

Interdisciplinary Research Cost Function Network for Life Sciences

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Appliquées de Toulouse (MIAT), France

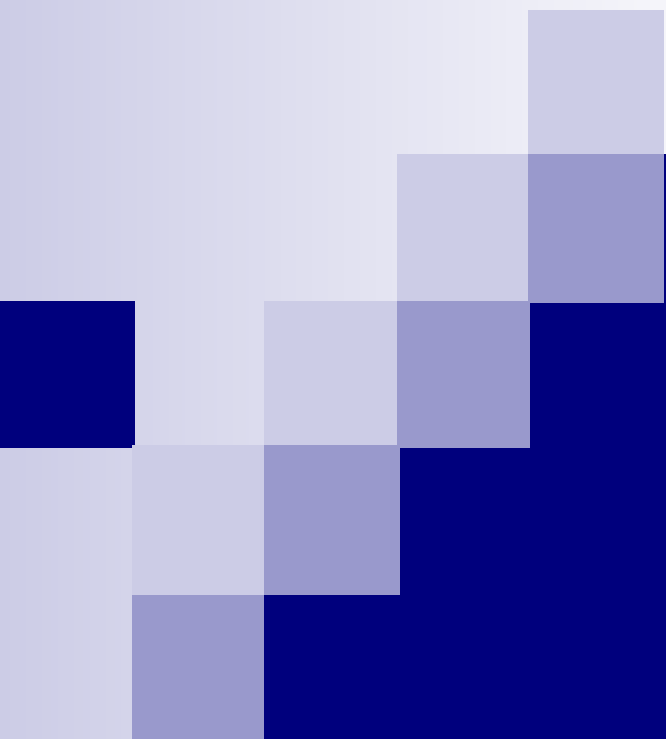


Doctoral Program
CP 2022, Haifa, Israel
08/01/2022



Plan

- Case studies
- *Bioinformatics & CFN*
 - Mendelian error detection
 - Computational Protein Design
- *Agronomy & AI*
 - Mixed Fruit-Vegetable Crop Allocation Problem
- Some tips for your thesis

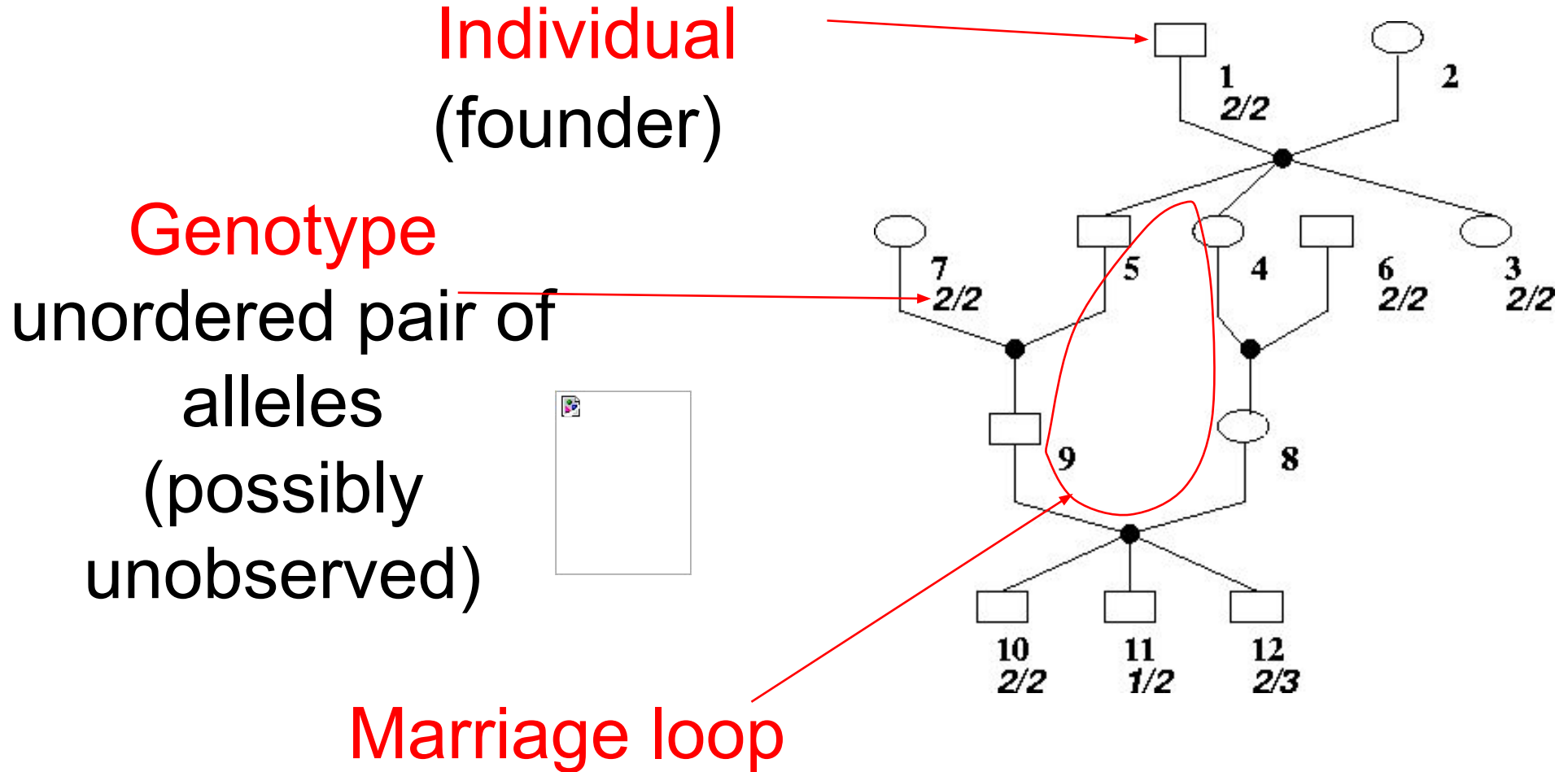


Mendelian error detection in complex pedigree

Simon de Givry, Marti Sanchez,
Isabelle Palhière, Zulma Vitezica,
and Thomas Schiex
INRAE, Toulouse, France

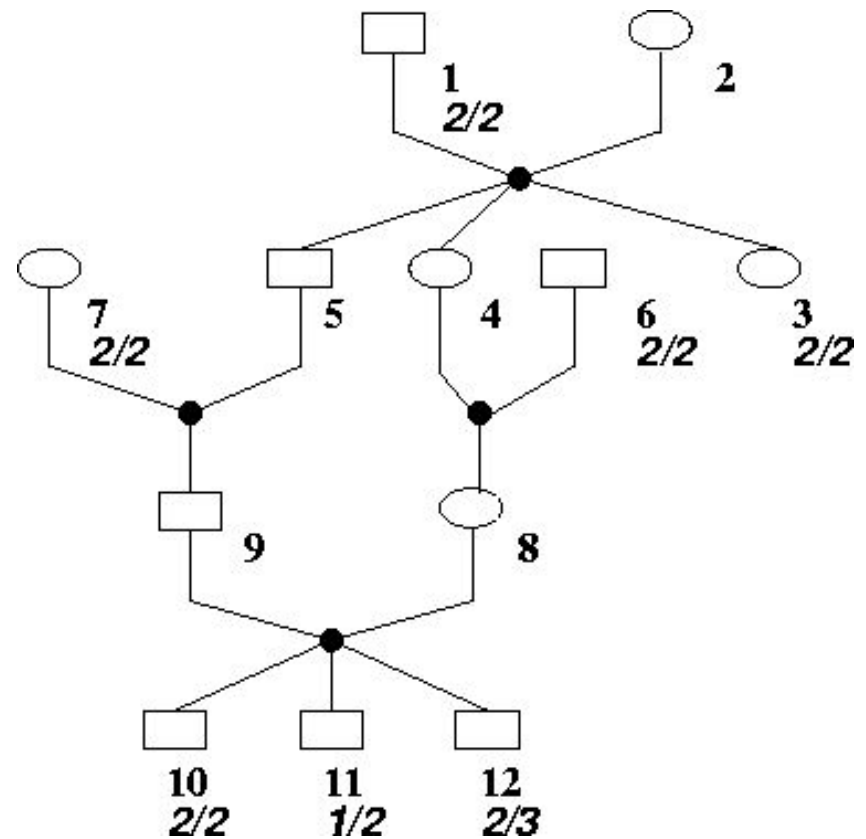
Cleaning the data

*In 2005, about 1% genotyping errors
Today, less than 0.1% (NGS)*



Task 1: Consistency Checking

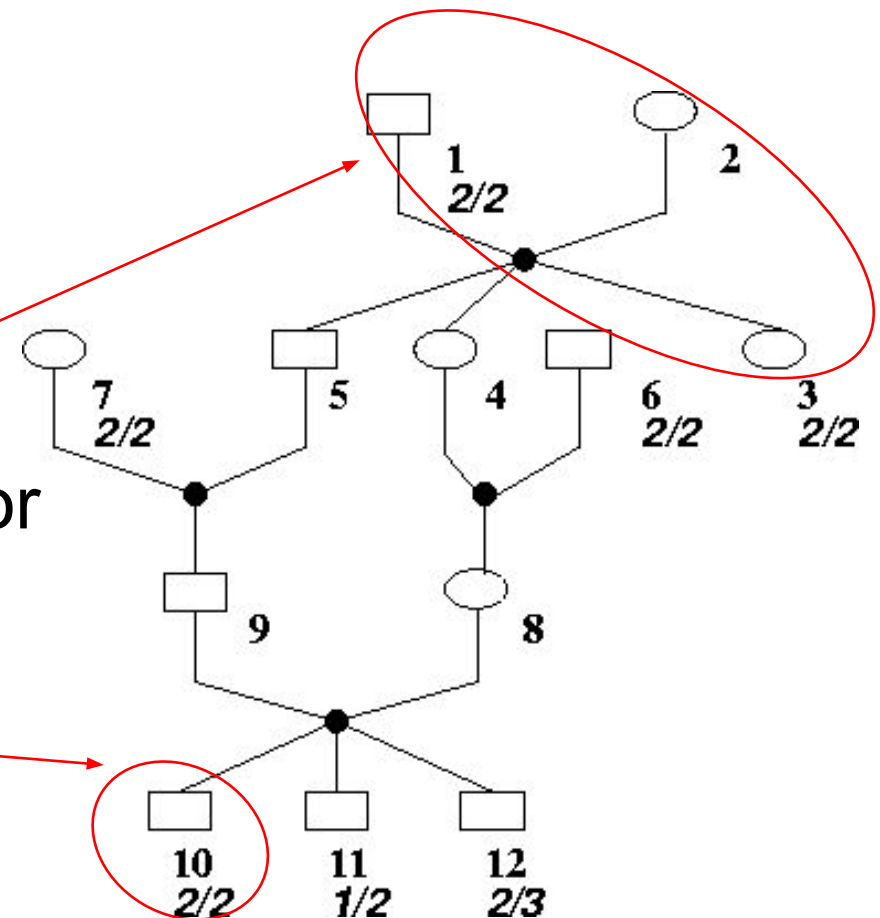
- Assuming the pedigree is correct, checks if it exists a **complete genotype assignment** consistent with the observed genotypes and with the Mendelian laws of inheritance
- Complexity results (Aceto et al., 2003)
 - NP-complete for a pedigree with loops and more than three alleles
 - Polynomial if no loops or just two alleles (SNP)



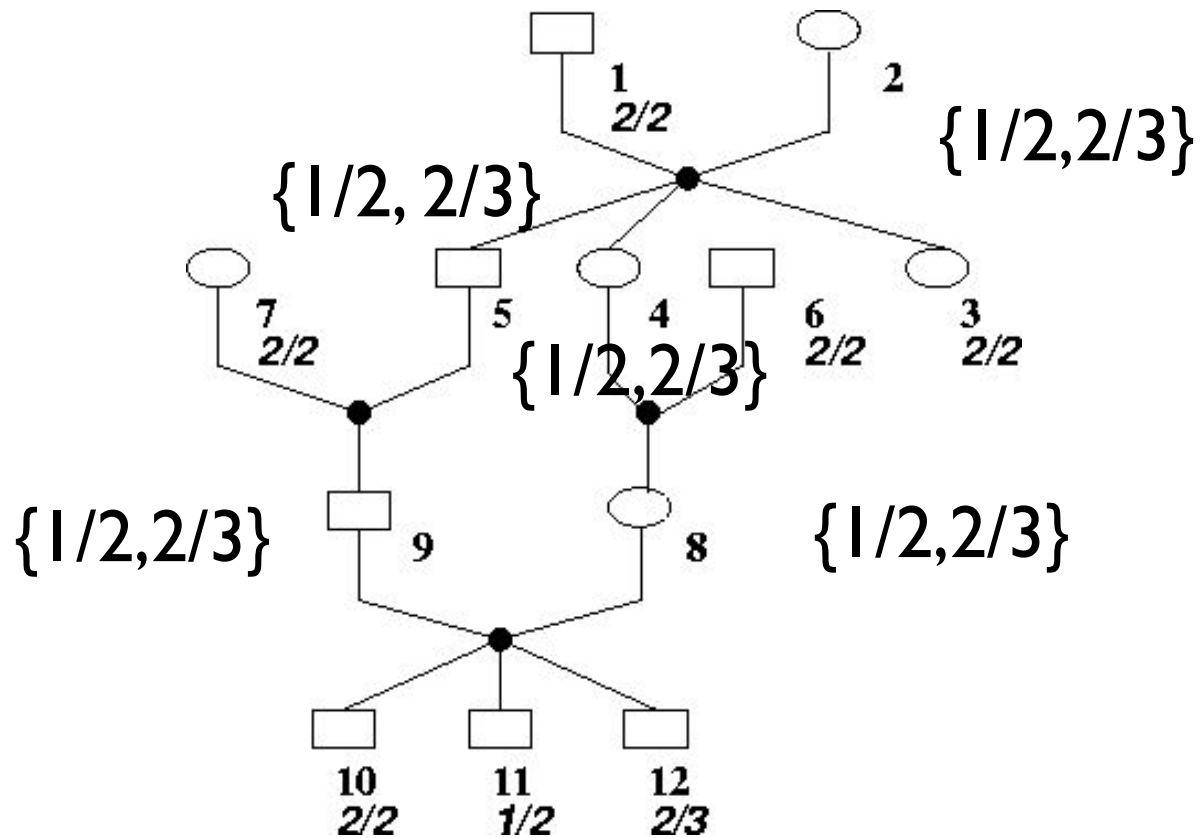
Constraint Satisfaction Problem

- **X**: one variable per individual
- **D**: domain of every variable is defined as the set of all possible genotypes
 $\{ 1/1, 1/2, 1/3, 2/2, 2/3, 3/3 \}$

- **C**:
 - Ternary constraints to encode Mendelian laws for any non founder
 - Unary constraints to encode genotyping data



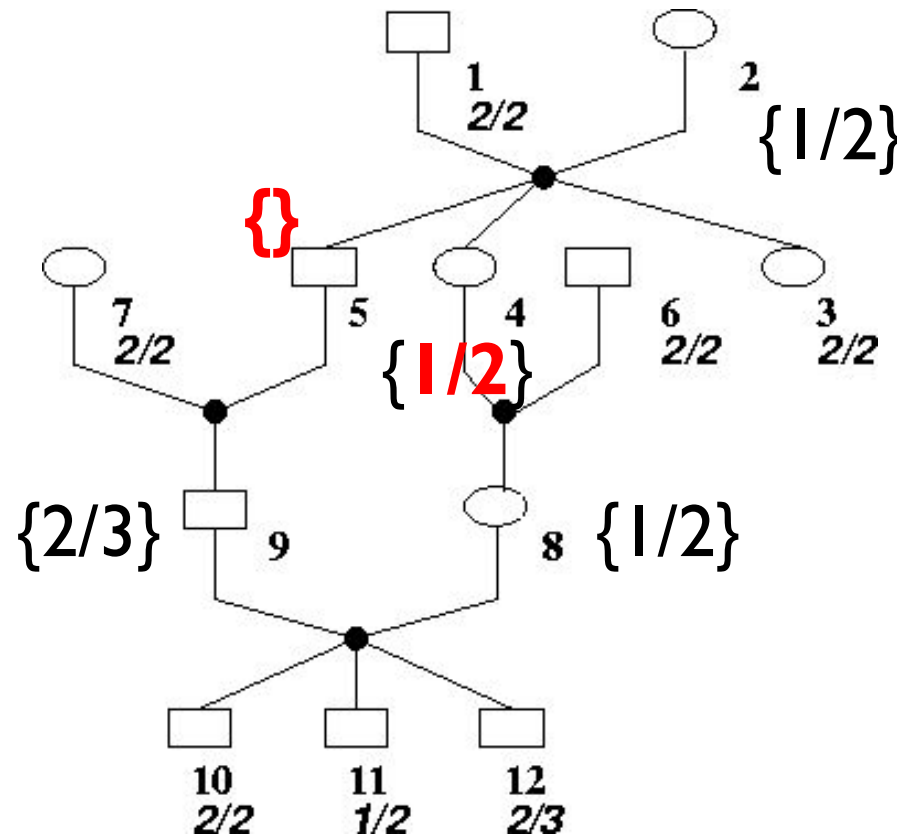
Generalized Arc Consistency



(Mackworth, AIJ 1977)

(Lange, Goradia, Am J Hum Genet 1987)

