Project started in december 2009

OptiMAS: a decision support tool to conduct Marker Assisted Selection (MAS) programs Generation F. Valente, F. Gauthier, J. Joets

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La fenne Du Moujon

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Initial user cases: 8 crops across 32 countries

Cereals	
Maize	Africa: Angola, Ethiopia, Kenya, Malawi, Mozambique, Tanzania, Uganda, Zambia, Zimbabwe Asia: China, India, Indonesia, Thailand, The Philippines, Vietnam
Rice	Africa: Benin, Burkina Faso, Ethiopia, Gambia, Ghana, Guinea, Liberia, Madagascar, Mali, Mozambique, Nigeria, Rwanda, Senegal, Tanzania, Uganda Asia: Bangladesh, Cambodia, China, India, Indonesia, Laos, Nepal, Pakistan, Sri Lanka, Vietnam
Sorghum	Mali
Wheat	Africa: Ethiopia, Kenya Asia: China, India
Legume	S.
Beans	Ethiopia, Kenya, Tanzania, Malawi
Chickpeas	Africa: Ethiopia, Kenya Asia: India
Cowpeas	Burkina Faso, Mozambique, Senegal
	id tubers
Cassava	Ghana, Nigeria, Tanzania, Uganda 👘 🏒 👘 👘
	Cereals
	Legumes
	Root and tubers







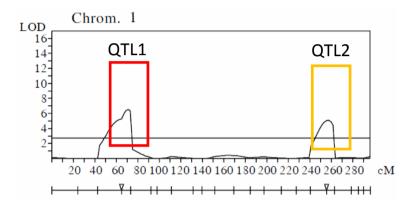






Breeding decisions adapted to different crops / projects

P1 (allele « a ») \times P2 (allele « b »)



- QTL will be detected for different traits of interest and favorable alleles will be found
- Aim: to create a target genotype with all the favorable alleles at the QTL positions

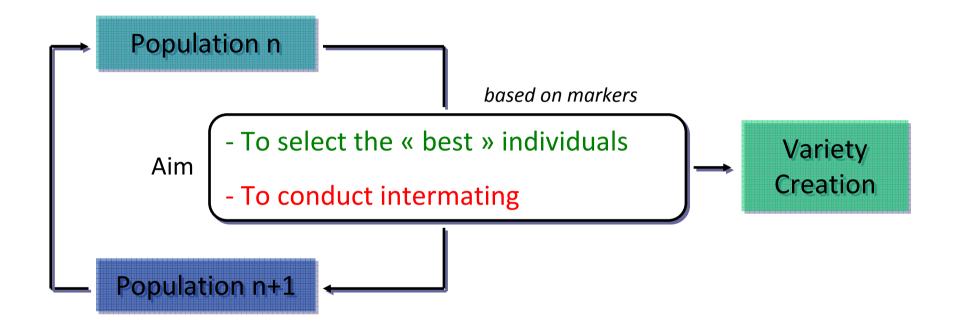


Mostly biparental population(s), but several cases where populations are connected through common parent(s)

Common objective: to create new genetic materials assembling favorable QTL alleles from 2 parents or several parents (get challenging as the No. of QTLs increases!)

→ Need user oriented tool to <u>assemble favorable alleles</u> through Marker Assisted Recurrent Selection (MARS) strategies

Implementation of Marker Assisted Recurrent Selection in OptiMAS

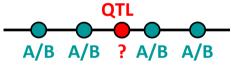


- Objective 1: to develop algorithms to compute probabilities of allele transmission through generations
- ✓ Objectif 2: to identify the best individuals
- Objective 3: to develop methodology to identify the best intermating scheme to accumulate favourable alleles and to extract varieties

Background elements on Objective 1: probabilities of allele transmission at QTLs

Aim: to evaluate individuals, we need to know which parental alleles are transmitted at the QTL position, based on the information from neighboring markers

QTL position rarely located at a marker \rightarrow QTL alleles are unknown and must be inferred from flanking markers

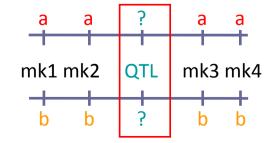


Selection of a subset of informative markers to follow the favorable parental alleles (based on haplotypes)

(bi/multi allelic context)

e.g 4 markers / QTL

Identification of all the possible phased genotypes for each QTL and computation of their probabilities



probabilities of allele transmission

- Information available (input files: pedigree/genotype & genetic map)
 - ✓ Pedigree
 - Molecular markers (observed genotypes)

Id	P1	P2	Step	Cycle	Group	mrk1	mrk2	mrk3
Fr2	а	а	IL	IL		А	А	А
Fr252	b	b	IL	IL		В	В	В
f1	Fr2	Fr252	CR	F1		A/B	A/B	A/B
ind1	f1	f1	S2	F3		А	А	А
ind2	f1	f1	S2	F3		А	А	А
ind3	f1	f1	S2	F3		A/B	-	А
ind4	f1	f1	S2	F3		В	В	A/B
ind5	f1	f1	S2	F3		В	В	В
ind6	f1	f1	S2	F3				
ind7	f1	f1	S2	F3		А	А	А

probabilities of allele transmission

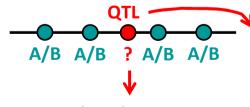
Information available (input files: pedigree/genotype & genetic map)

