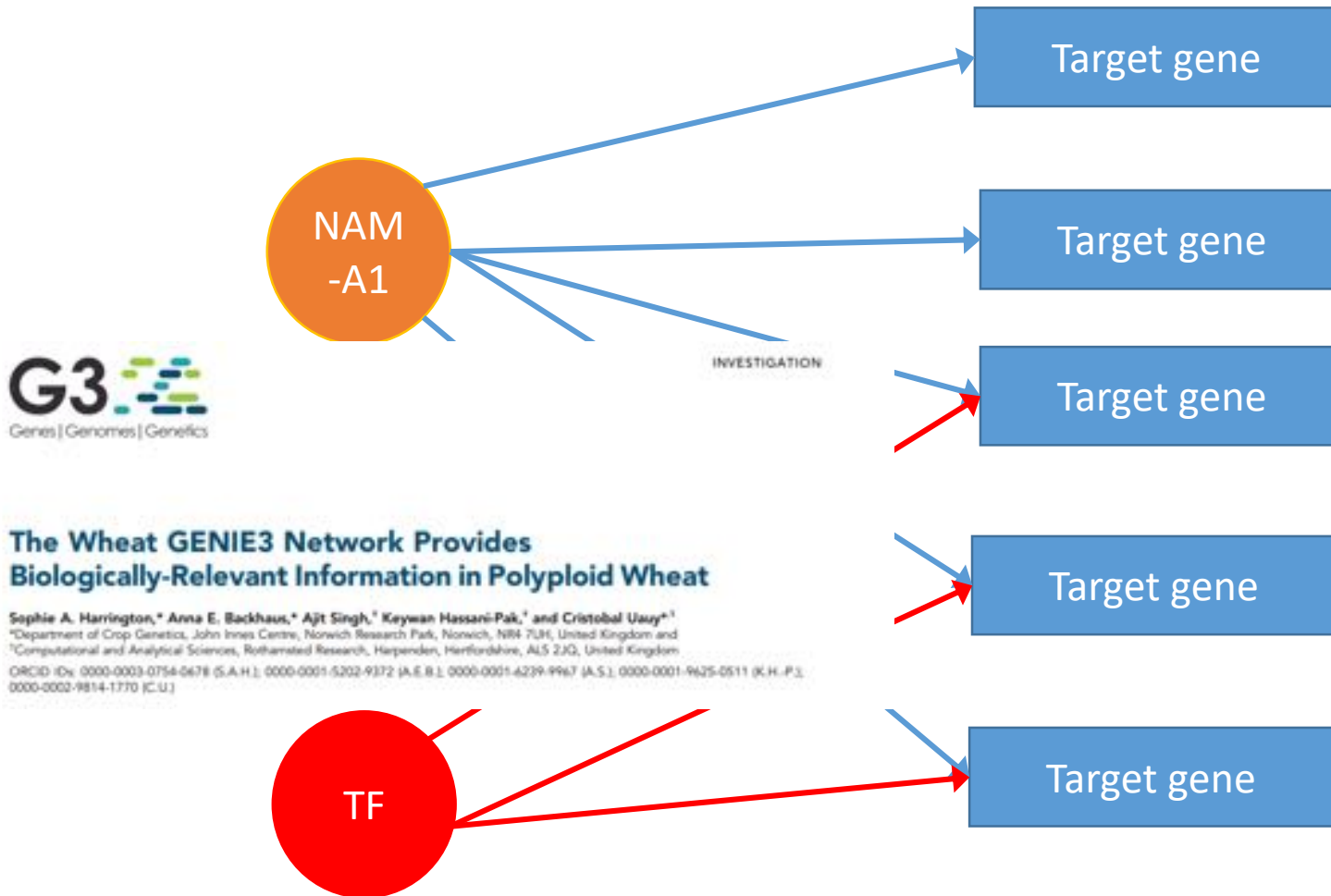
# Wheat eFP

[http://bar.utoronto.ca/efp\\_wheat/cgi-bin/efpWeb.cgi](http://bar.utoronto.ca/efp_wheat/cgi-bin/efpWeb.cgi)