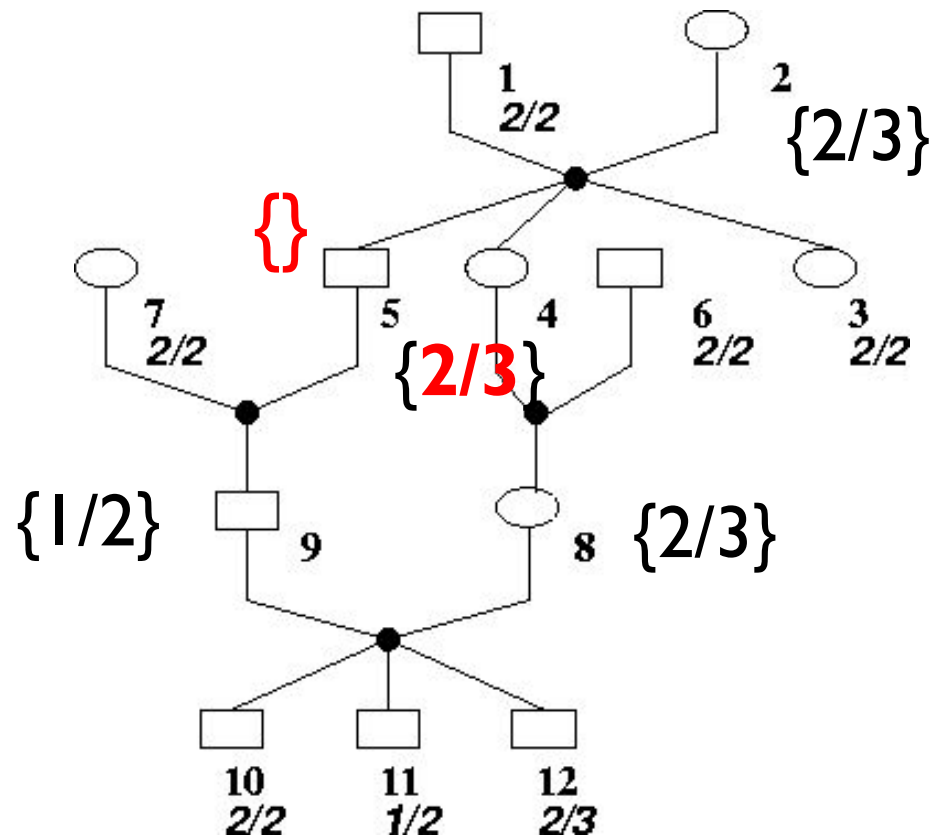
Backtrack search on loop-breaker individuals



(Dechter, AIJ 1990)

(O'Connell, Weeks, Am J Hum Genet 1997)

Backtrack search on loop-breaker individuals



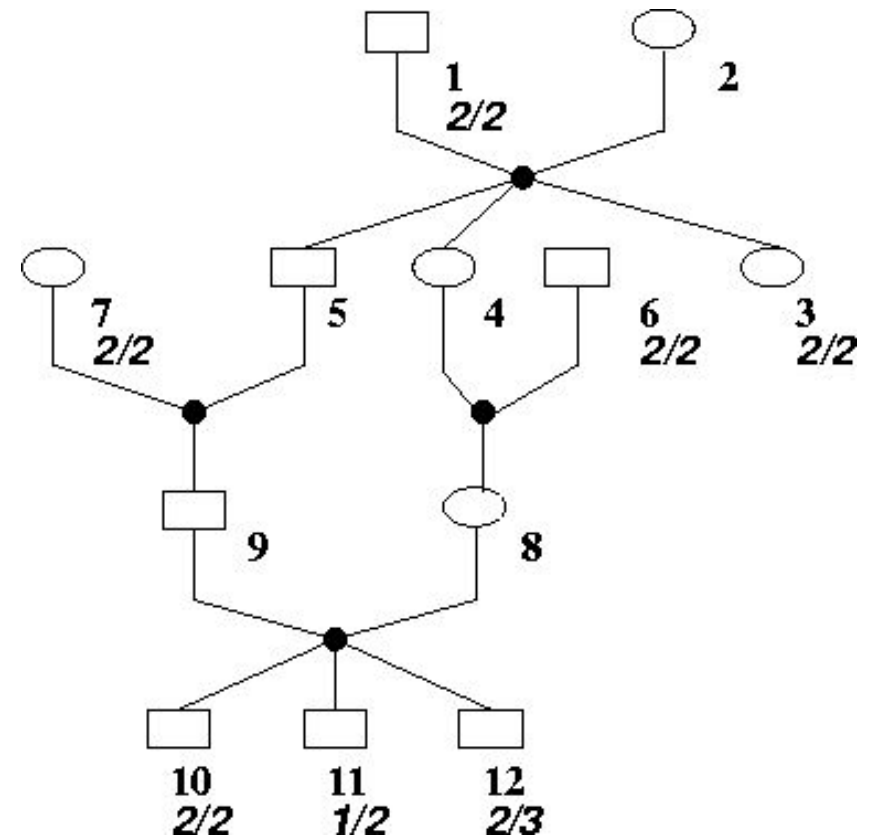
(Dechter, AIJ 1990)

(O'Connell, Weeks, Am J Hum Genet 1997)

Task 2: Error Detection

- Finds a complete assignment with the minimum number of errors

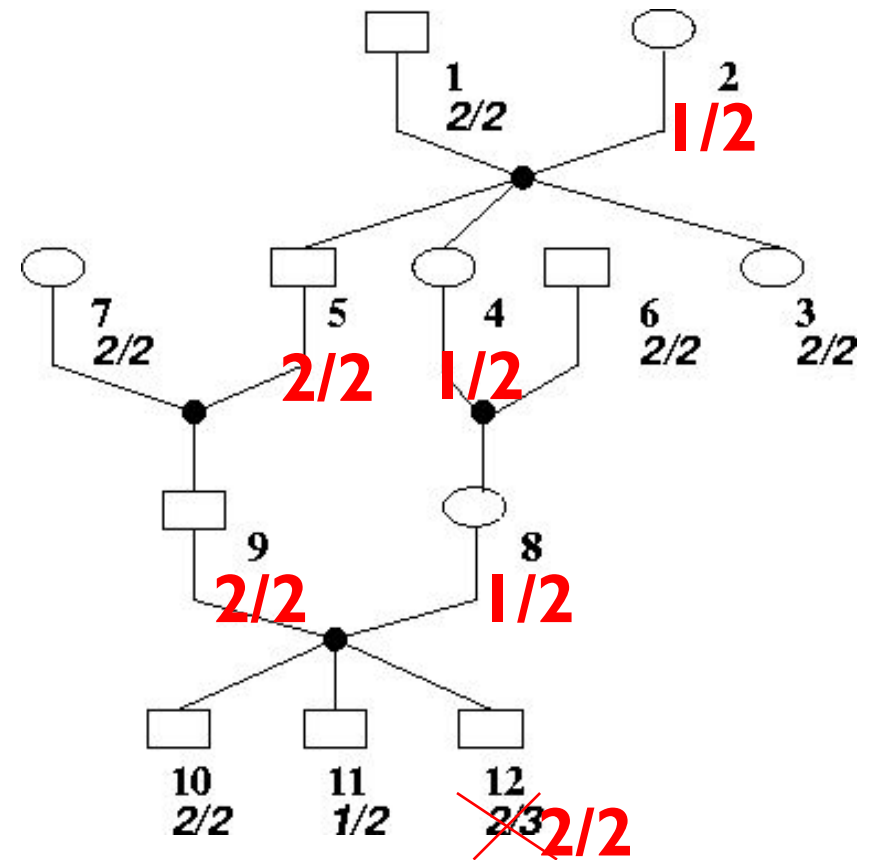
→ parsimony principle



Task 2: Error Detection

- Finds a complete assignment with the minimum number of errors

→ parsimony principle



Cost Function Network

(Shapiro, Haralick, IEEE PAMI 81)

(Freuder, Wallace, AIJ 92)

- (X, D, F)
 - $X = \{X_1, \dots, X_n\}$ n variables (Schiex, Fargier, Verfaillie, IJCAI 95)
 - $D = \{D_1, \dots, D_n\}$ n finite domains of maximum size d
 - $F = \{f_{S_1}, \dots, f_{S_e}\}$, e cost functions

f_{S_i} : associates a finite or infinite (k) positive integer to each tuple of S_i

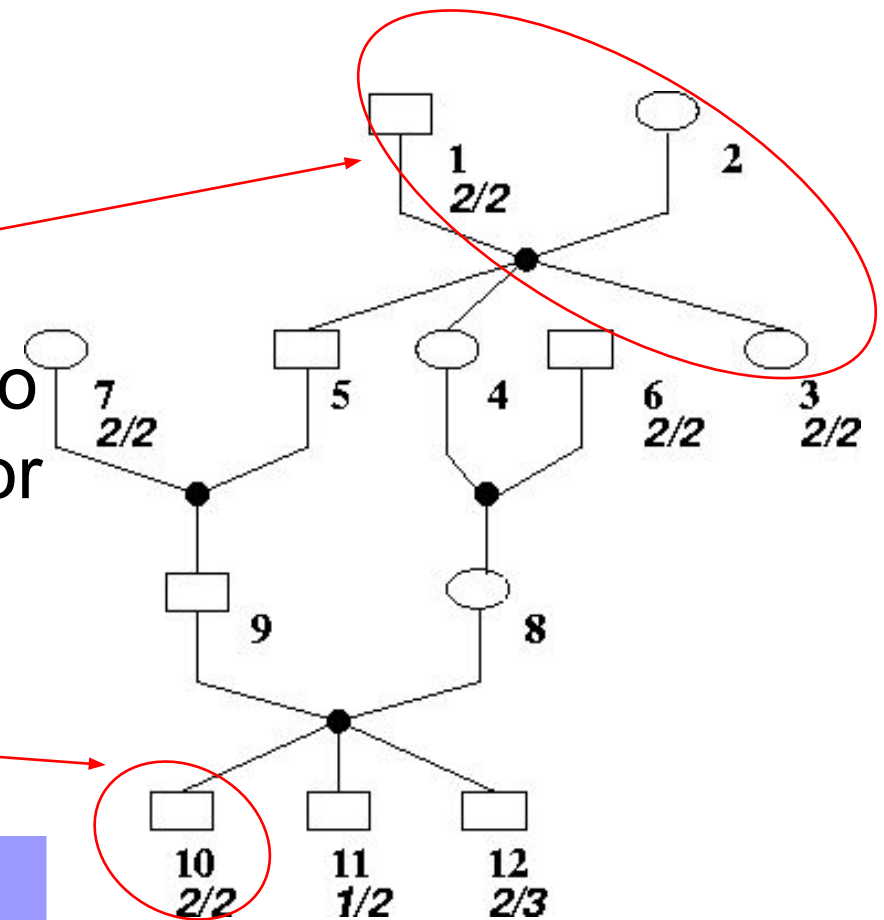
- Weighted CSP: find a complete assignment A minimizing

$$\sum_{f \in F} f_S (A[S])$$

NP-hard problem

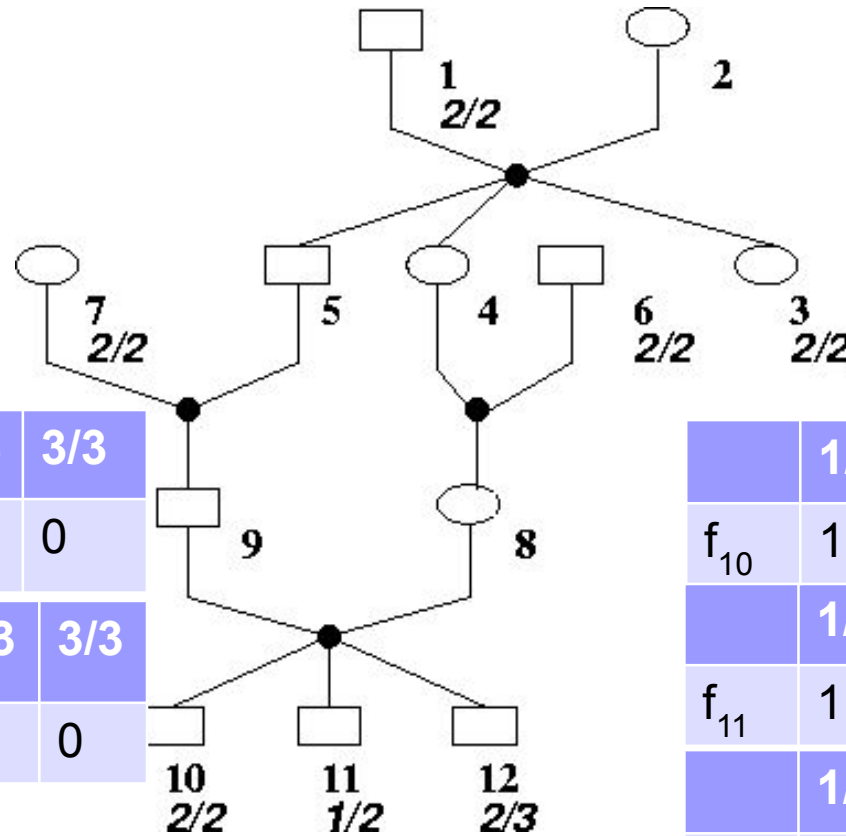
Cost Function Network

- **X**: one variable per individual
- **D**: domain of every variable is defined as the set of all possible genotypes
 $\{ 1/1, 1/2, 1/3, 2/2, 2/3, 3/3 \}$
- **F**:
 - Ternary **hard** constraints to encode Mendelian laws for any non founder
 - Unary **soft** constraints to encode genotyping data



	1/1	1/2	1/3	2/2	2/3	3/3
f_{10}	1	1	1	0	1	1

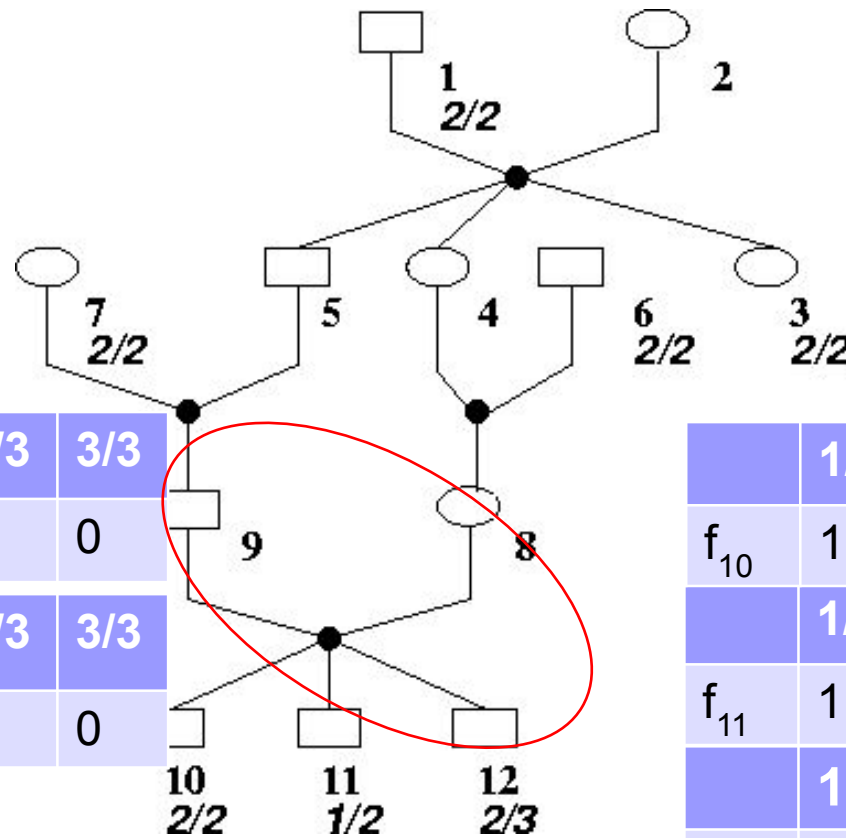
Generalized **Soft** Arc Consistency



	1/1	1/2	1/3	2/2	2/3	3/3
f_8	0	0	0	0	0	0
	1/1	1/2	1/3	2/2	2/3	3/3
f_9	0	0	0	0	0	0

	1/1	1/2	1/3	2/2	2/3	3/3
f_{10}	1	1	1	0	1	1
	1/1	1/2	1/3	2/2	2/3	3/3
f_{11}	1	0	1	1	1	1
	1/1	1/2	1/3	2/2	2/3	3/3
f_{12}	1	1	1	1	0	1

Generalized **Soft** Arc Consistency



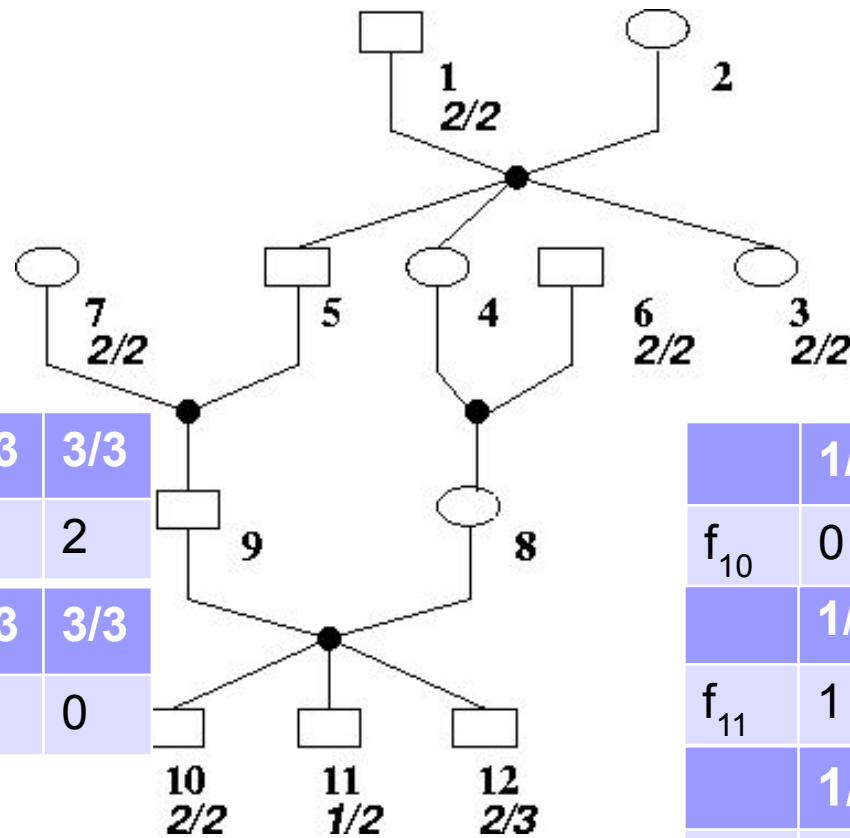
	1/1	1/2	1/3	2/2	2/3	3/3
f_8	1	0	0	0	0	0
	1/1	1/2	1/3	2/2	2/3	3/3
f_9	0	0	0	0	0	0

	1/1	1/2	1/3	2/2	2/3	3/3
f_{10}	1	1	1	0	1	1
	1/1	1/2	1/3	2/2	2/3	3/3
f_{11}	1	0	1	1	1	1
	1/1	1/2	1/3	2/2	2/3	3/3
f_{12}	0	0	0	1	0	1

Equivalence Preserving Transformation

(Schiex, CP 2000),...