- ✓ Pedigree
- Molecular markers (observed genotypes)
- QTL position
- ✓ Distance between loci (Haldane \rightarrow recombination rate)

Locus	Chr	QTL	Pos	All+
marker1	1	1	42.2	
marker2	1	1	64.0	
qtl1	1	1	70.0 م	а
marker3	1	1	d 72.5	
marker4	1	1	90.8	
marker5	1	2	237.1	
marker6	1	2	252.2	
qtl2	1	2	254.0	b
marker7	1	2	259.5	
marker8	1	2	274.8	

Objective 1: probabilities of allele transmission

- Information available (input files: pedigree/genotype & genetic map)
 - ✓ Pedigree
 - Molecular markers (observed genotypes)
 - QTL position
 - \checkmark Distance between loci (Haldane \rightarrow recombination rate)
- Algorithm: compute the probabilities of allele transmission in different MAS schemes and mating designs (intercrossing, selfing, backcrossing, DH, RIL)
 - \checkmark 2 versions: command line \rightarrow integrated in a new GUI



p(a/a) = 0.9, p(a/b) = 0.1, $p(b/b) = 0 \rightarrow$ probability of each genotype at QTL given markers

Molecular Score

MS = 0.95 \rightarrow Probability to carry favorable allele at the QTL position

Presentation of mean features of OptiMAS

■ New Graphical User Interface (GUI) organized in 3 modules corresponding to the ≠ steps of the selection program

OptiMAS: a decision support tool to conduct	Marker Assisted Selection programs - 1.0		
File Visualization Data Help			
Detimas	Computation of genotypic probabiliti	es - Estimation of gene	tic values
A Step 1 - Prediction Predictio	Homo / Hetero Estimation of parental alleles probability	Graphs	Weight 1,00 + Apply

<u>Step 1</u>: Computation of genotypic probabilities – Estimation of genetic values

Molecula	ar score pi	rediction	Homo	/Hetero	Esti	mation of p	parental	alleles prob	ability	Graphs								
🤍 Find	F) 🔳	View	🛺 Weigł	nt QT	1 01 🗔	D		<u> </u>							Weight	1,00	9 (Apply
Id	P1	P2	Cycle	Group	MS 🛧	Weight	UC	No.(+/+)	No.(-/-)	No.(+/-)	No.(?)	QTL1	QTL2	QTL3	QTL4	QTL5	QTL6	QTL7
B8	A1005	A1005	C2	-	0.8366	0.8366	10	8	1	0	2	0.0000	0.8598	0.9761	0.8752	0.8959	0.9745	0.9960
B158	A251	A1005	C2	G1	0.8024	0.8024	10	7	1	1	2	0.0000	0.8792	0.8925	0.9405	0.9731	0.9693	0.9960
B28	A1006	A251	C2	G1	0.7740	0.7740	9,7071	6	1	1	3	0.0000	0.9716	0.4938	0.9492	0.9137	0.9199	0.9797
B13	A1006	A1005	C2	-	0.7609	0.7609	9,7071	6	1	2	2	0.0000	0.8844	0.9559	0.9179	0.9731	0.4963	0.9895
B38	A1040	A1005	C2	-	0.7494	0.7494	9,366	6	1	1	3	0.0000	0.9349	0.4907	0.9429	0.5822	0.9260	0.9860
B37	A1040	A1005	C2	5.	0.7433	0.7433	9	7	1	1	2	0.0000	0.9528	0.2334	0.9172	0.8735	0.9718	0.9860
B40	A1040	A1040	C2	3	0.7404	0.7404	9,366	7	1	2	1	0.0000	0.9775	0.4376	0.9592	0.4974	0.9231	0.9759
B242	A37	A1005	C2	2	0.7305	0.7305	9,366	6	1	1	3	0.0000	0.7259	0.8959	0.8879	0.2896	0.9722	0.9860
B124	A23	A167	C2		0.7223	0.7223	9,366	6	1	1	3	0.4812	0.6698	0.9559	0.9429	0.2787	0.9614	0.9924
B125	A23	A167	C2	-	0.7032	0.7032	8,7071	6	2	1	2	0.4812	0.6536	0.9559	0.9429	0.0058	0.9616	0.9924
A1040_	sf683	df37	CI		0.6615	0.6615	8,618	5	1	5	0	0.0000	0.9771	0.4842	0.9592	0.4974	0.9231	0.9759
A1040	A1040_	A1040_	÷	-	0.6615	0.6615	8,618	5	1	0	5	0.0000	0.9771	0.4842	0.9592	0.4974	0.9231	0.9759
A1005_	df108	sf650	C1	-	0.6418	0.6418	8,618	5	1	4	1	0.0000	0.8928	0.4896	0.8751	0.4878	0.9747	0.9960
A1005	A1005_	A1005_	5		0.6417	0.6417	8,618	5	1	0	5	0.0000	0.8927	0.4895	0.8751	0.4878	0.9747	0.9960
A1003	A1003_	A1003_	3	3	0.6413	0.6413	8	6	3	0	2	0.0000	0.9757	0.9738	0.9135	0.0000	0.4986	0.9759
A1003_	df108	df37	C1	2	0.6413	0.6413	8	6	3	1	1	0.0000	0.9757	0.9738	0.9135	0.0000	0.4987	0.9759
A251_	sf683	df108	CI	4	0.6364	0.6364	8,618	3	1	5	2	0.0000	0.5014	0.4896	0.9544	0.4974	0.9090	0.9960
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□ Individuals in lines, MS value $[0-1] \rightarrow 1 = ideotype$