# Genie3 network

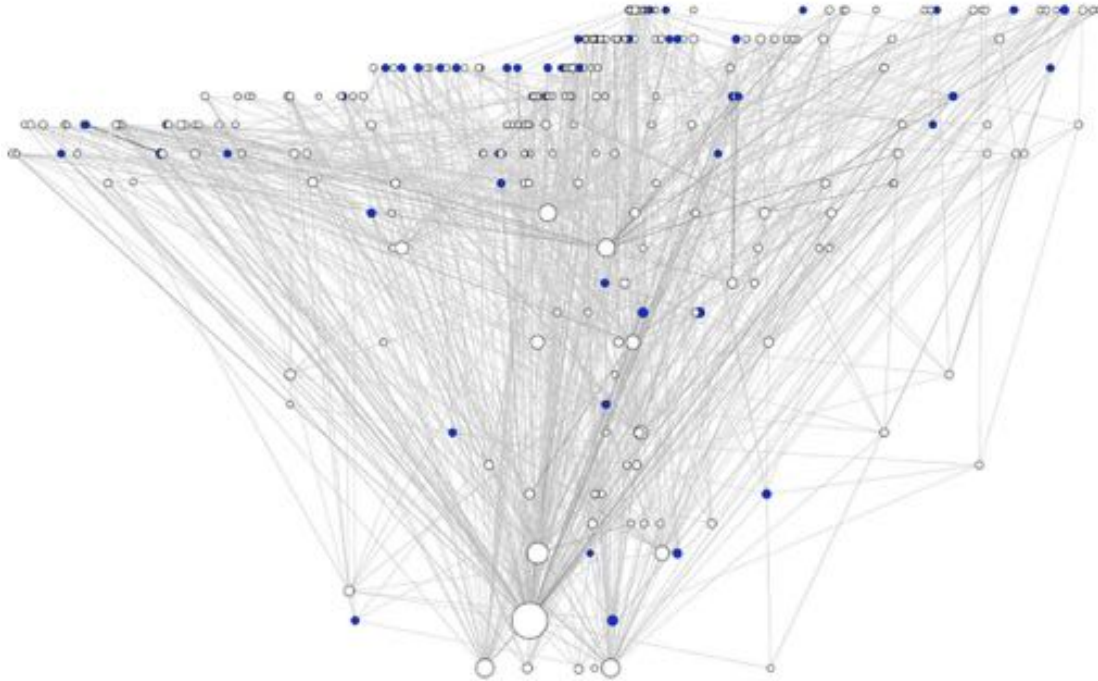
(Available at <http://knetminer.rothamsted.ac.uk/>)



## The Wheat GENIE3 Network Provides Biologically-Relevant Information in Polyploid Wheat

Sophie A. Harrington,<sup>\*</sup> Anna E. Backhaus,<sup>\*</sup> Ajit Singh,<sup>\*</sup> Keywan Hassani-Pak,<sup>†</sup> and Cristóbal Uauy<sup>\*†</sup>  
<sup>\*</sup>Department of Crop Genetics, John Innes Centre, Norwich Research Park, Norwich, NR4 7UH, United Kingdom and  
<sup>†</sup>Computational and Analytical Sciences, Rothamsted Research, Harpenden, Hertfordshire, AL5 2JQ, United Kingdom  
ORCID iDs: 0000-0003-0754-0678 (S.A.H.); 0000-0001-5202-9372 (A.E.B.); 0000-0001-4239-9967 (A.S.); 0000-0001-9625-0511 (K.H.-P.); 0000-0002-9814-1770 (C.U.)

# Future TFs to characterise



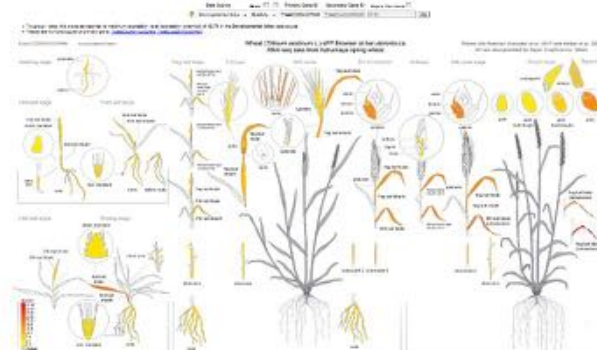
20 TFs with additional evidence

Rice orthologs frequently involved in:

- Nitrate response
- Drought

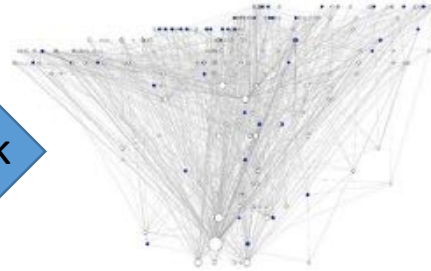


### 3) Can we use gene networks to escape polyploidy?

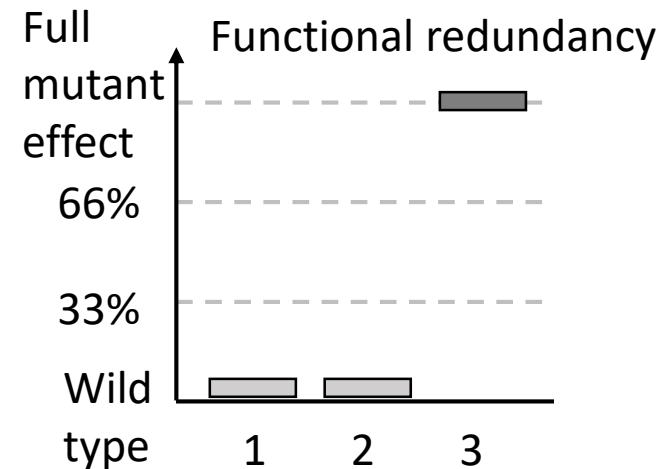
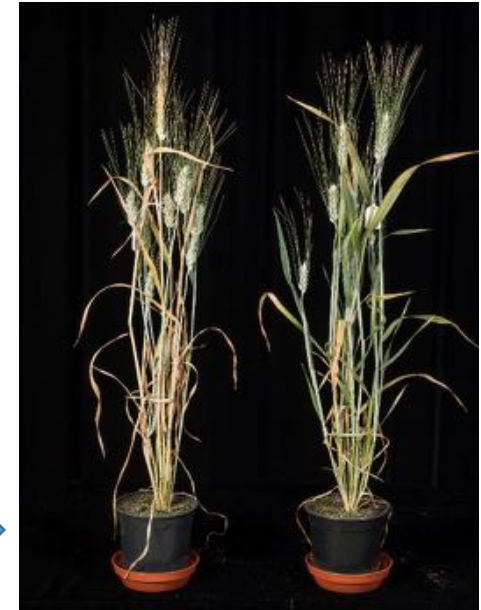


RNA-seq

Gene network



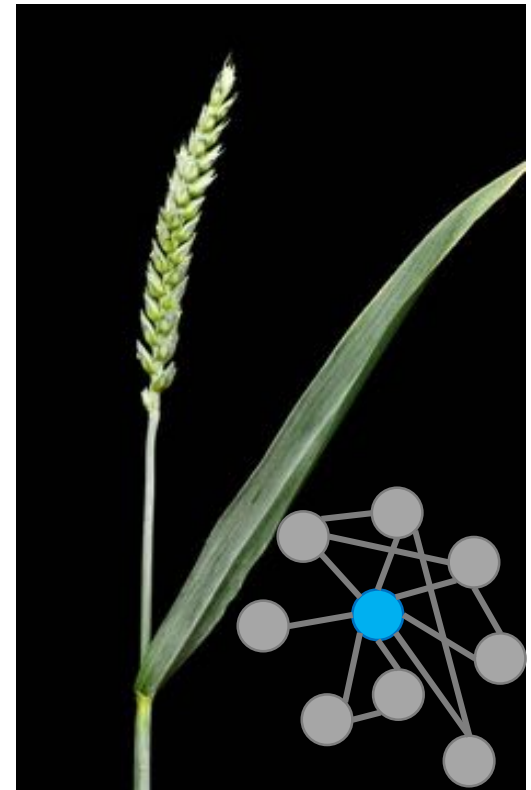
Predict biology



# Conclusions

- 1) Why is polyploidy relevant?
- 2) Why now?
- 3) Can we use gene networks to escape polyploidy?

- Polyploidy challenge and opportunity
- New resources for wheat (genome, TILLING, expression)
- Gene networks can unmask polyploidy



# Acknowledgements

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**Cristobal Uauy**



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Matt Scott  
James Simmonds



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# Acknowledgements

## Conclusions

- 1) Why is polyploidy relevant?  
*Challenge and opportunity*
- 2) Why now?  
*New resources for wheat*
- 3) Can we use gene networks  
to escape polyploidy?  
*Gene networks can unmask  
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