Generalized **Soft** Arc Consistency



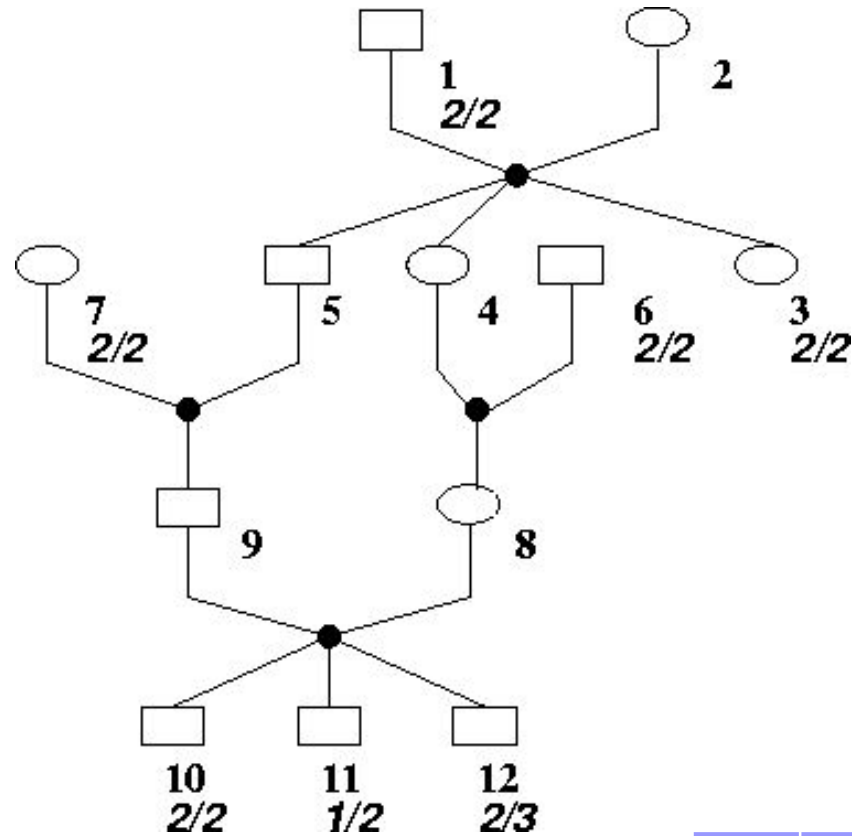
	1/1	1/2	1/3	2/2	2/3	3/3
f_8	2	0	1	0	0	2
f_9	0	0	0	0	0	0

	1/1	1/2	1/3	2/2	2/3	3/3
f_{10}	0	0	0	0	0	0
f_{11}	1	0	0	1	0	0
f_{12}	0	0	0	1	0	1

Equivalence Preserving Transformation on trios

(Schiex, CP 2000),..., (Sanchez et al, Constraint 2008)

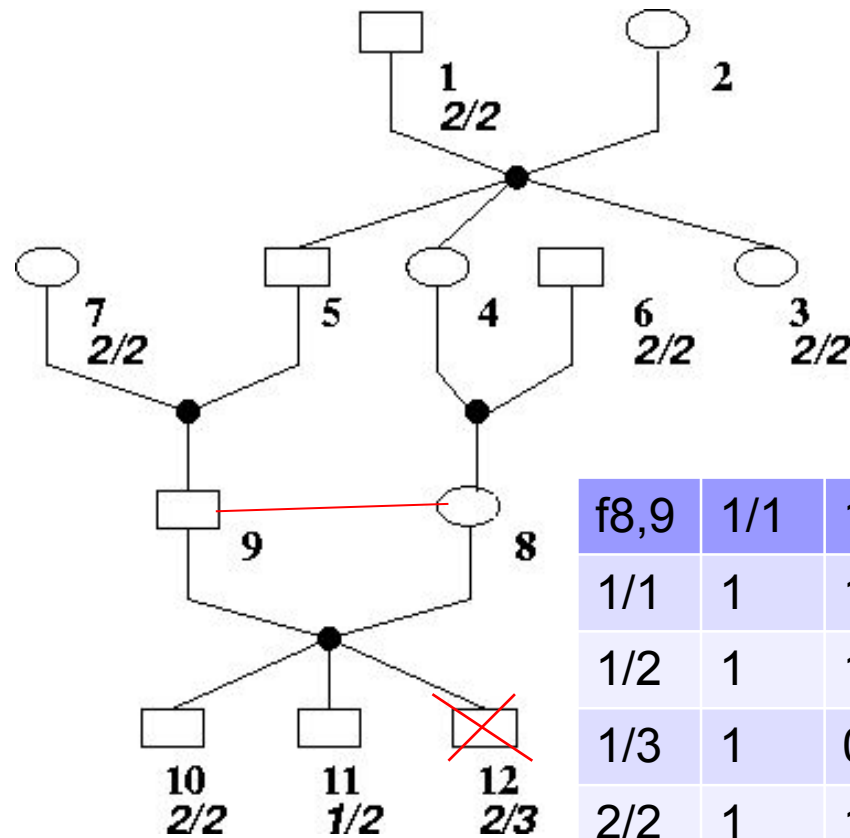
Variable elimination



	1/1	1/2	1/3	2/2	2/3	3/3
f_{12}	1	1	1	1	0	1

(Dechter, AIJ 1999)

Variable elimination



f8,9	1/1	1/2	1/3	2/2	2/3	3/3
1/1	1	1	1	1	1	1
1/2	1	1	0	1	0	0
1/3	1	0	1	0	0	1
2/2	1	1	0	1	0	0
2/3	1	0	0	0	0	0
3/3	1	0	1	0	0	1

(Dechter, AIJ 1999), (Larrosa, CP 2000)

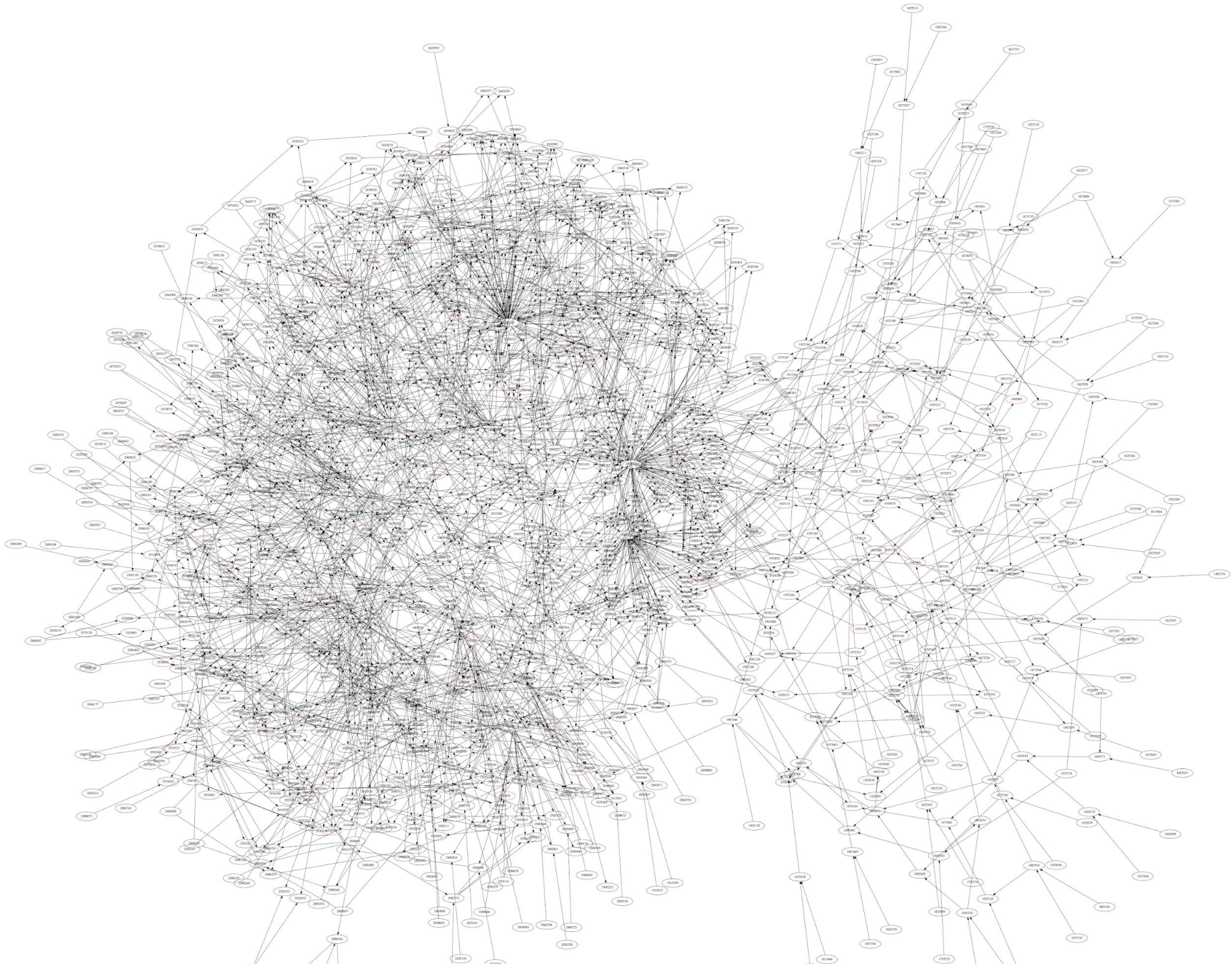
(Vitezica et al, World Congress on Genetics Applied to Livestock Production 2006)

(Sanchez et al, Constraints 2008)

Real data

*CPU time in seconds to find and prove optimality
on a linux PC 3 GHz with 16 GB using toulbar2 v0.5*

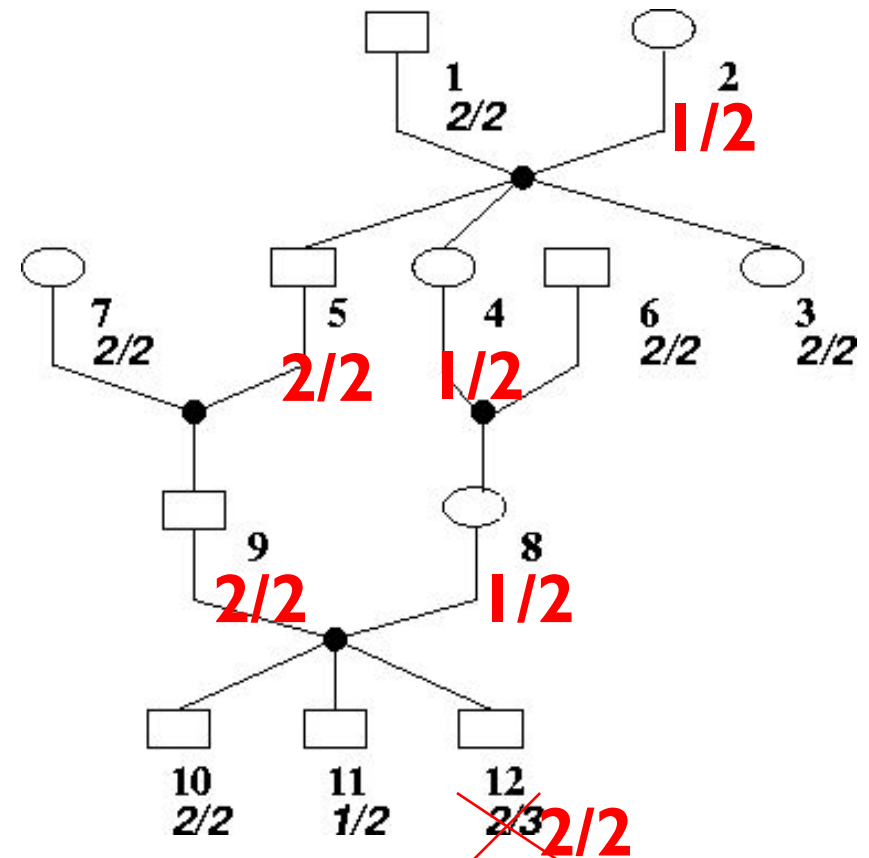
	ind	vars	genotyped	alleles	nf	ngen	treewidth ub	B&B-VE(2)		
								errors	time	nodes
<i>eye</i>	36	36	28	6	11	4	2	1	0.02	0
<i>cancer</i>	49	48	37	8	18	5	2	1	0.21	0
<i>parkinson</i>	37	34	13	4	7	7	5	0	0	6
<i>berrichon_{1nc}</i>	129516	9947	2448	4	8821	17	262	2	4.73	8805
<i>berrichon₁</i>	129516	10017	2483	4	8786	17	330	23	5.81	8384
<i>berrichon_{2nc}</i>	27255	19337	10215	4	4719	19	-	41	5.89	6170
<i>berrichon₂</i>	27255	19562	10215	4	2381	19	-	106	17.23	15445
<i>langlade₁</i>	1355	1209	711	9	298	13	84	38	12.28	391
<i>langlade₂</i>	1355	1223	715	7	298	13	82	89	60.56	17857
<i>langlade₃</i>	1355	1258	787	5	298	13	85	39	14.19	6731
<i>langlade₄</i>	1355	1186	672	8	298	13	83	43	59.7	3520
<i>moissac₁</i>	283	260	183	2	81	5	6	0	0	5
<i>moissac₂</i>	283	244	167	7	81	5	6	0	0.51	6
<i>moissac₃</i>	283	225	151	3	81	5	6	0	0	4
<i>moissac₄</i>	283	256	179	2	81	5	6	0	0	5
<i>moissac₅</i>	283	237	161	8	81	5	6	0	1.02	5
<i>moissac₆</i>	283	201	131	11	81	5	5	0	5.64	6



1515 sheep (LangladeM7), 243 founders, 3 alleles, 880 genotypings, 13 generations
(minimum of 23 errors, most probable correction removes 43 genotypes)

Task 3: Error Correction using Probabilistic Model

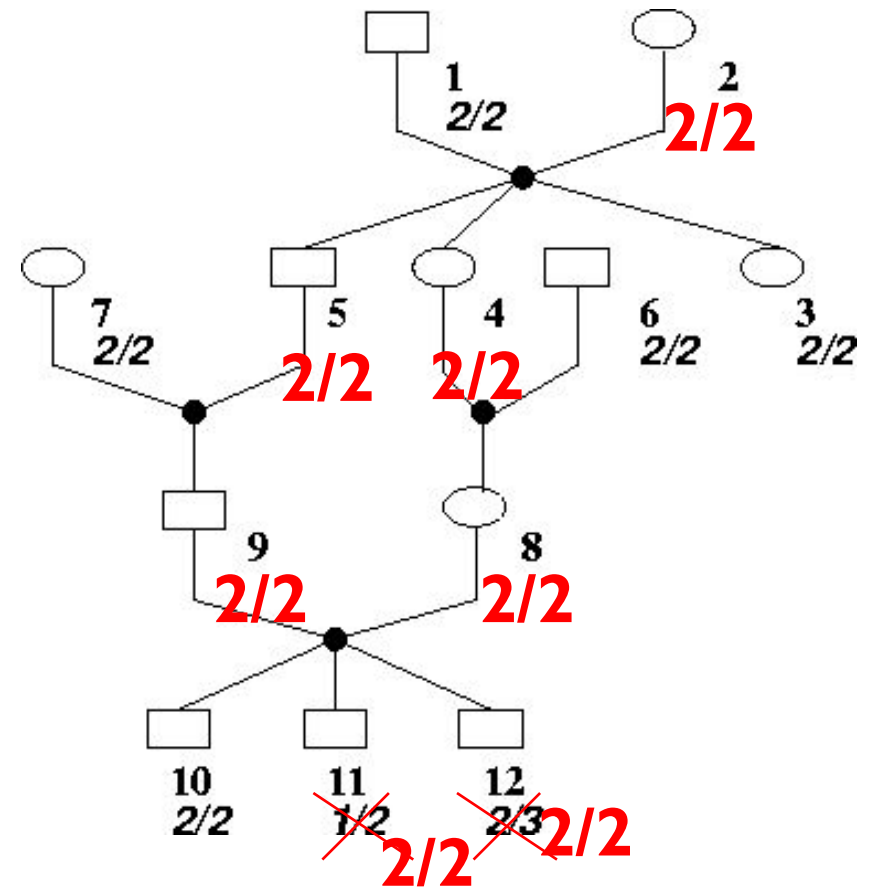
- Finds a complete assignment with maximum posterior probability
- Bayesian network