<u>Step 1</u>: Computation of genotypic probabilities – Estimation of genetic values

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 | 0.9761 | 0.8752 | 0.8959 | 0.9745 | 0.9960
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All: f 1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 0.9 All: f 1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 1 2 0.0000 0.9 87 A1005 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.9 87 A167 C2 - 0.7305 0.7325 9,366 6 1 1 3 0.4812 0.6 23 A167 C2 - 0.7032 0.7032 8,7071 6 2 1 2 0.4812 0.6 1040 A1040 - - 0.6615 0.6615 8,618 5 1 0 0.0000 0.8 6 0.0000 0.8 6 0.0000 0.8 6 | 1040 A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 0.0000 0.9 QTL: 2
All+: f 1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 0.9 QTL: 2
All+: f 1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 0.0000 0.9 87 A1005 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.9 87 A107 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.7 81 A167 C2 - 0.7032 0.7032 8,7071 6 2 1 2 0.4812 0.6 833 df37 C1 - 0.6615 0.6615 8,618 5 1 0 0.0000 0.9 $d_{d:s=0.029221}$ $d_{d:s=0.02921}$ 1005 A1040_ - | A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 0.0000 0.7 A1005 C2 - 0.7433 0.7433 0.7433 9 7 1 1 2 0.0000 0.7 A1005 C2 - 0.7403 0.7404 9,366 7 1 1 2 0.0000 0.7 A1005 C2 - 0.7404 0.7404 9,366 7 1 1 2 0.0000 0.7 A1005 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.7 A107 C2 - 0.7302 0.7322 9,366 6 1 1 3 0.4812 0.6 A167 C2 - 0.7032 0.7032 8,707 6 2 1 2 0.4812 0.6 A167 C1 - 0.6615 0.6615 8,618 5 1 0 5 0.0000 0.6 1005 <td>A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 0.000 0.7 All+: f 0.040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 0.9 0.040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 0.0000 0.9 Genetype 0.040 A1040 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.7 87 A1057 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.7 87 A167 C2 - 0.7032 0.7022 8,7071 6 2 1 2 0.4812 0.6 883 df37 C1 - 0.6615 8,618 5 1 0 0.0000 0.9 des-0.00418 s:s=0.000186 d:d=0.0000 des-0.013427 des-0.013427 des-0.013427 des-0.013427 des-0.013427</td> | A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 0.000 0.7 All+: f 0.040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 0.9 0.040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 0.0000 0.9 Genetype 0.040 A1040 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.7 87 A1057 C2 - 0.7305 0.7305 9,366 6 1 1 3 0.0000 0.7 87 A167 C2 - 0.7032 0.7022 8,7071 6 2 1 2 0.4812 0.6 883 df37 C1 - 0.6615 8,618 5 1 0 0.0000 0.9 des-0.00418 s:s=0.000186 d:d=0.0000 des-0.013427 des-0.013427 des-0.013427 des-0.013427 des-0.013427 |

- □ Prob(++/--/+-) at the QTL position + detailed genotype in terms of parental alleles
- **Estimation (probability) of parental allele transmission**

<u>Step 1</u>: Computation of genotypic probabilities – Estimation of genetic values

Rind	View Weight QTL 01	a a a	1.		1.0	-	1		Weight	1,00	<u>+</u>] [_	Apply
Id	P1 P2 Cycle Group MS A Weight U	C No.(+/+)	No.(-/-)	No.(+/-)	No.(?)	QTL1	QTL2	QTL3	QTL4	QTL5	QTL6	QTL7
B8 /	Visualization of genotypes	8	1	0	2	0.0000	0.8598	0.9761	0.8752	0.8959	0.9745	0.9960
B158 /	Visualization of genotypes	7	1	1	2	0.0000	0.8792	0.8925	0.9405	0.9731	0.9693	0.9960
B28 /	The probabilities to be homozygous /	16	1	1	3	0.0000	0.9716	0.4938	0.9492	0.9137	0.9199	0.9797
B13 /	heterozygous, at the QTL positions, have been	16	1	2	2	0.0000	0.8844	0.9559	0.9179	0.9731	0.4963	0.9895
B38 /	computed according to favourable / unfavourable grouping of founder alleles.	6	1	1	3	0.0000	0.9349	0.4907	0.9429	0.5822	0.9260	0.9860
B37		7	1	1	2	0.0000	0.9528	0.2334	0.9172	0.8735	0.9718	0.9860
B40	Set a threshold and select a color to display a new view of the molecular score table based on	7	1	2	1	0.0000	0.9775	0.4376	0.9592	0.4974	0.9231	0.9759
B242	genotypes.	6	1	1	3	0.0000	0.7259	0.8959	0.8879	0.2896	0.9722	0.9860
B124	Customize cut-off/colours:	6	1	1	3	0.4812	0.6698	0.9559	0.9429	0.2787	0.9614	0.9924
B125	Prob(+/+) ≥ \$75 +	16	2	1	2	0.4812	0.6536	0.9559	0.9429	0.0058	0.9616	0.9924
A1040_ s	Prob(+/-) ≥ 0,75 ÷	5	1	5	0	0.0000	0.9771	0.4842	0.9592	0.4974	0.9231	0.9759
A1040	Prob(+/-) ≥ 0,75 😴	5	1	0	5	0.0000	0.9771	0.4842	0.9592	0.4974	0.9231	0.9759
A1005_ d	Prob(-/-) ≥ 0,75 🖕	5	1	4	1	0.0000	0.8928	0.4896	0.8751	0.4878	0.9747	0.9960
A1005		5	1	0	5	0.0000	0.8927	0.4895	0.8751	0.4878	0.9747	0.9960
A1003	The rest: uncertain genotypes (?)	6	3	0	2	0.0000	0.9757	0.9738	0.9135	0.0000	0.4986	0.9759
A1003_ d		6	3	1	1	0.0000	0.9757	0.9738	0.9135	0.0000	0.4987	0.9759
A251_ s	Reset	3	1	5	2	0.0000	0.5014	0.4896	0.9544	0.4974	0.9090	0.9960