Prior on genotyping error: 1%
(and equifrequent alleles)

Task 3: Error Correction using Probabilistic Model

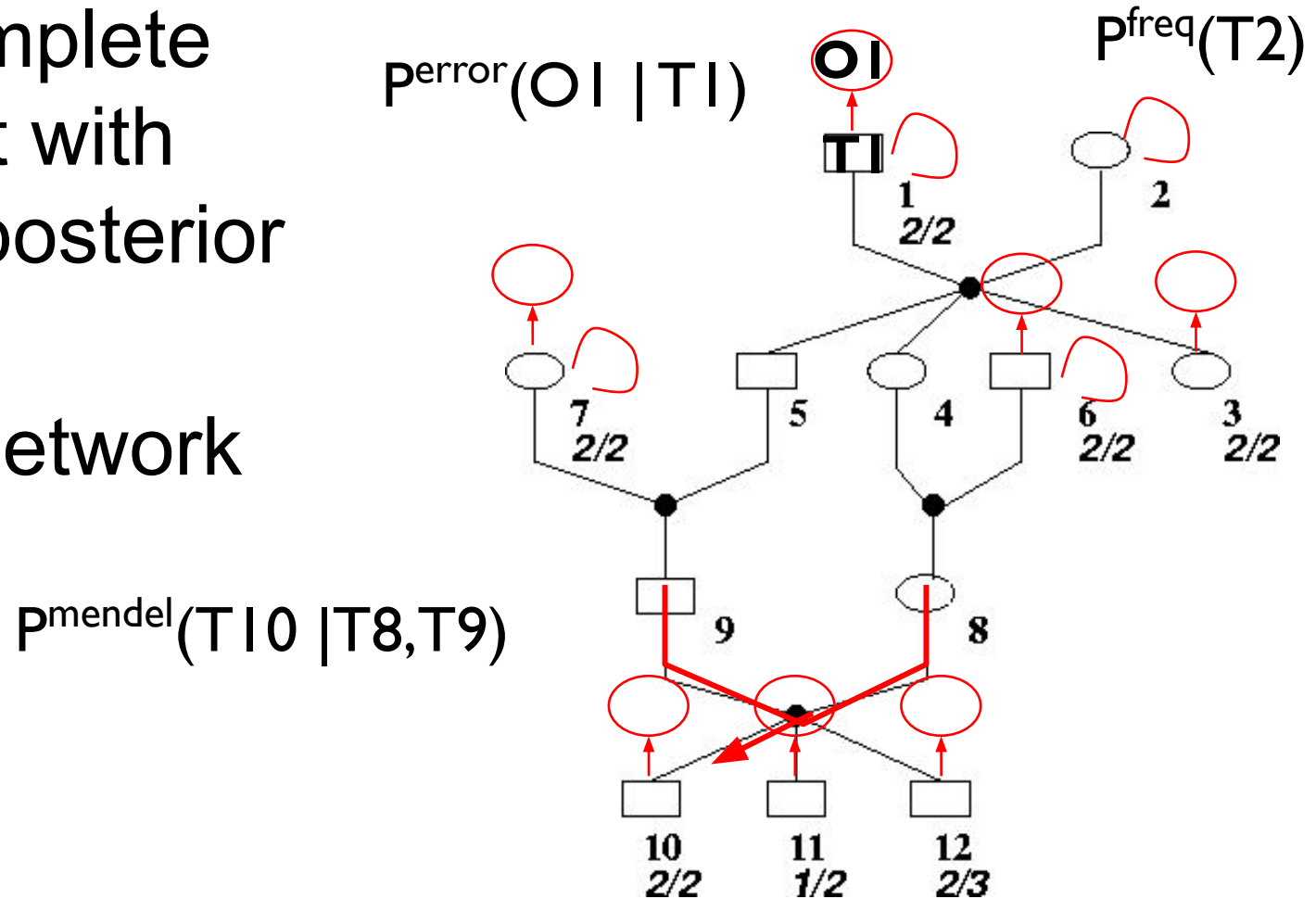
- Finds a complete assignment with maximum posterior probability
- Bayesian network



Prior on genotyping error: 10%
(and equiprequent alleles)

Task 3: Error Correction using Probabilistic Model

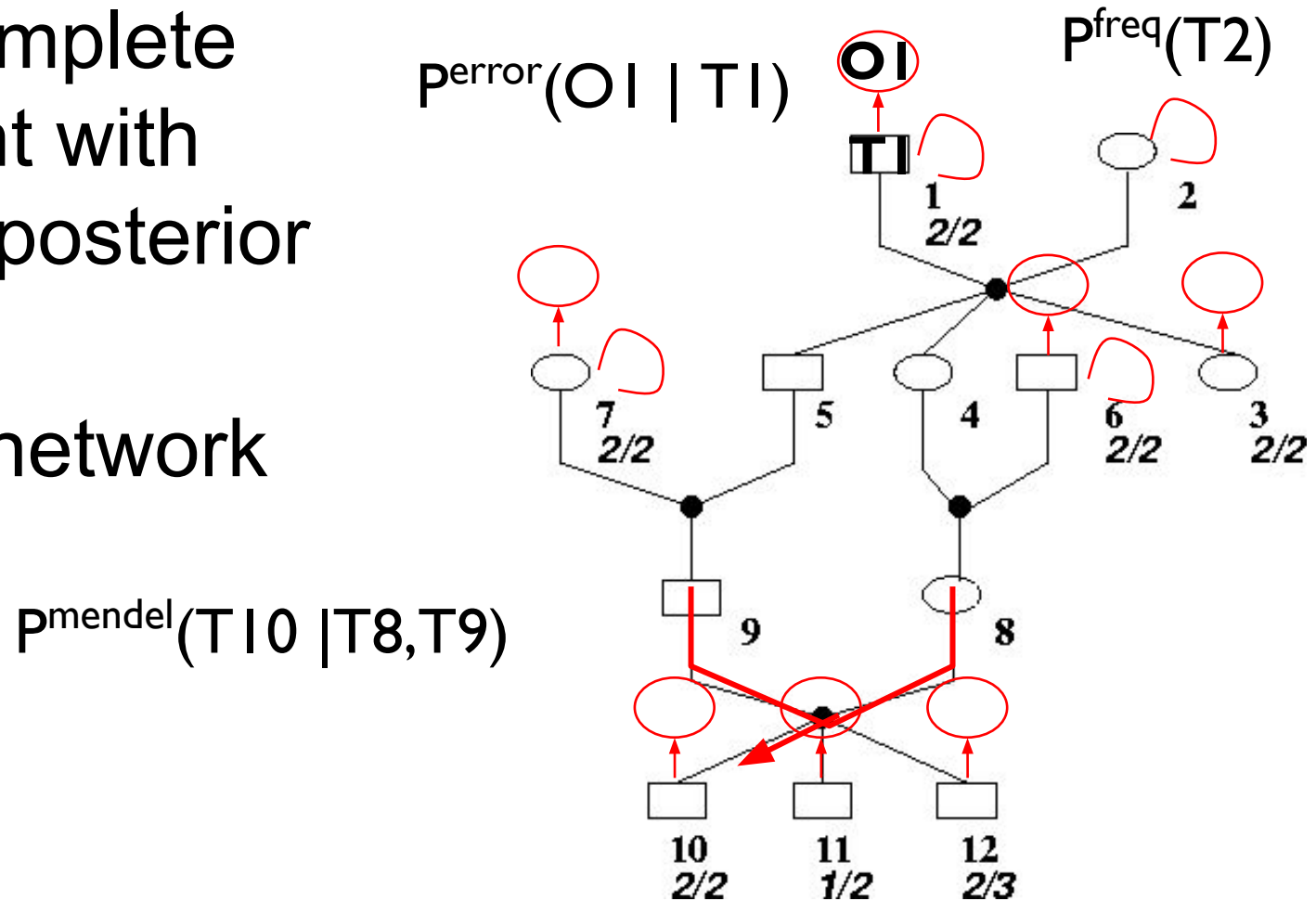
- Finds a complete assignment with maximum posterior probability
- Bayesian network



$$P(O, T) = \prod P^{\text{error}}(O_i | T_i) \times \prod P^{\text{mendel}}(T_i | \text{parents}(i)) \times \prod P^{\text{freq}}(T_i)$$

Task 3: Error Correction using Probabilistic Model

- Finds a complete assignment with maximum posterior probability
- Bayesian network

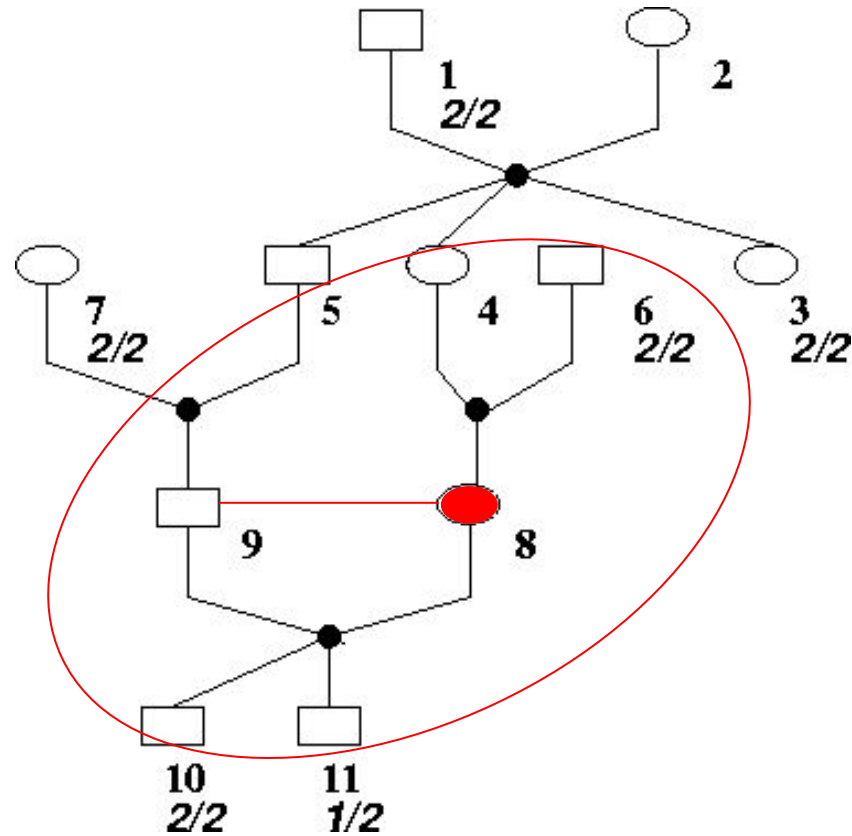


~~$$P(O, T) = \prod P^{\text{error}}(O_i | T_i) \times \prod P^{\text{mendel}}(T_i | \text{parents}(i)) \times \prod P^{\text{freq}}(T_i)$$~~

-log transform →

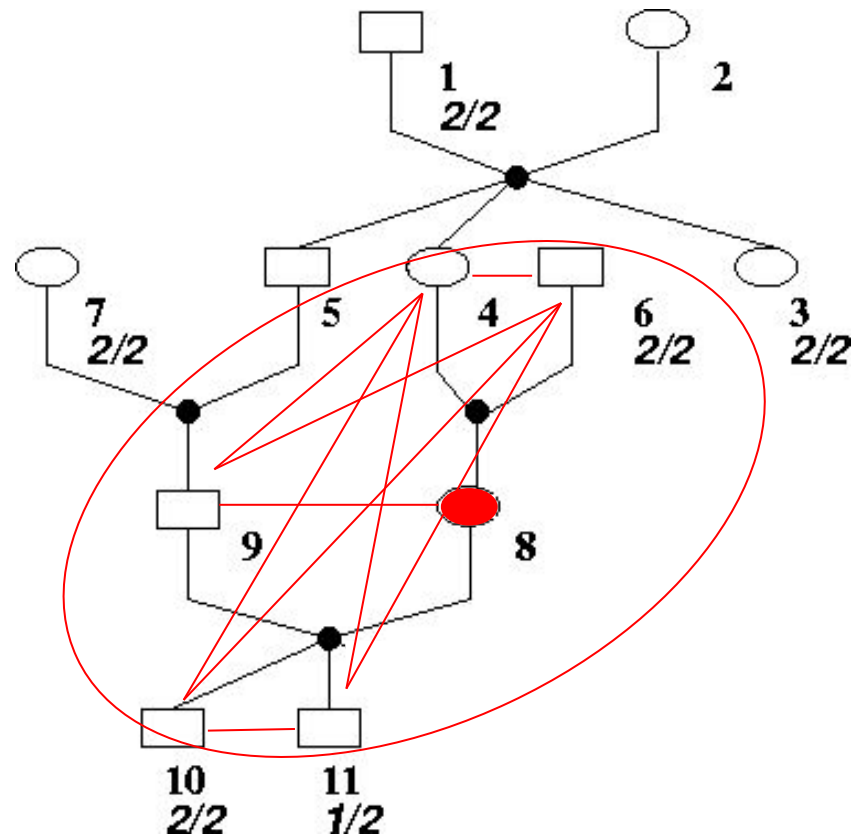
$$\sum_{f_S \in F} f_S(A[S])$$

Variable elimination ..continued



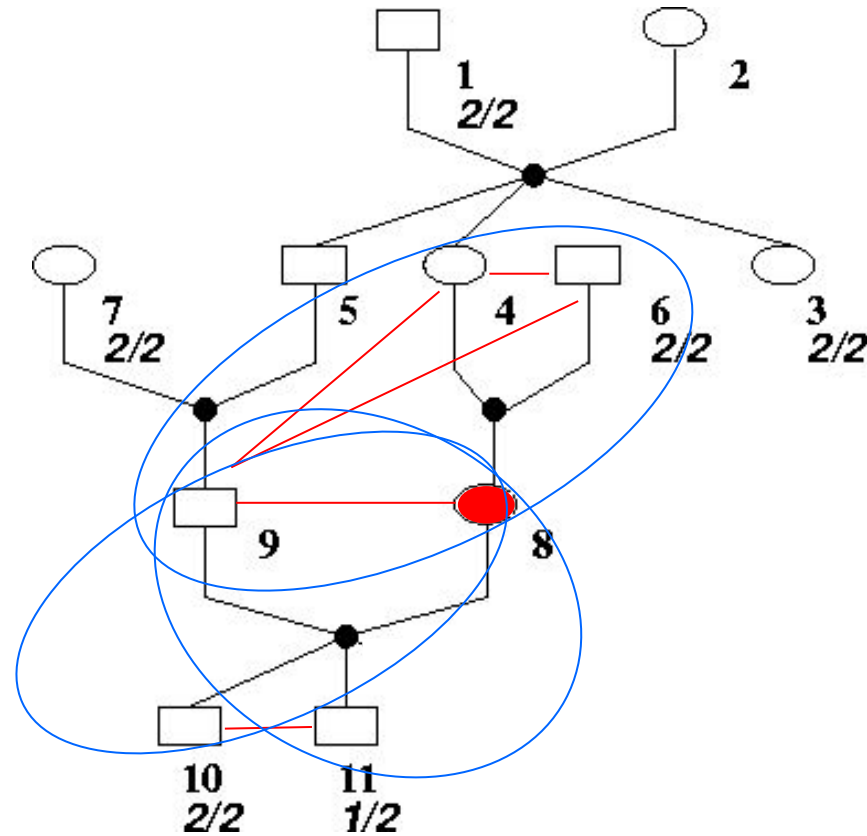
(Dechter, AIJ 1999)

Variable elimination ..continued



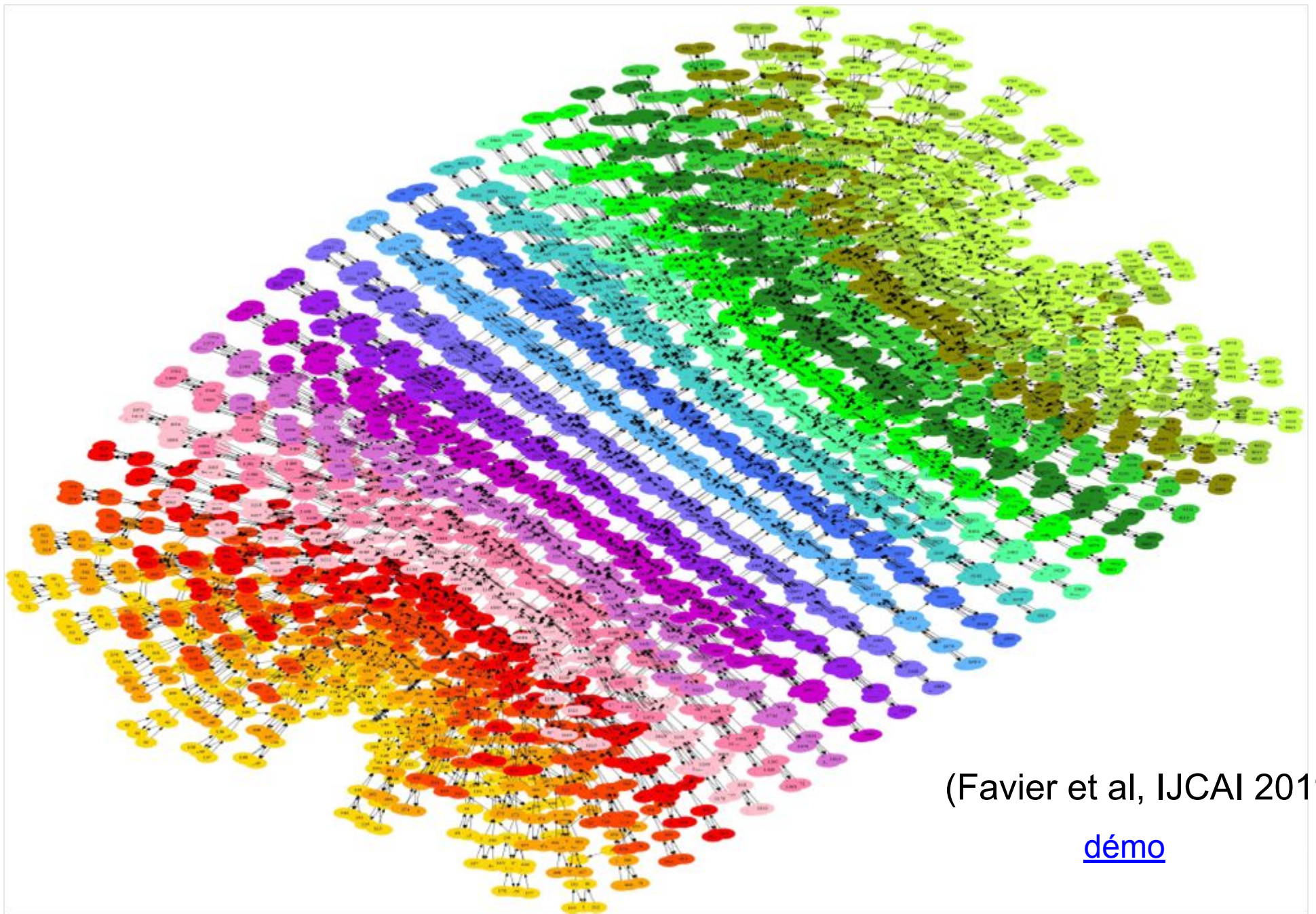
(Dechter, AIJ 1999)

Variable elimination ..continued



Cost function decomposition

(Favier et al, IJCAI 2011)



(Favier et al, IJCAI 2011)

[d emo](#)

Optimal haplotype reconstruction in half-sib families (Legarra et al, WCGALP 2010)

Guaranteed Discrete Energy Optimization on Large Protein Design Problems

Guaranteed Discrete Energy Optimization on Large Protein Design Problems. *Journal of chemical theory and computation*.
Fast search algorithms for computational protein design. *Journal of computational chemistry*.

Thomas Schiex (INRA)
D. Simoncini, D. Allouche, S. Barbe (LISBP)



* Slides by T. Schiex

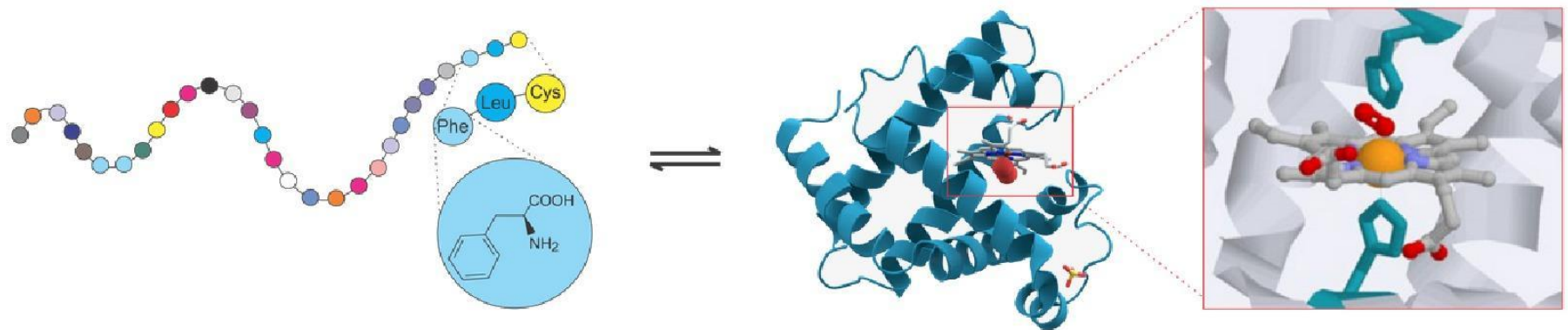
Summer 2016 - US tour

May 2022 – LIPME INRAE

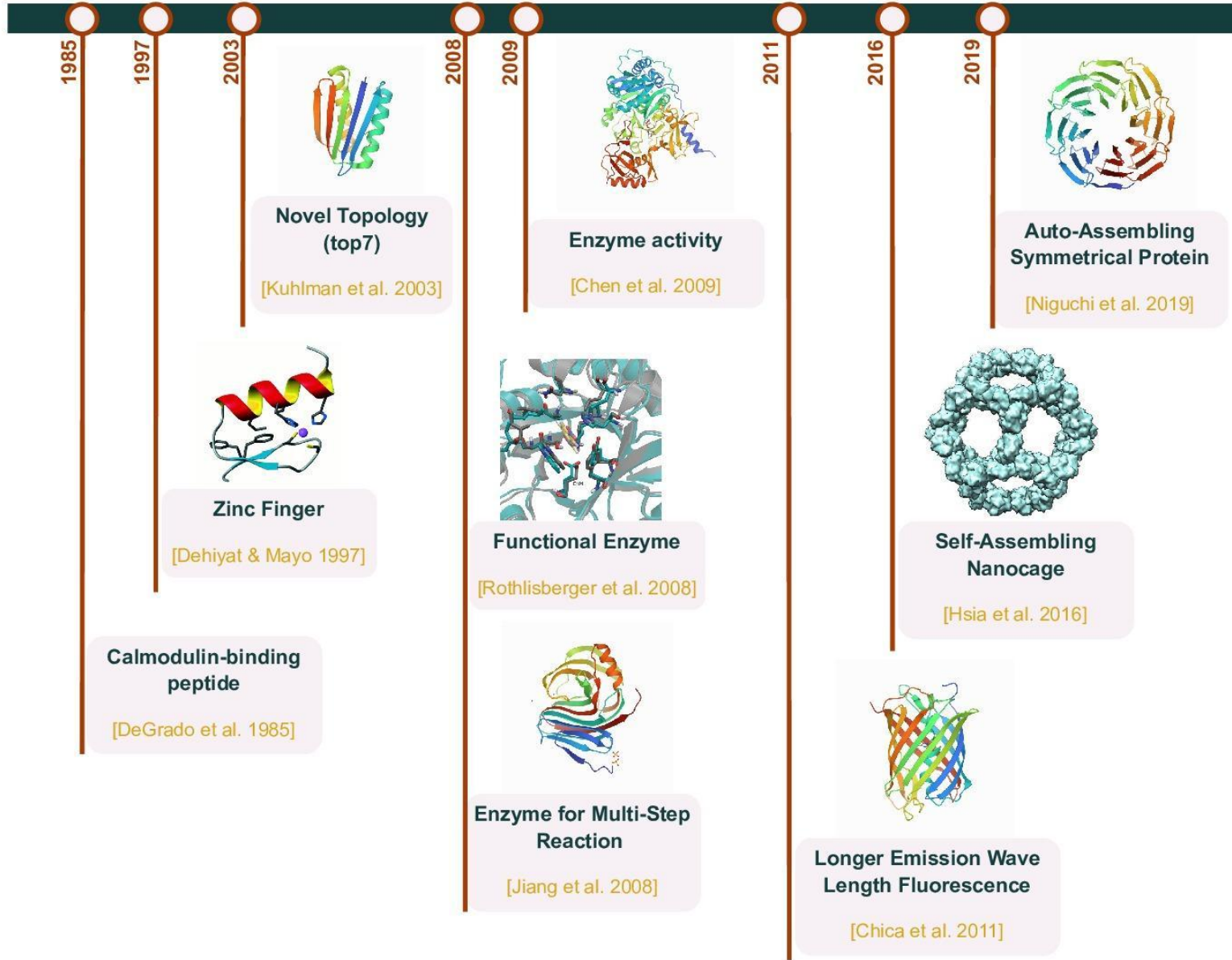
Why should we want to design proteins?

Eco-friendly chemical/structural nano-agents

- New drugs for health (human, animals, plants)
- New catalysts (environment, recycling, biofuels, food and feed, cosmetics...),
- New components for nanotechnologies
- Relying on inexpensive atomic level 3D-printers (bacterias, yeast, ...)



⁰Thanks to the Zhang lab. for this image.



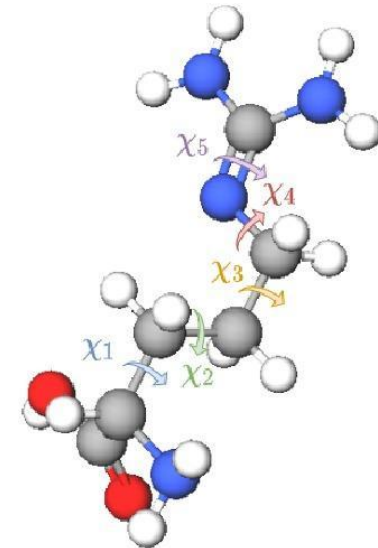
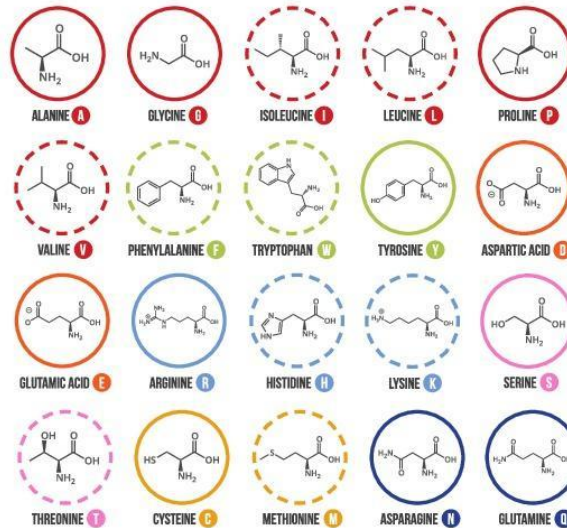
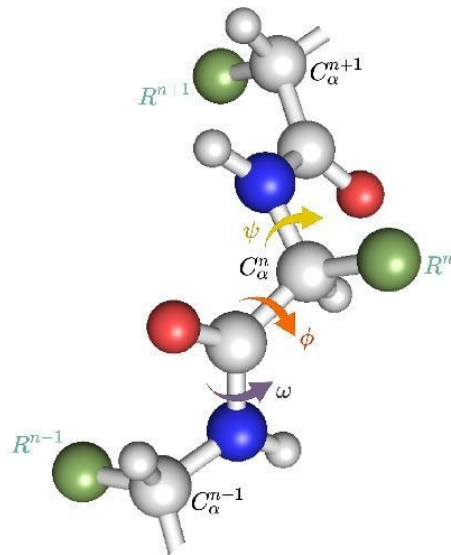
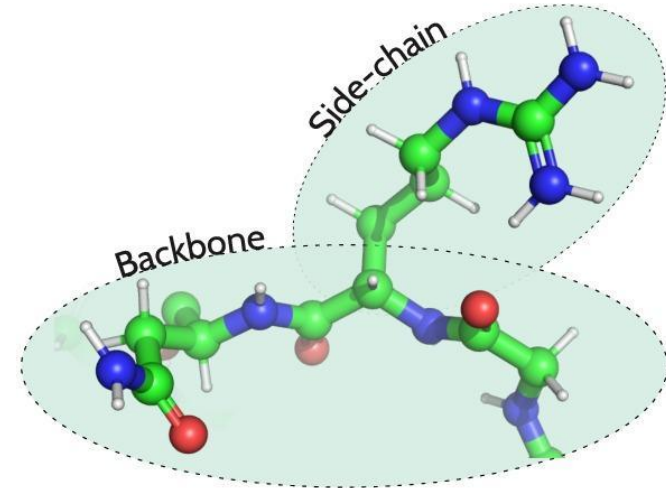
Informal definition (globular proteins)

Produce a sequence s of amino-acids that *spontaneously adopts* a conformation X that *performs some function*.

What defines a conformation ?

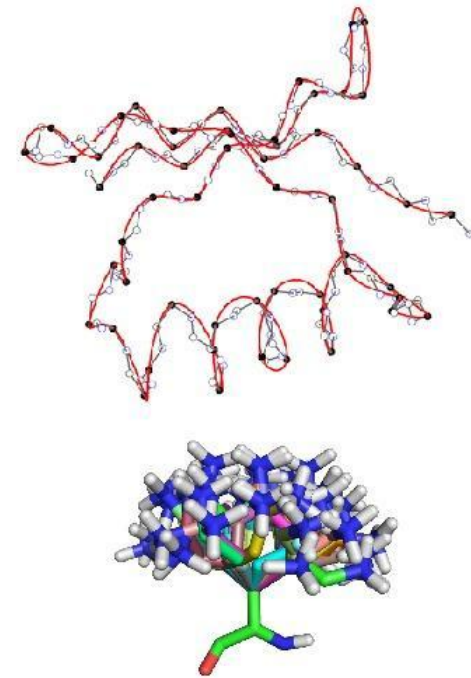
Conformation

- backbone: dihedral angles ϕ_i, ψ_i
- sequence: amino-acid choice S_i
- side-chains: torsion angles χ_{ij}



Rigid backbone design

- 1 Side-chain flexibility: discretized in rotamers
- 2 1 rotamer to choose at each flexible/mutable position



Search Space

- 1 Fully discrete description, defined by a choice of rotamer (AA \times conformation) for each position.
- 2 Search space can be $\approx 250^n$ or more (Dunbrack)

Using Talaris14²⁰ (pairwise decomposable)

- Bonded (dihedral torsion angles,...)
- van der Waals (attractive+repulsive)
- Electrostatic (statistical)
- Rotamer statistics (Dunbrack)
- Hydrogen bonds
- Implicit solvation
- Cutoff functions

$$E(c) = E_{\emptyset} + \sum_{i=1}^n E_i(r) + \sum_{i < j} E_{ij}(r, s)$$

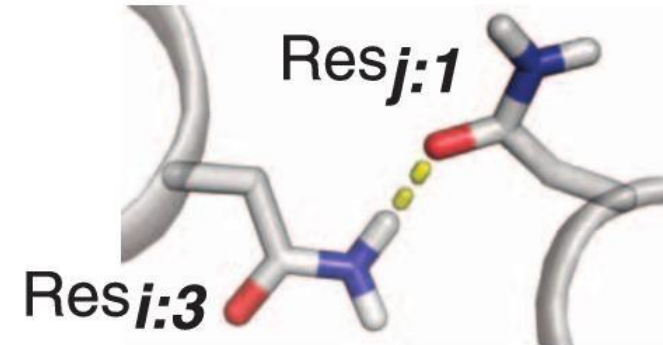
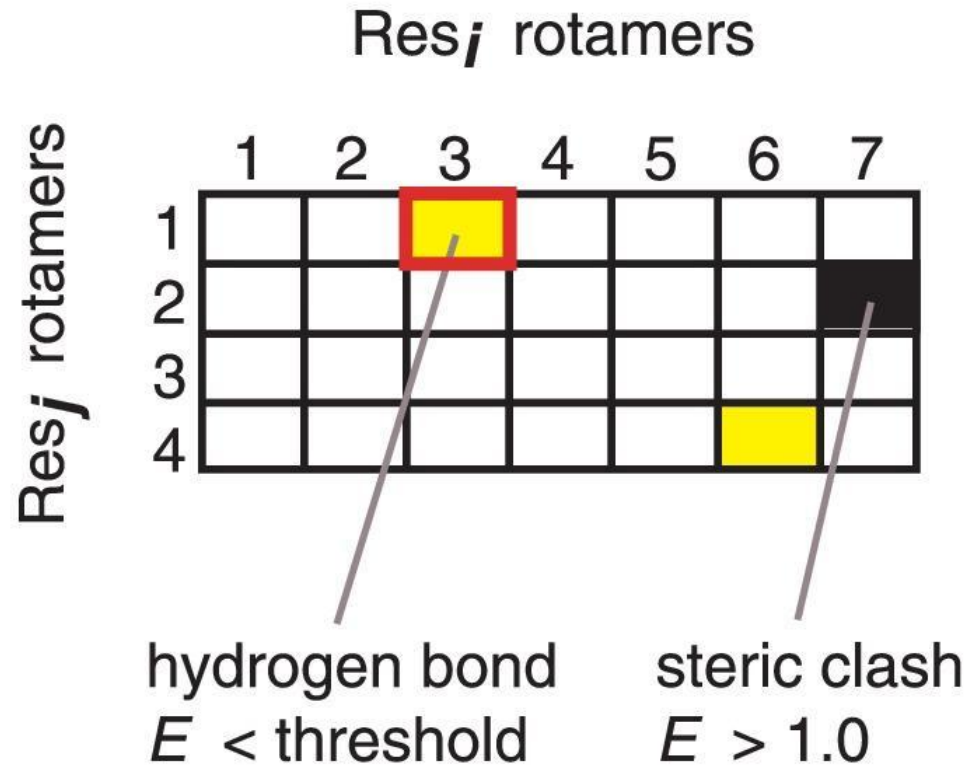
Using Talaris14²⁰ (pairwise decomposable)

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- Hydrogen bonds
- Implicit solvation
- Cutoff functions

$$E(c) = E_{\emptyset} + \sum_{i=1}^n E_i(r) + \sum_{i < j} E_{ij}(r, s)$$


$$\sum_{f \in F} f_S (A[S])$$

NP-hard

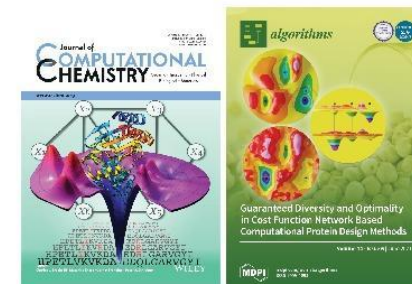


Large input (> 1GB)

NP-hard problem

Toulbar2 is able to...