□ Visualization of genotypes via colors for each QTL (how many QTL are fixed ?)

<u>Step2</u>: Selection of individuals

Manual selection

🔍 Fin	nd 📃 📃	View	4 Weig	ht QT	L 01 💷	D				1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1 1	10 20				Veight	1,00	<u>+</u>	Apply
Id	P1	P2	Cycle	Group	MS ^	Weight	UC	No.(+/+)	No.(-/-)	No.(+/-)	No.(?)	QTL1	QTL2	QTL3	QTL4	QTL5	QTL6	QTL7
8	A1005	A1005	C2	-	0.8366	0.8366	10	8	1	0	2	0.0000	0.8598	0.9761	0.8752	0.8959	0.9745	0.9960
158	A251	A1005	C2	G1	0.8024	0.8024	10	7	1	1	2	0.0000	0.8792	0.8925	0.9405	0.9731	0.9693	0.9960
28	A1006	A251	C2	G1	0.7740	0.7740	9,7071	6	1	1	3	0.0000	0.9716	0.4938	0.9492	0.9137	0.9199	0.9797
13	A1006	A1005	C2	-	0.7609	0.7609	9,7071	6	1	2	2	0.0000	0.8844	0.9559	0.9179	0.9731	0.4963	0.9895
38	A1040	A1005	C2	-	0.7494	0.7494	9,366	6	1	1	3	0.0000	0.9349	0.4907	0.9429	0.5822	0.9260	0.9860
													10.0	1.1				
37 40	Add	to list ALU4U	List S List S	ld to list election election Manual	1		,366	7 7	1	1 2	2	0.0000		20.000		0.8735 0.4974	0.9718 0,9231	0.9860 0.9759
		mill south the late	List S List S	election election	1		,366	7 7 Selection o List of	4	s Grapł		0.0000 edigree		0.4376			1000	and the second second
		mill south the late	List S List S	election election Manual	1 2 Selection	n	,366		1 findividual ^f selected ir	s Grapł		0.0000 edigree t1_Manu	0.9775 al_Selecti	0.4376 on	0.9592	0.4974	0,9231	0.9759
-		mill south the late	List S List S	election election	1 2 Selection		,366	List of List Select List Select	1 findividual fselected ir tion 1 tion 2	s Graph ndividuals	Lis	0.0000 edigree	0.9775	0.4376	0.9592		1000	0.9759
		mill south the late	List S List S	election election Manual	1 2 Selection	n	,366	List of List Select	1 findividual fselected ir tion 1 tion 2	s Graph ndividuals		0.0000 edigree t1_Manu Id	0.9775 al_Selecti P1	0.4376 on P2	0.9592	0.4974	0.9231 Group	0.9759
-		mill south the late	List S List S	election election Manual	1 2 Selection	n	366	List of List Select List Select	1 findividual fselected ir tion 1 tion 2	s Graph ndividuals	Lis	0.0000 edigree tt1_Manu Id B13	0.9775 al_Selecti P1 A1006	0.4376 on P2 A1005	0.9592 Cy C2	0.4974 /cle	0.9231 Group	0.9759
_		mill south the late	List S List S	election election Manual	1 2 Selection	n	,366	List of List Select List Select	1 findividual fselected ir tion 1 tion 2	s Graph ndividuals	Lis 1 2 3	0.0000 edigree t1_Manu Id B13 B40	0.9775 al_Selecti P1 A1006 A1040	0.4376 on P2 A1005 A1040	0.9592 Cy Cy C2 C2	0.4974 /cle -	0.9231 Group	0.9759

Step2: Selection of individuals

- Manual selection
- **Truncation selection based on:**
 - ✓ Molecular score (MS)

I	Truncation select	ion (MTS) –					
	N _{Sel} 10	Criterion	Molecular Score 🖨	List	List2_Truncation_Selecti	Option	Run