- provide a proven minimum energy solution²⁵
- exhaustively enumerate sequences close to it
- provide sequence libraries with guaranteed diversity.¹⁸
- in sequence-conformation spaces of size $> 10^{400}$



Rosetta's Monte Carlo Simulated Annealer increasingly fails to find the optimal sequence^a

^aDavid Simoncini et al. "Guaranteed Discrete Energy Optimization on Large Protein Design Problems". In: *Journal of Chemical Theory and Computation* 11.12 (2015), pp. 5980–5989. DOI: 10.1021/acs.jctc.5b00594.

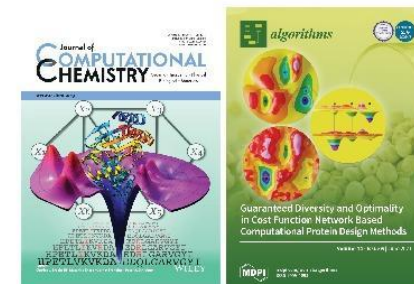
Clément Viricel's PhD on predicting changes in binding affinity (counting) (Viricel et al, Bioinformatics 2018)

Large input (> 1GB)

NP-hard problem

Toulbar2 is able to...

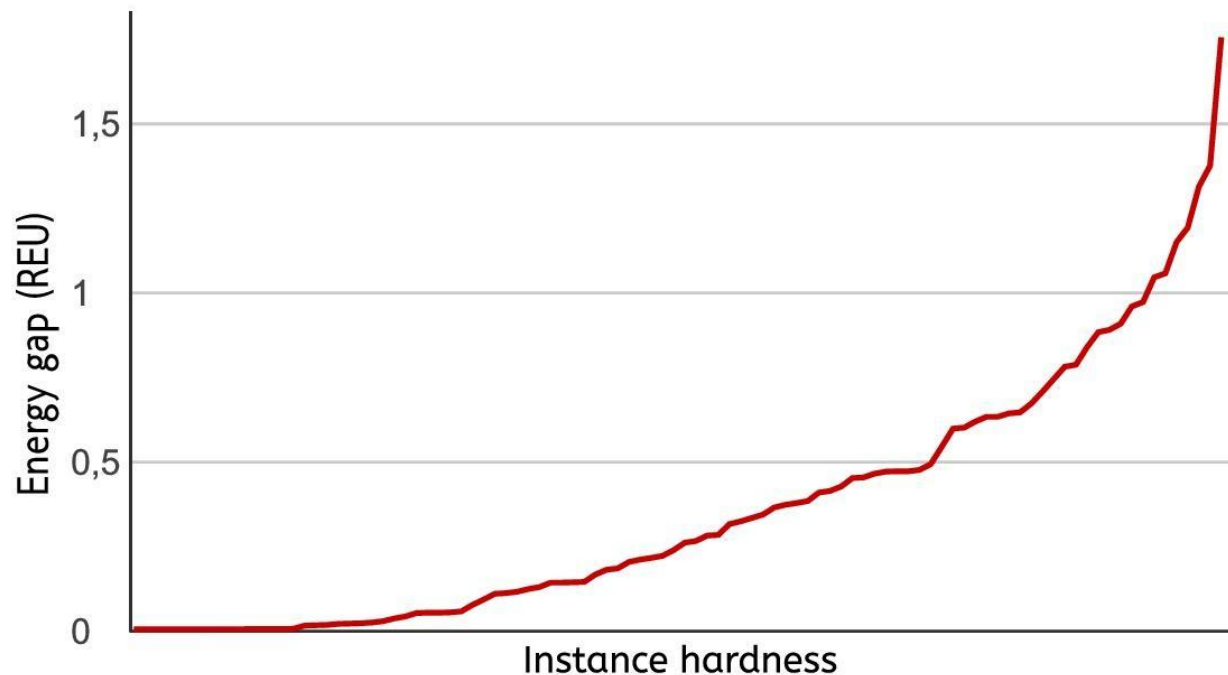
- provide a proven minimum energy solution²⁵
- exhaustively enumerate sequences close to it
- provide sequence libraries with guaranteed diversity.¹⁸
- in sequence-conformation spaces of size $> 10^{400}$



Manon Ruffini's PhD on diversity encoding and negative design (Ruffini et al, ICTAI 2019, Algorithms 2021)

Rosetta's Monte Carlo Simulated Annealer increasingly fails to find the optimal sequence^a

^aDavid Simoncini et al. "Guaranteed Discrete Energy Optimization on Large Protein Design Problems". In: *Journal of Chemical Theory and Computation* 11.12 (2015), pp. 5980–5989. DOI: 10.1021/acs.jctc.5b00594.



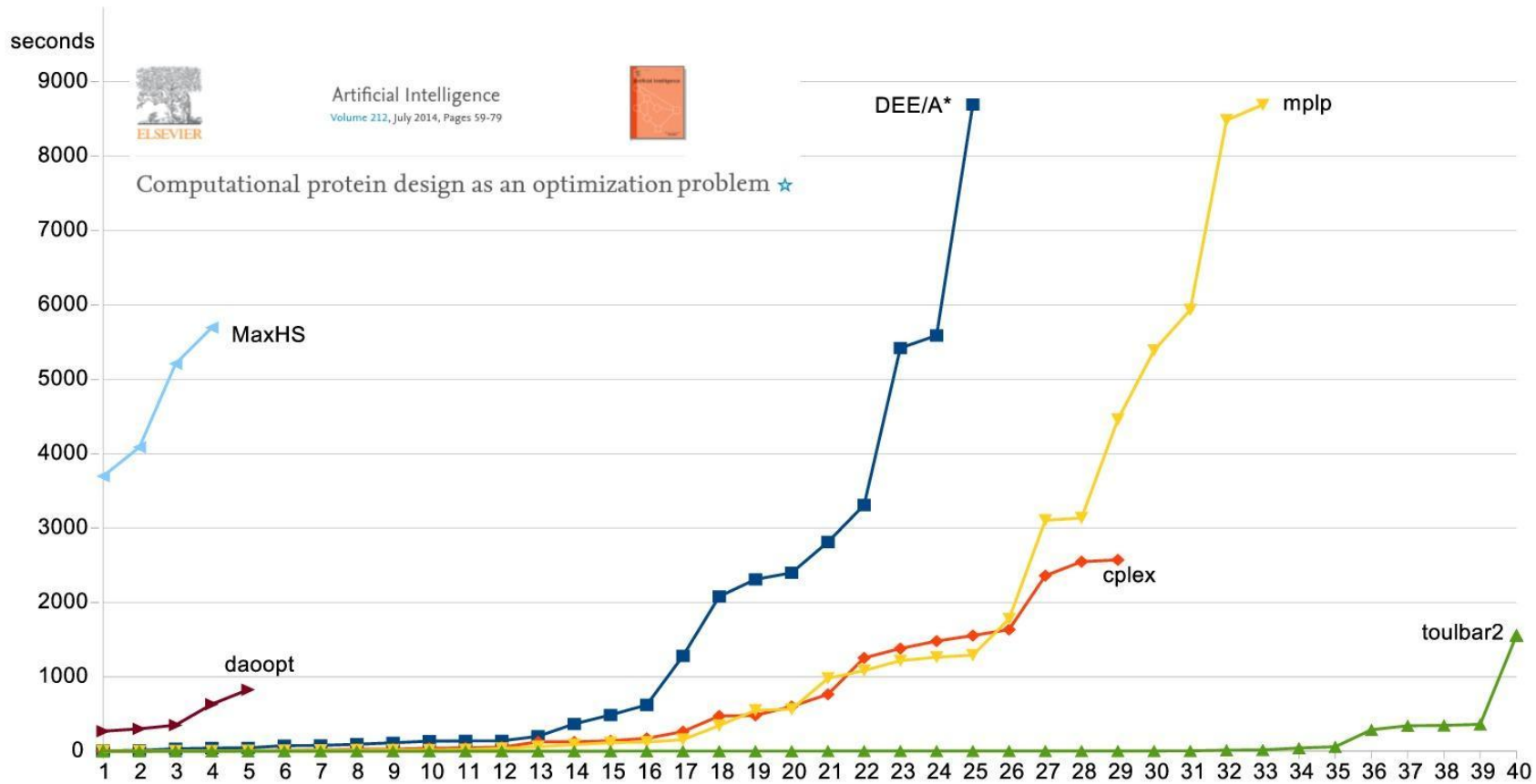
Asymptote: Size matters!

Asymptotic convergence can be arbitrarily slow...

Guaranteed Discrete Energy Optimization on Large Protein Design Problems

David Simoncini[†], David Allouche[†], Simon de Givry[†], Céline Delmas[†], Sophie Barbe^{‡§±}, and Thomas Schiex^{**}

Toulbar2 vs. CPLEX, MaxHS...(real instances)

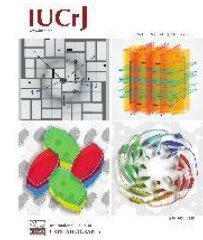


of instances solved (X) within a per instance cpu-time limit (Y)

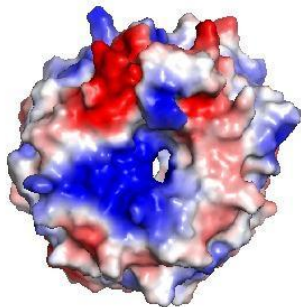
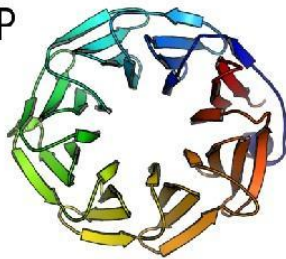
Designing a self-assembling β -propeller

Coll. A. Voet (KU Leuven), D. Simoncini¹⁶

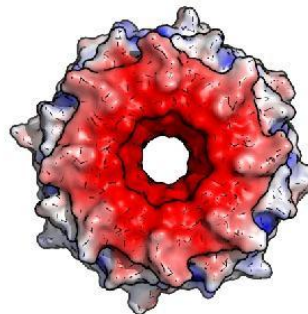
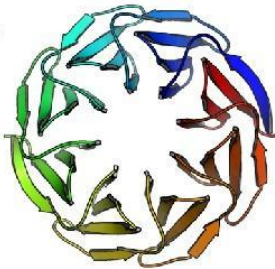
-  Tako: (R)evolution + Rosetta/talaris14 8 fold
-  Ika: toulbar2 + talaris14 4 fold



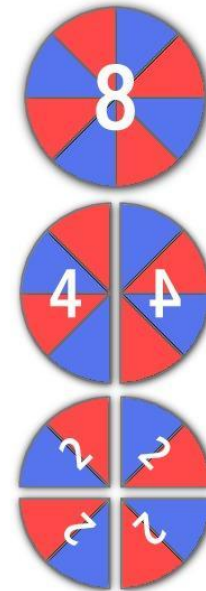
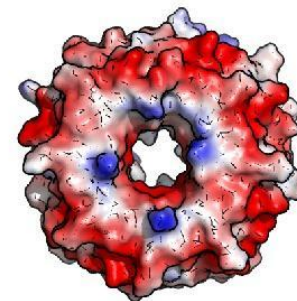
20VP



Tako



Ika





A Mixed Integer Programming Reformulation of the Mixed Fruit-Vegetable Crop Allocation Problem

Sara Maqrot¹, Simon De Givry¹, Gauthier Quesnel¹
Marc Tchamitchian² *

¹UR 875, MIAT-INRA Toulouse France

²UR 767, Ecodeveloppement, INRA-Avignon France

June 28th 2017

IEA/AIE 2017, Arras, France

* Jury member of Mahuna Akplogan's PhD on crop allocation problem

Introduction



Fruit cropping

+



Vegetable cropping

=



Mixed fruit-vegetable cropping
Avignon France, 2015

Agriculture's challenges (examples)

- Local market, diversified food.
- Reducing pollution (chemical products).
- Preservation of resources (water, energy).
- Better biodiversity.



Localization in France

Casdar SMART 2014-2016
National Project in France

Introduction

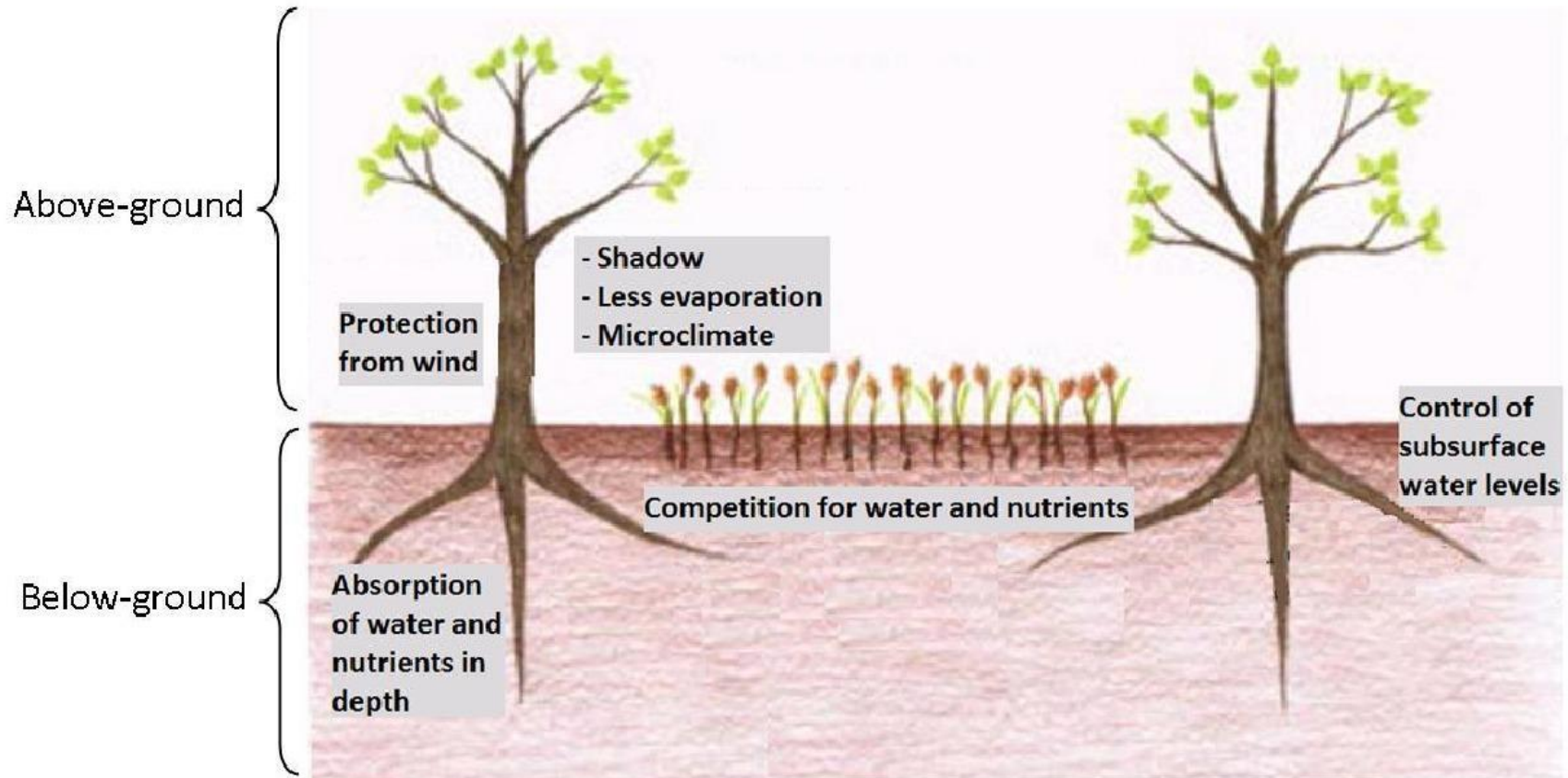
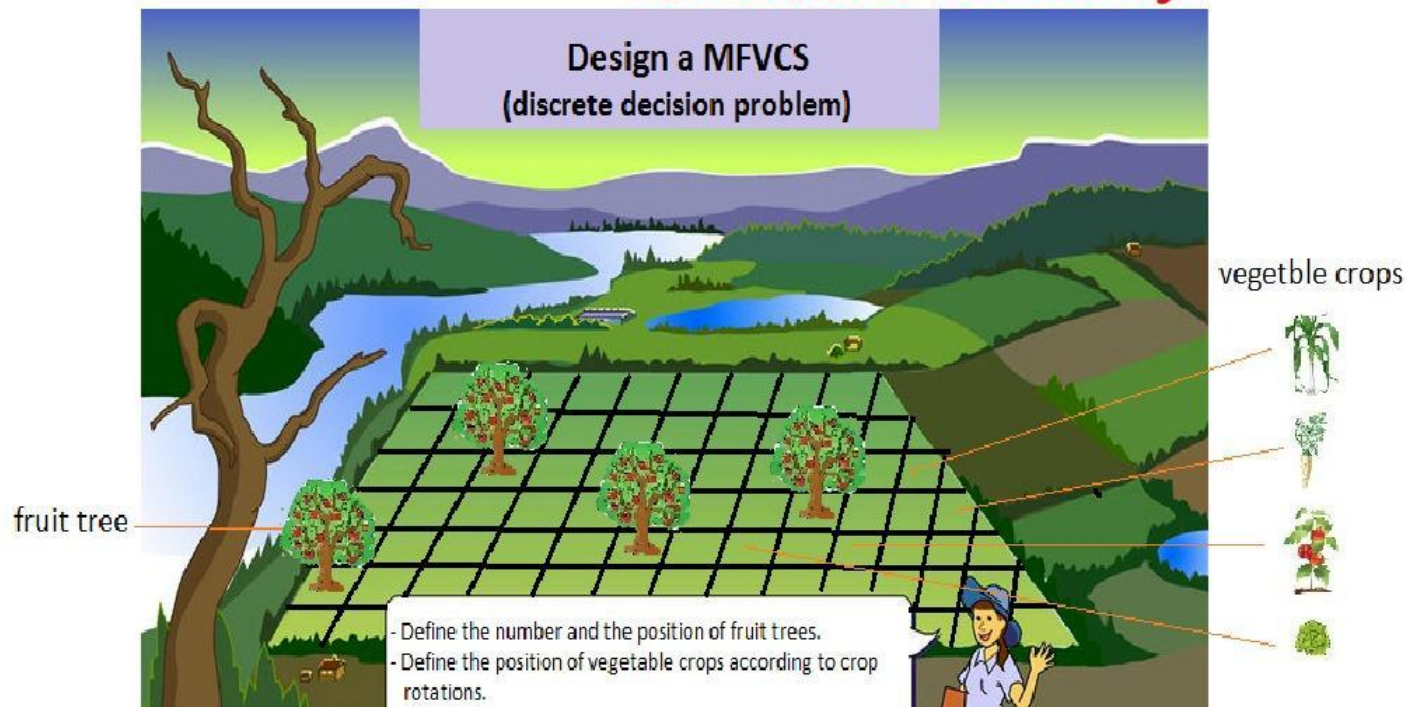


Figure – Example of above- and below-ground interactions between crops and trees [Kaeser et al., 2010]

Introduction

The aim of our study



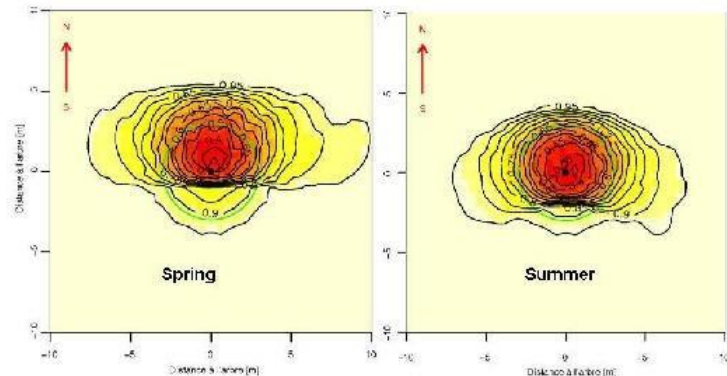
Interactions



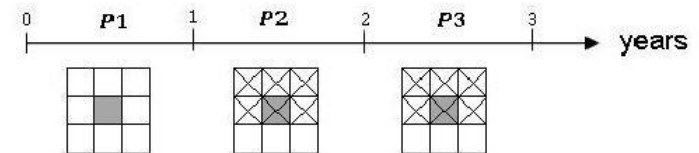
Designing a mixed fruit-vegetable system having diversified production while **optimizing the use of resources** (light, water and nutrients), by exploiting interactions between trees and crops.

Above-ground interactions : shade and micro-climate conditions

Fruit tree



Solar radiation interception simulation of an apple tree
(Source : PSH - INRA Avignon).



Evolution of potential shade in spring and summer (checked cells).

Interactions with vegetable crops

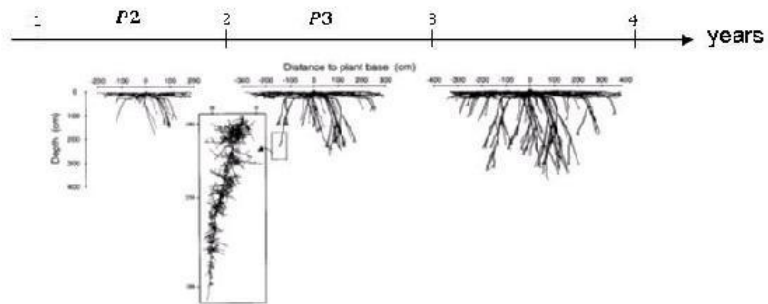
Crop sensitivity to shade.

- - : negative effect
- 0 : neutral effect
- + : positive effect

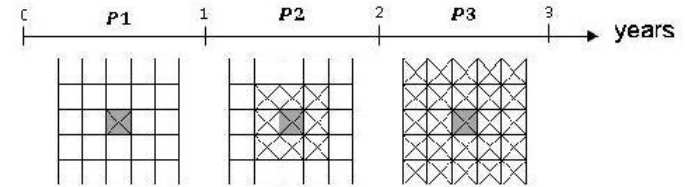
Crops	Shade seasons	
	Spring	Summer
Lettuce	0	+30
Tomato	0	+20
Onion	-10	0
Melon		-10
Carrot		0

Below-ground interactions : root extension dynamics

Fruit tree



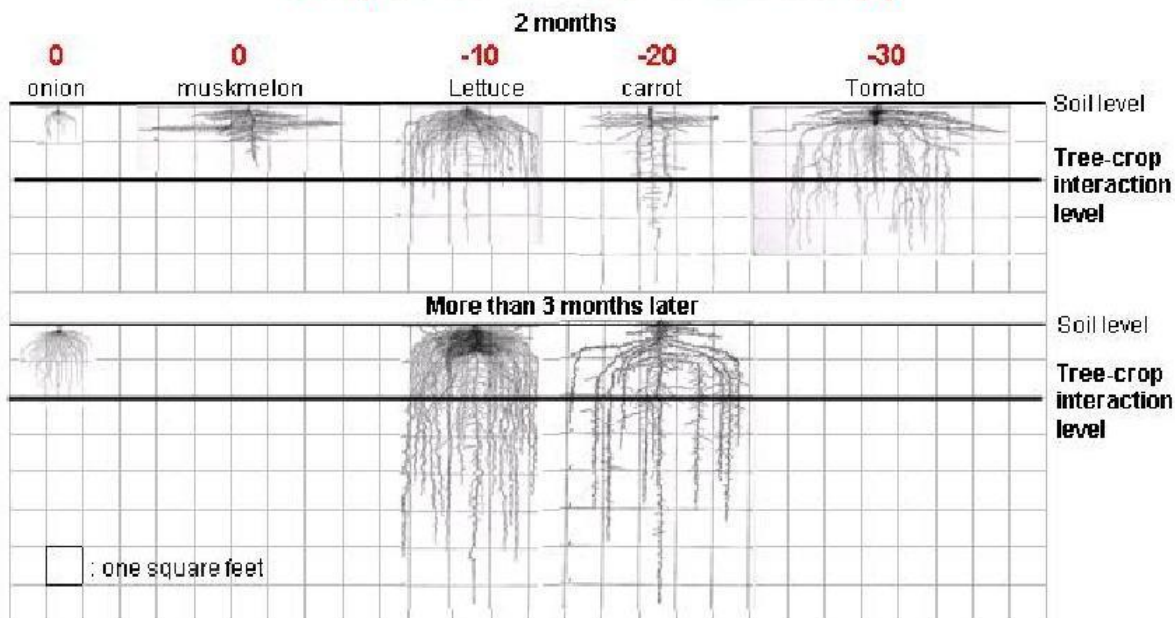
Simulated plum root system [Vercambre et al., 2003]



Surface view of root extensions of an apple tree.

Interactions with vegetable crops

Competition for water in summer (-)



[Weaver and Bruner, 1927]

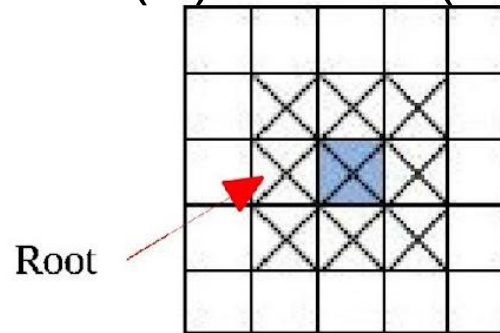
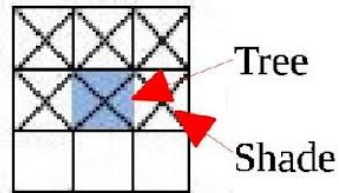
Sharing for water (+) Absorption of surplus water

winter	spring	summer	autumn
+10	+10		+10



Question

Plants N trees to get the right number of cells with shade(S) or root (R)

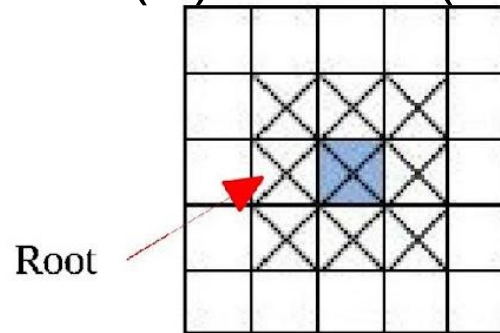
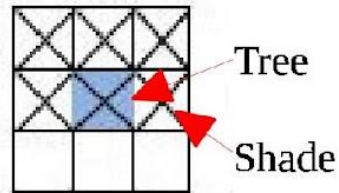


$$N=2, SR=5, -R=5, --=13$$

$$N=2, SR=10, -R=0, --=13$$

Question

Plants N trees to get the right number of cells with shade(S) or root (R)



$N=2, SR=5, -R=5, --=13$

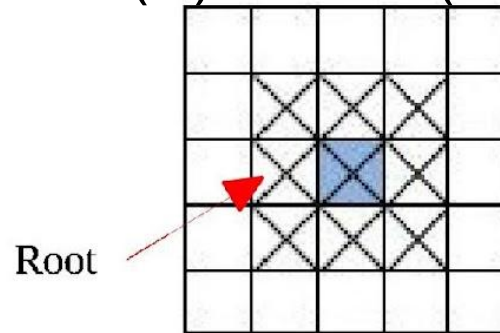
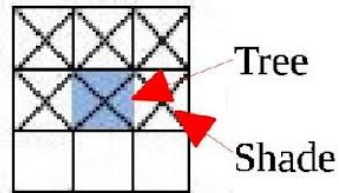
		SR	A	SR
		-R	-R	-R
SR	SR			
A	SR			
-R	-R			

$N=2, SR=10, -R=0, --=13$

* A: Apple
Tree

Question

Plants N trees to get the right number of cells with shade(S) or root (R)



$N=2, SR=5, -R=5, --=13$

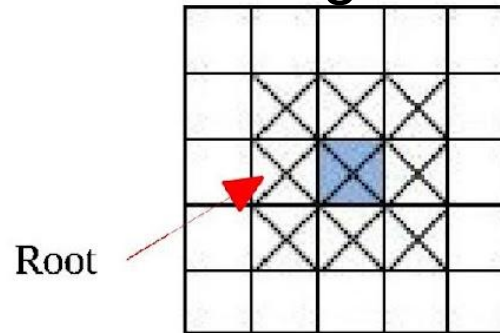
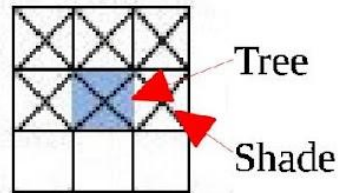
		SR	A	SR
		-R	-R	-R
SR	SR			
A	SR			
-R	-R			

$N=2, SR=10, -R=0, --=13$

	SR	SR	SR	
	SR	A	SR	
	SR	SR	SR	
	SR	A	SR	

Question

Adds vegetables (each one on 4 or 5 cells) such that it maximizes green situations



$N=2, SR=5, -R=5, --=13$

		SR	A	SR
		-R	-R	-R
SR	SR			
A	SR			
-R	-R			

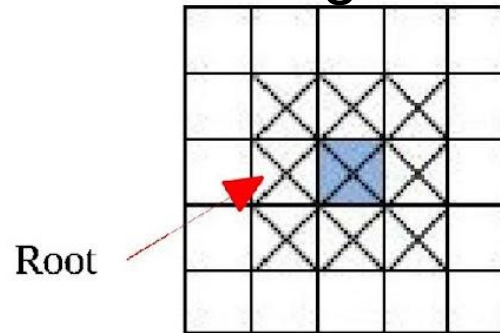
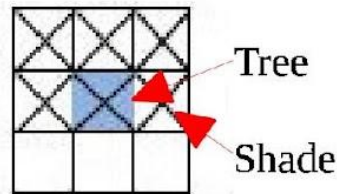
$N=2, SR=10, -R=0, --=13$

	SR	SR	SR	
	SR	A	SR	
	SR	SR	SR	
	SR	A	SR	

	Shade and root (SR)	Sun and root (-R)	Sun and no root (--)
Lettuce			
Tomato			
Onion			
Melon			
Carrot			

Question

Adds vegetables (each one on 4 or 5 cells) such that it maximizes green situations



$N=2, SR=5, -R=5, --=13$ $Obj=23$

T	M	L	A	L
T	T	O	O	O
L	L	M	T	M
A	L	C	C	C
M	O	C	T	C

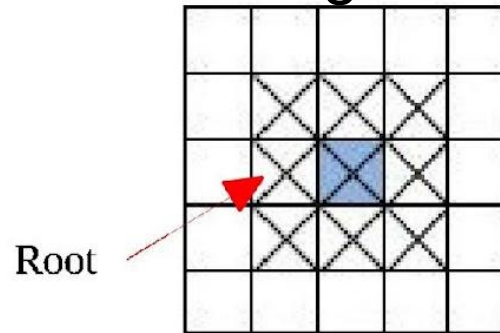
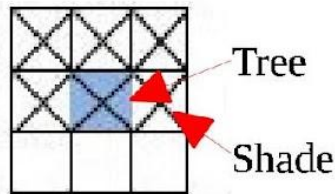
$N=2, SR=10, -R=0, --=13$

	SR	SR	SR	
	SR	A	SR	
	SR	SR	SR	
	SR	A	SR	

	Shade and root (SR)	Sun and root (-R)	Sun and no root (--)
Lettuce			
Tomato			
Onion			
Melon			
Carrot			

Question

Adds vegetables (each one on 4 or 5 cells) such that it maximizes green situations



$N=2, SR=5, -R=5, --=13 \text{ Obj}=23$

T	M	L	A	L
T	T	O	O	O
L	L	M	T	M
A	L	C	C	C
M	O	C	T	C

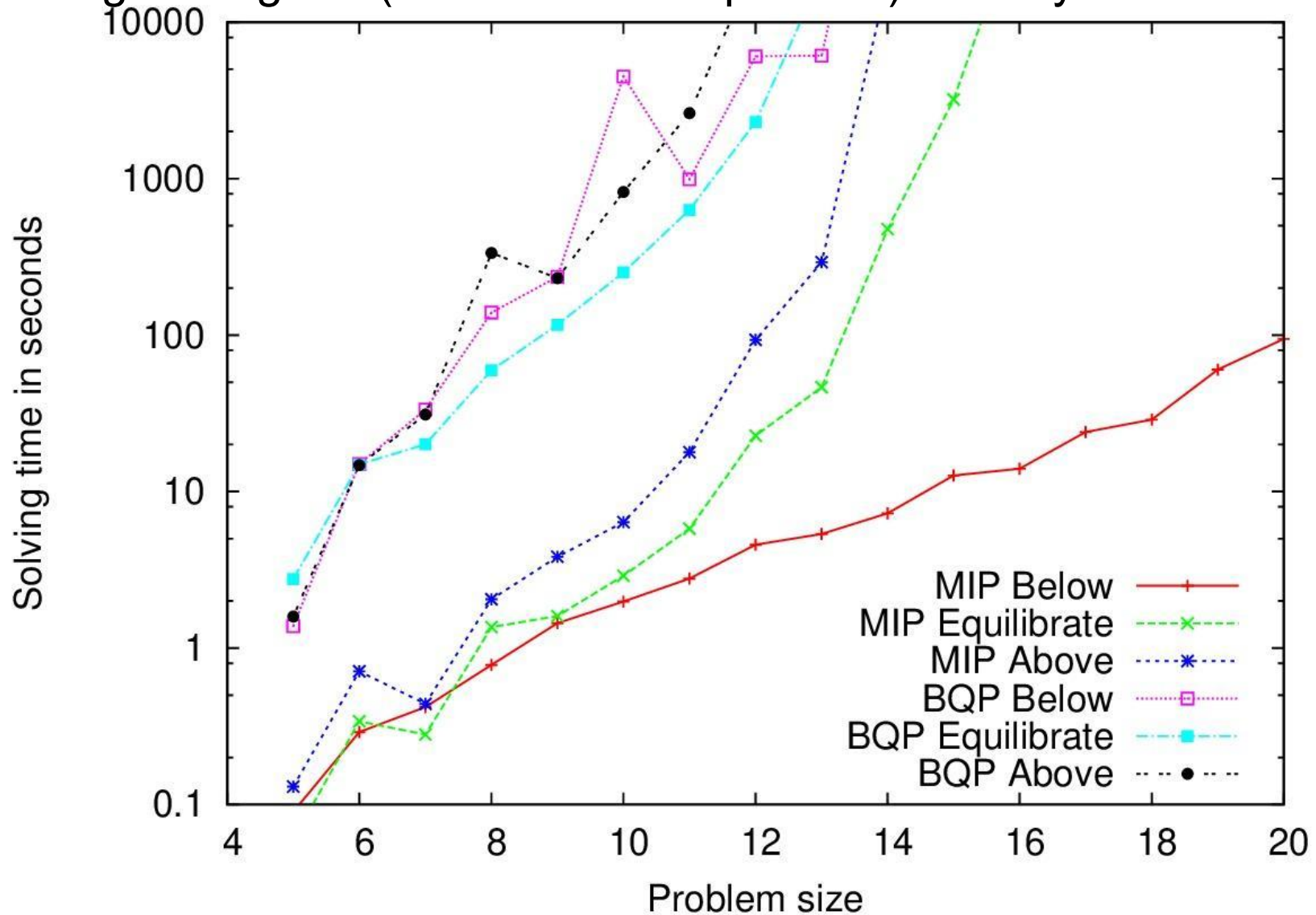
$N=2, SR=10, -R=0, --=13 \text{ Obj}=19$