Id B8	P1 A1005	IS_Select P2	N											4
B8	a de la compañía de la	P2	Cycle	-										
	A1005			Group	MS ^	Weight	UC	No.(+/+)	No.(-/-)	No.(+/-)	No.(?)	QTL1	QTL2	QTL3
B158		A1005	C2	-	0.8366	0.8366	10	8	1	0	2	0.0000	0.8598	0.9761
	A251	A1005	C2	G1	0.8024	0.8024	10	7	1	1	2	0.0000	0.8792	0.8925
B28	A1006	A251	C2	G1	0.7740	0.7740	9,7071	6	1	1	3	0.0000	0.9716	0.4938
B13	A1006	A1005	C2	-	0.7609	0.7609	9,7071	6	1	2	2	0.0000	0.8844	0.9559
B38	A1040	A1005	C2	-	0.7494	0.7494	9,366	6	1	1	3	0.0000	0.9349	0.4907
B37	A1040	A1005	C2	-	0,7433	0.7433	9	7	1	1	2	0.0000	0.9528	0.2334
B40	A1040	A1040	C2	-	0.7404	0.7404	9,366	7	1	2	1	0.0000	0.9775	0.4376
B242	A37	A1005	C2	÷	0.7305	0.7305	9,366	6	1	1	3	0.0000	0.7259	0.8959
B246	A37	A1040	C2	-	0.7303	0.7303	9,366	6	1	3	1	0.0000	0.6036	0.8755
B293	A9	A1040	C2	-	0.7268	0.7268	9,366	6	1	3	1	0.0000	0.9549	0.9741
			,îl											
	838 837 840 8242 8246 8293	B38 A1040 B37 A1040 B40 A1040 B242 A37 B246 A37 B293 A9	B38 A1040 A1005 B37 A1040 A1005 B40 A1040 A1040 B242 A37 A1040 B246 A37 A1040 B293 A9 A1040	B38 A1040 A1005 C2 B37 A1040 A1005 C2 B40 A1040 A1040 C2 B242 A37 A1005 C2 B246 A37 A1040 C2 B293 A9 A1040 C2	B38 A1040 A1005 C2 - B37 A1040 A1005 C2 - B40 A1040 A1040 C2 - B242 A37 A1005 C2 - B246 A37 A1040 C2 - B293 A9 A1040 C2 -	B38 A1040 A1005 C2 - 0.7494 B37 A1040 A1005 C2 - 0.7433 B40 A1040 A1040 C2 - 0.7404 B242 A37 A1005 C2 - 0.7305 B246 A37 A1040 C2 - 0.7303 B293 A9 A1040 C2 - 0.7268	B38 A1040 A1005 C2 - 0.7494 0.7494 B37 A1040 A1005 C2 - 0.7433 0.7433 B40 A1040 A1040 C2 - 0.7404 0.7404 B242 A37 A1045 C2 - 0.7305 0.7305 B246 A37 A1040 C2 - 0.7303 0.7303 B293 A9 A1040 C2 - 0.7268 0.7268	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 B37 A1040 A1005 C2 - 0.7433 0.7433 9 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 B242 A37 A1005 C2 - 0.7305 0.7305 9,366 B246 A37 A1040 C2 - 0.7303 0.7303 9,366 B293 A9 A1040 C2 - 0.7268 0.7268 9,366	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 6 B37 A1040 A1005 C2 - 0.7433 0.7433 9,366 7 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 B40 A1040 A1040 C2 - 0.7305 0.7305 9,366 6 B242 A37 A1005 C2 - 0.7305 0.7305 9,366 6 B246 A37 A1040 C2 - 0.7303 0.7303 9,366 6 B293 A9 A1040 C2 - 0.7268 0.7268 9,366 6	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 6 1 B37 A1040 A1005 C2 - 0.7433 0.7433 9 7 1 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 B242 A37 A1005 C2 - 0.7305 0.7305 9,366 6 1 B246 A37 A1040 C2 - 0.7303 0.7303 9,366 6 1 B293 A9 A1040 C2 - 0.7268 0.7268 9,366 6 1	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 6 1 1 B37 A1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 B242 A37 A1005 C2 - 0.7305 0.7305 9,366 6 1 1 B246 A37 A1040 C2 - 0.7305 0.7305 9,366 6 1 3 B293 A9 A1040 C2 - 0.7268 0.7268 9,366 6 1 3	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 B37 A1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 1 B242 A37 A1005 C2 - 0.7305 0.7305 9,366 6 1 3 3 B246 A37 A1040 C2 - 0.7303 0.7303 9,366 6 1 3 1 B293 A9 A1040 C2 - 0.7268 0.7268 9,366 6 1 3 1	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 0.0000 B37 A1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 B40 A1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 1 0.0000 B424 A37 A1005 C2 - 0.7305 0.7305 9,366 6 1 3 0.0000 B246 A37 A1040 C2 - 0.7303 0.7303 9,366 6 1 3 1 0.0000 B293 A9 A1040 C2 - 0.7268 0.7268 9,366 6 1 3 1 0.0000	B38 A1040 A1005 C2 - 0.7494 0.7494 9,366 6 1 1 3 0.0000 0.9349 B37 A1040 A1005 C2 - 0.7433 0.7433 9 7 1 1 2 0.0000 0.9328 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 0.0000 0.9528 B40 A1040 A1040 C2 - 0.7404 0.7404 9,366 7 1 2 1 0.0000 0.9578 B242 A37 A1005 C2 - 0.7305 0.7305 9,366 6 1 3 0.0000 0.7259 B246 A37 A1040 C2 - 0.7303 0.7303 9,366 6 1 3 1 0.0000 0.6036 B293 A9 A1040 C2 - 0.7268 0.7268 9,366 6 1 3 1 0.0000 0.6036 0.6036 1