M	C	C	M	C
M	L	<u>O</u>	L	T
T	L	A	<u>O</u>	<u>O</u>
C	<u>O</u>	T	L	M
T	L	A	T	M

	Shade and root (SR)	Sun and root (-R)	Sun and no root (--)
Lettuce			
Tomato			
Onion			
Melon			
Carrot			

Experimental Results using cplex 12.7

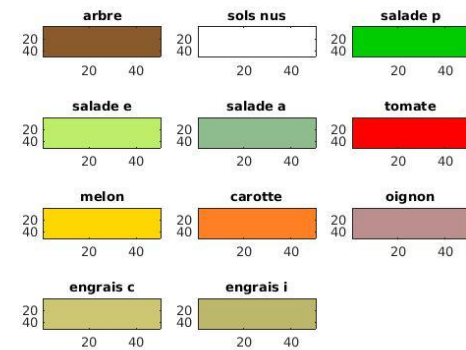
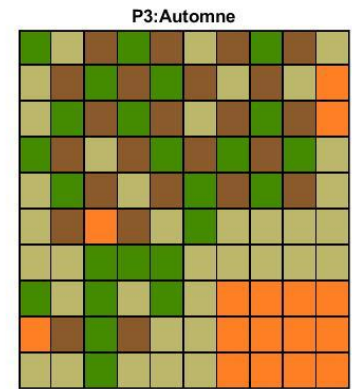
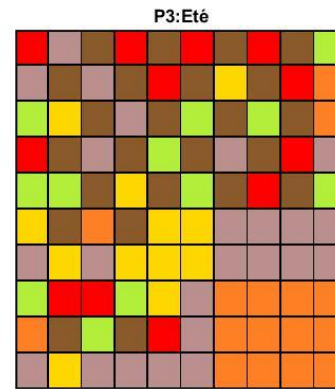
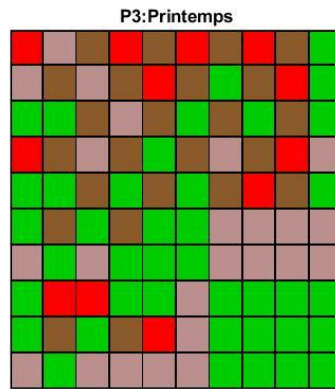
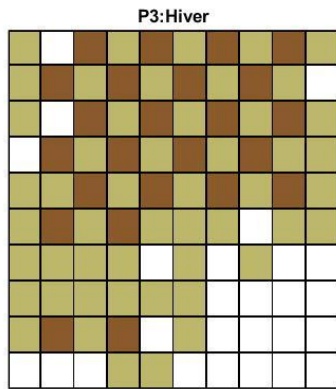
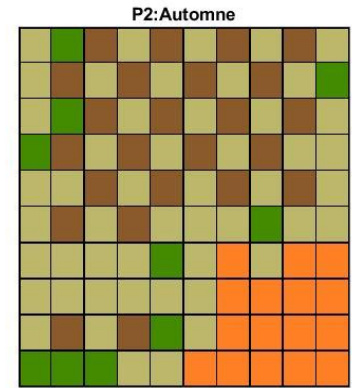
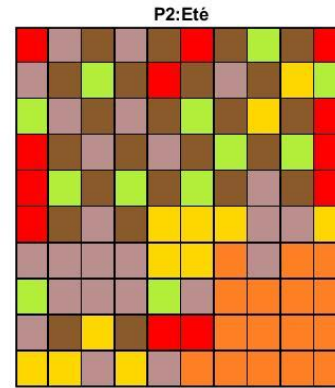
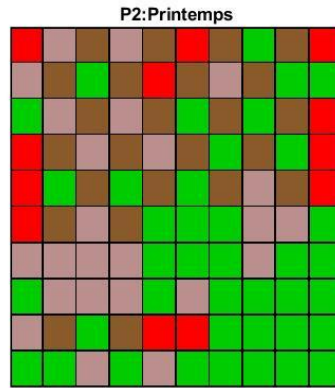
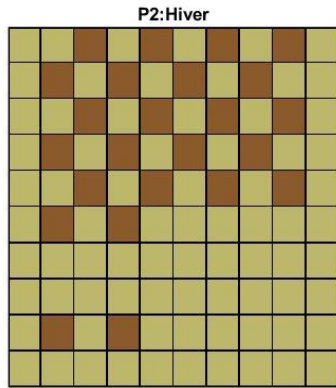
Mixed Integer Program (Benders decomposition) / Binary Quadratic Program



Time to reconstruct a full crop allocation plan from MIP is negligible (greedy algorithm)

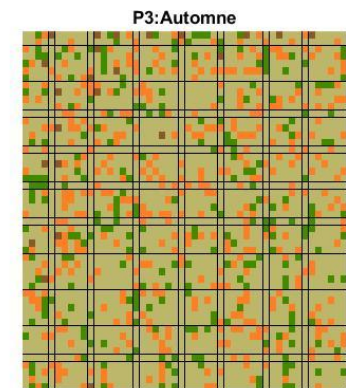
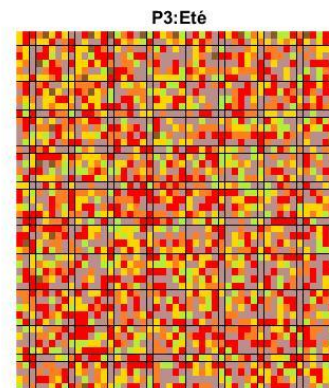
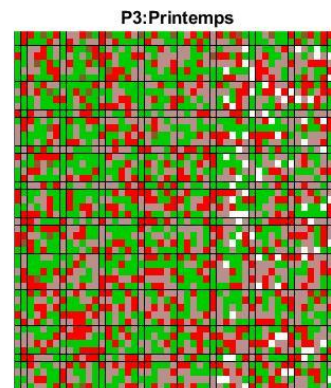
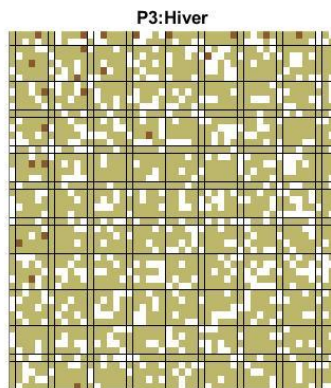
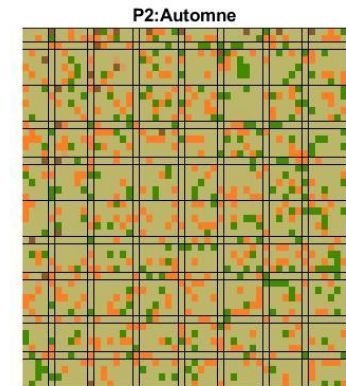
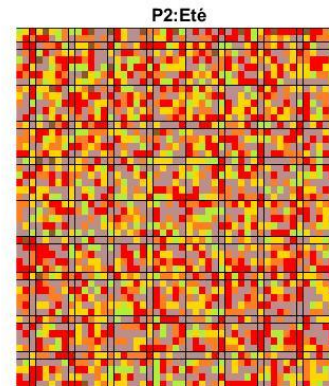
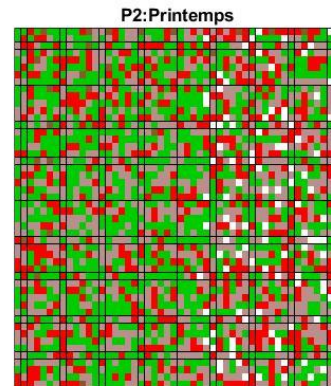
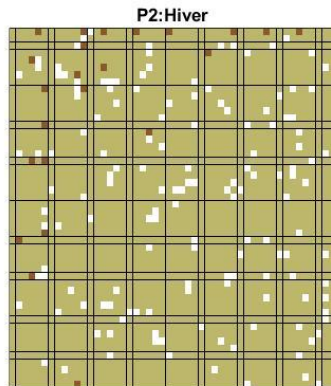
Mixed Fruit-Vegetable Crop Allocation Problem

10 x 10



(iEMSs 2016, IEA/AIE 2017)

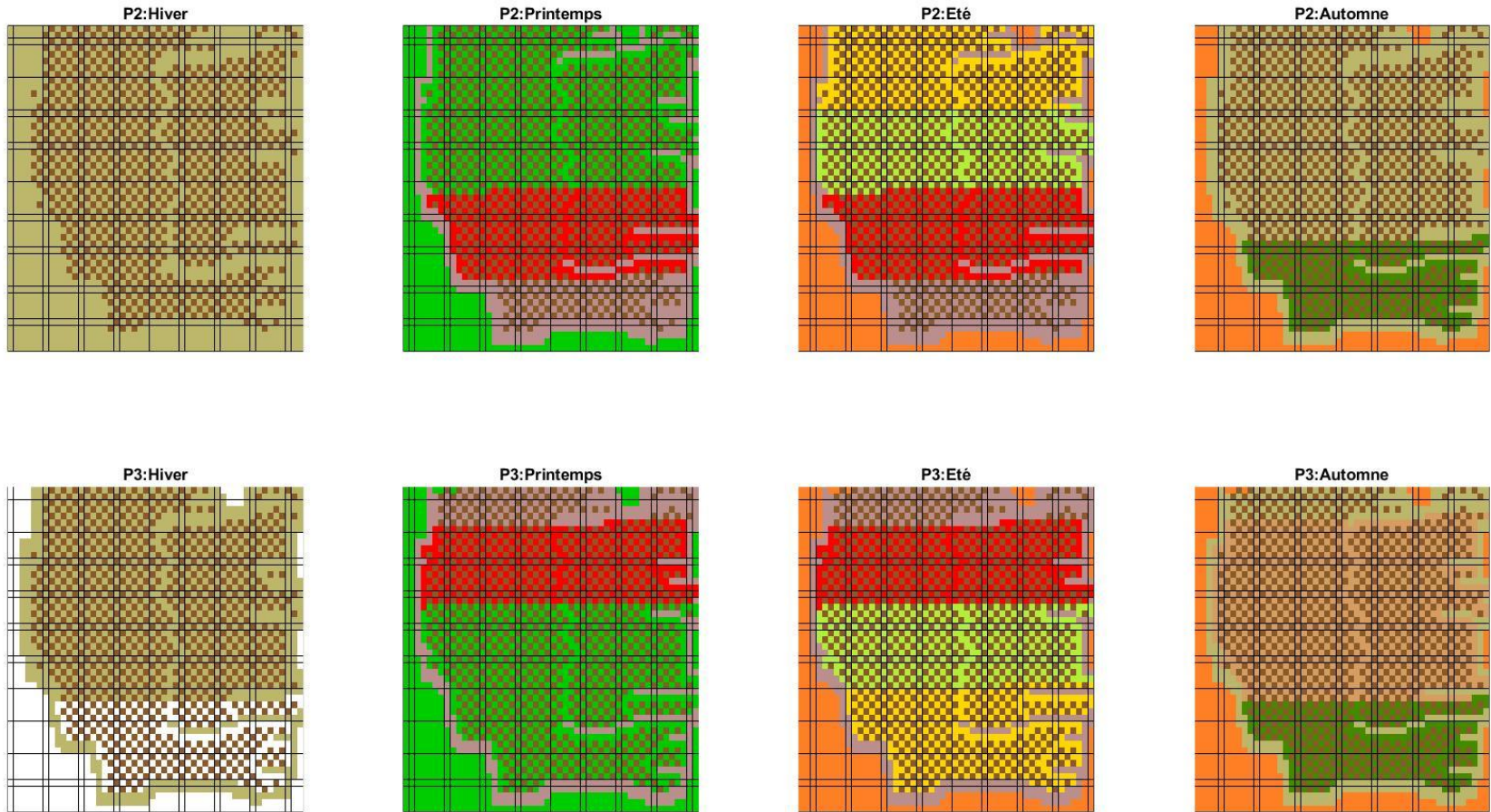
50 x 50



Best solution found by *baryonyx** 0.4 after 1h on 30 cores of AMD Opteron 2.3GHz (01LP-SPP model, 2 million Boolean variables!)

* <https://github.com/quesnel/baryonyx>

50 x 50



MIP best solution found by cplex 12.8 after 1h on 30 cores of AMD Opteron 2.3GHz (MIP model with 12500 Boolean variables and 276 continuous variables)

Some tips

- Collaborate with researchers from the other discipline
- Get access to (new) **real data**
- Collect articles, software, contacts to build an overview of the field
- Select journals / (interdisciplinary) conferences to publish
- Validate the results by **domain experts**
- Enjoy learning a new (complex) discipline!
- PhD subject is a bet! (from your supervisors ;-)

Before the thesis

- How to be recruited
 - Long meeting with the supervisor(s)
 - Master first research experience (technical report)
 - Relevance of your Master courses to the subject (course marks/ranking)
 - **Funding** (National/European projects are usually more restrictive, already specified, less innovation than University doctoral school funding)
- Building the subject / project
 - A particular story between (at least) **two** supervisors
 - Better if they have already collaborate in the past
 - Useful for the society ? => complex subjects requiring interdisciplinary approaches
 - Sometimes good subjects need a long time to be mature (between the supervisors)
 - **Subject can be revised**, it depends on the student interests (eg. more theory vs practical)
 - Associate the student to its construction
 - See as a whole in the student curriculum

During the thesis

- Supervision

- Diversity of supervisors (at least 2!) => more people, more pressure, more difficult to make choices
- Identifying all the actors for the thesis and their role
- Working environment
- Frequency of meetings (in person), **regularity**
- Places for informal meetings
- Collective agility and flexibility (subject evolution)
- Anticipate and real-time modifications
- **Strategy of publications**: where (conferences/journals), when (deadlines), what (reselling), who (author ordering)
 - it depends on the scientific discipline (computer science, biology,...)
 - doctoral schools usually ask for at least one (two?) selective publication(s)
 - after that you have more freedom!
 - see past [Best Dissertation Awards at CP](#) for some good examples
- Associate the PhD student to the decision process
- **Benevolent attitude, recognition at work**
 - for the student: be rigorous, open-mind, be a diplomat
 - for the supervisors: mutual trust, back and forth between freedom and guidance

During the thesis

- **Developing skills**
 - Which skills? (theory, engineer, relational/meeting, supervision of trainees,...)
 - Recognition, value for the company
 - PhD committee during the thesis (mid-term evaluation)
 - Doctoral school supports (interview, courses)
 - Interdisciplinary conferences
 - **Foreign lab visits**

After the thesis

- **Anticipate** after-the-thesis during your thesis!
 - What is your career plan?
 - Inform supervisors of your career plan (academy / industry)
 - List of your skills
 - Career counselor
 - Doctoral school training / workshops (like CP Doctoral Program!)
- **Post-doc strategy**
 - Social network (of supervisors)
 - supervisors should help
 - Curriculum vitae
 - Publication strategy
 - Doing applied research in your second discipline (being a facilitator, risk taking)
- To be hired
 - Interdisciplinary institutes (University, National Research Institute,...)
 - Raise awareness of the jury members
 - Adapt to the job market (additional training,..)

Professional experiences

- **Marti Sanchez**, PhD in 2006, postdoc at INRAE (2006-2008) and Barcelona University, tenure track at Universitat Pompeu-Fabra, Barcelona, Spain, in 2022 (work in neuro-robotics)
- **Aurélie Favier**, PhD in 2011, 2-year postdoc at University College Cork, secondary school teacher in Bordeaux, France since 2014
- **Jimmy Vandel**, PhD in 2012, postdocs at CEA Grenoble, LIRMM Montpellier and Lille University, permanent research engineer at CNRS, Lille, France since 2020 (work on a bioinformatics and biostatistics platform)
- **Mahuna Akplogan**, PhD in 2013, recruited in 2012 by a private French company (R&D consultant in optimization, work on nurse scheduling, healthcare and rescue,...)
- **Clément Viricel**, PhD in 2017, temporary associate professor at Lyon University, freelance data-scientist (work on neural nets)
- **Sara Maqrot**, PhD in 2019, temporary associate professor at Toulouse University (2018-2020), research engineer at ONERA (2021-2022) and permanent position at Berger Levrault, Toulouse, France since 2022
- **Manon Ruffini**, PhD in 2021 ([My Thesis in 180s](#)), recruited in Aibstract, Albi, France since 2021 (work on automatic music generation)