Step2: Selection of individuals

- Manual selection
- Truncation selection based on:
 - ✓ Molecular score (MS)
 - Weighted MS: give more or less importance to the different QTL

O QTLs Weights	η															
Different weights for each QTL can be assigned:	List	:3_Trun	cation_W	/eight_S	election	1										+
1. Select the QTLs that have to be weighted		Id	P1	P2	Cycle	Group	MS 🛧	Weight	UC	No.(+/+)	No.(-/-)	No.(+/-)	No.(?)	QTL1	QTL2	QTL3
2. Set a weight 3. Click update	1	B8	A1005	A1005	C2	-	0.8366	0.7079	10	8	1	0	2	0.0000	0.8598	0.9761
4. Click Apply	2	B158	A251	A1005	C2	G1	0.8024	0.6790	10	7	1	1	2	0.0000	0.8792	0.8925
All None Weight 1,00 🗣 Update	3	B28	A1006	A251	C2	G1	0.7740	0.6550	9,7071	6	1	1	3	0.0000	0.9716	0.4938
	4	B13	A1006	A1005	C2	-	0.7609	0.6438	9,7071	6	1	2	2	0.0000	0.8844	0.9559
QTL Weight	5	B124	A23	A167	C2	-	0.7223	0.6852	9,366	6	1	1	3	0.4812	0.6698	0.9559
1 QTL1 3,00 🜩	6	B125	A23	A167	C2	-	0.7032	0.6691	8,7071	6	2	1	2	0.4812	0.6536	0.9559
2 QTL2 1,00 🜩	7	B110	A212	A1005	C2	-	0.6863	0.6481	9,2247	4	0	4	3	0.4376	0.7319	0.4876
3 QTL3 1,00 🜩	8	B57	A167	A1040	C2	-	0.6860	0.6545	8,8229	4	0	6	1	0.4812	0.6066	0.4230
4 QTL4 1,00 🜩	9	B123	A212	A91	C2	-	0.6717	0.6390	8,618	4	1	3	3	0.4590	0.6119	0.4879
5 QTL5 1,00 🜩	10	B47	A166	A167	C2	-	0.6309	0.6570	8,366	4	2	2	3	0.8010	0.4966	0.8733
																►
7 QTL7 1,00 🗢	No	ind =	10		N	lo. group	= 1									
Reset Cancel Apply	Me	an = 0.	727447		V	ar = 0.00	372879									
Cancer Apply																

Step2: Selection of individuals

- Manual selection
- Truncation selection based on:
 - ✓ Molecular score (MS)
 - Weighted MS: give more or less importance to the different QTL
 - Utility criterion: select candidates based on the possibility of obtaining superior genotype in their progeny (favor heterozygous)
- QTL complementation selection

(Hospital et al., 2000)

<u>Step2</u>: Selection of individuals – QTL complementation selection (QCS) (Hospital et al., 2000)

List	4_Com	plementa	tion_Sel	ection															\$
	Id	P1	P2	Cycle	Group	MS 🛧	Weight	UC	No.(+/+)	No.(-/-)	No.(+/-)	No.(?)	QTL1	QTL2	QTL3	QTL4	QTL5	QTL6	QTL7
1	B8	A1005	A1005	C2	-	0.8366	0.7079	10	8	1	0	2	0.0000	0.8598	0.9761	0.8752	0.8959	0.9745	0.9960
2	B158	A251	A1005	C2	G1	0.8024	0.6790	10	7	1	1	2	0.0000	0.8792	0.8925	0.9405	0.9731	0.9693	0.9960
3	B28	A1006	A251	C2	G1	0.7740	0.6550	9,7071	6	1	1	3	0.0000	0.9716	0.4938	0.9492	0.9137	0.9199	0.9797
4	B13	A1006	A1005	C2	-	0.7609	0.6438	9,7071	6	1	2	2	0.0000	0.8844	0.9559	0.9179	0.9731	0.4963	0.9895
5	B38	A1040	A1005	C2	-	0.7494	0.6341	9,366	6	1	1	3	0.0000	0.9349	0.4907	0.9429	0.5822	0.9260	0.9860
6	B37	A1040	A1005	C2	-	0.7433	0.6290	9	7	1	1	2	0.0000	0.9528	0.2334	0.9172	0.8735	0.9718	0.9860
7	B40	A1040	A1040	C2	-	0.7404	0.6265	9,366	7	1	2	1	0.0000	0.9775	0.4376	0.9592	0.4974	0.9231	0.9759
8	B242	A37	A1005	C2	-	0.7305	0.6181	9,366	6	1	1	3	0.0000	0.7259	0.8959	0.8879	0.2896	0.9722	0.9860
													+						
9	B124	A23	A167	C2	-	0.7223	0.6852	9,366	6	1	1	3	0.4812	0.6698	0.9559	0.9429	0.2787	0.9614	0.9924
10	B125	A23	A167	C2	-	0.7032	0.6691	8,7071	6	2	1	2	0.4812	0.6536	0.9559	0.9429	0.0058	0.9616	0.9924

θ_{Ms}

nT

2

0,70

10

Take into account complementarities between candidates regarding the favorable alleles they carry

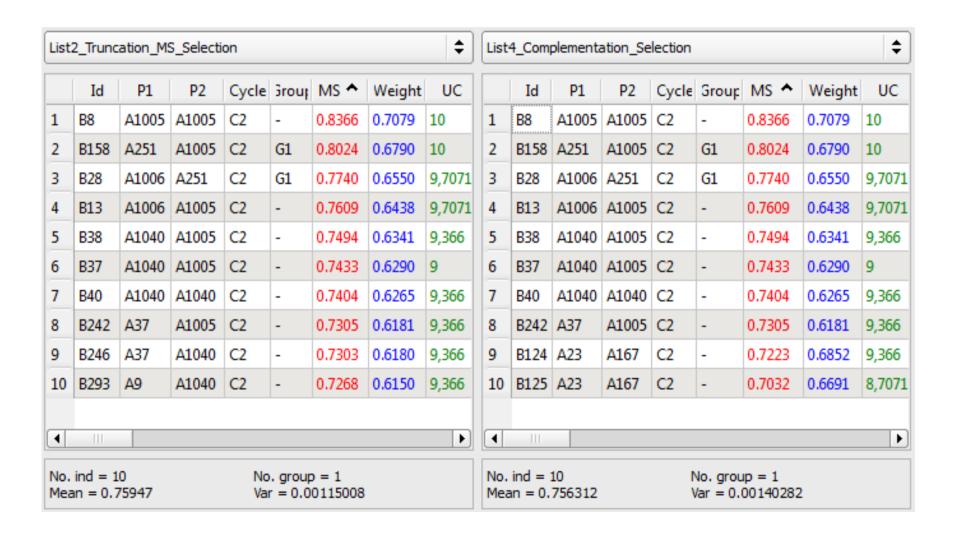
Prevent the loss of rare favourable alleles and fixation MSmin ≥ of unfavourable alleles at QTL with small effects. Nmax =

AQCS strategy is described by four parameters: (Hospital et al., 2000)

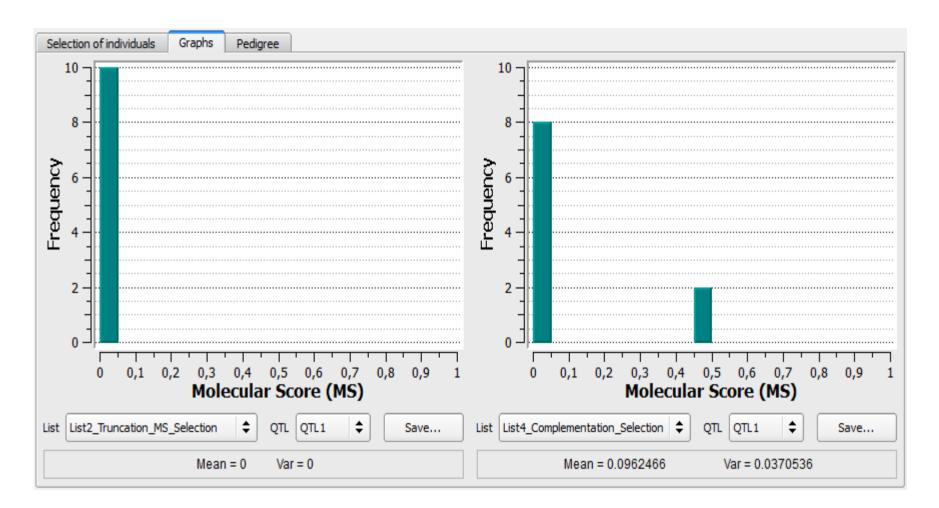
- The threshold for Molecular Score (MS), above which a favourable QTL allele is declared 'present'.
 - ▲ Each QTL is requested to be 'present' in at least n_T selected individuals.
 - The minimum threshold value (Molecula Score) for the addition of an individual.

Maximum number of individuals selected at the end of the complementation process.

<u>Step2</u>: Selection of individuals – Comparison between lists of selected individuals

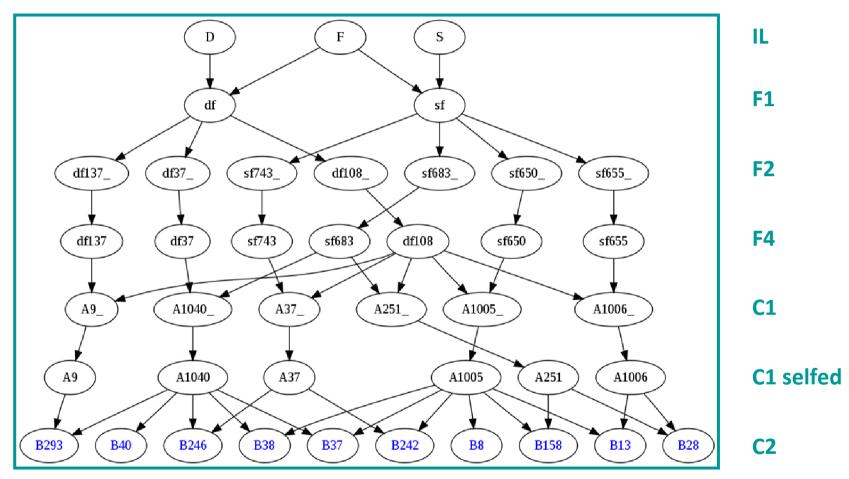


<u>Step2</u>: Selection of individuals – Comparison between lists of selected individuals



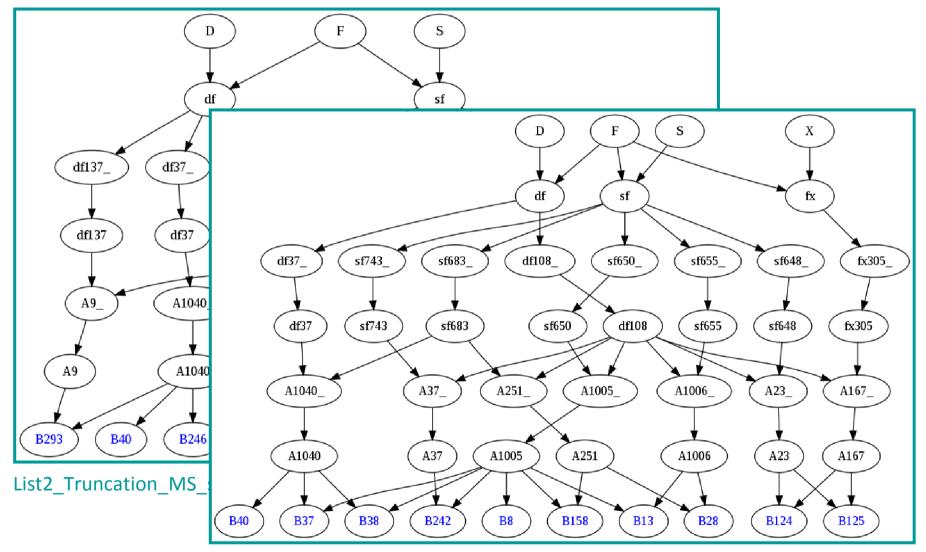
Distribution of individuals for each list regarding the molecular score (MS)
 Favorable allele (QTL1) lost without the QTL Complementation Selection (QCS)

<u>Step2</u>: Selection of individuals – Comparison between lists of selected individuals (pedigree)



List2_Truncation_MS_selection

To follow the contribution of selected individuals over generations and prevent possible bottlenecks <u>Step2</u>: Selection of individuals – Comparison between lists of selected individuals (pedigree)

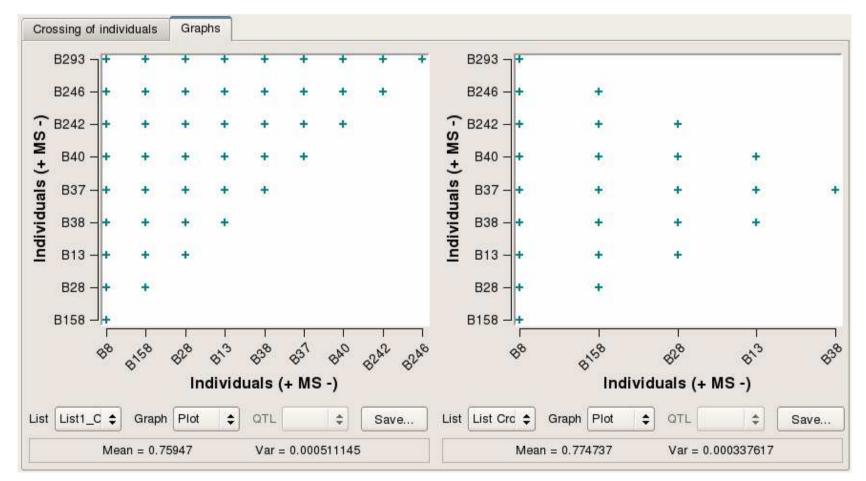


List4_Complementation_selection: 4 parental alleles present in the next generation

<u>Step3</u>: Identification of crosses to be made among selected individuals

Half-diallel between selected candidates

□ « Better-half » strategy which consists of avoiding crosses between the «worst» selected individuals (*Bernardo et al., 2006*) \rightarrow optimization of selection intensity

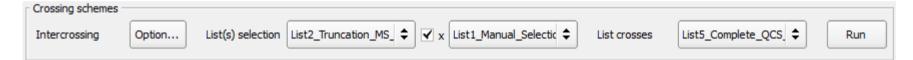


<u>Step3</u>: Identification of crosses to be made among selected individuals

Half-diallel between selected candidates

□ « Better-half » strategy which consists of avoiding crosses between the worst selected individuals (*Bernardo et al., 2006*) \rightarrow optimization of selection intensity

2 lists: factorial design



Constraints:

- Maximum number of crosses
- Contribution of individuals

Crossing so	inemes			-	-	
Number of crosses		🔿 Maximu				
Contributio	n of individuals	🔿 Unlimite	d @	1	*	
Criterion	Score	¢				
Method	Complete	¢				
	_	-				

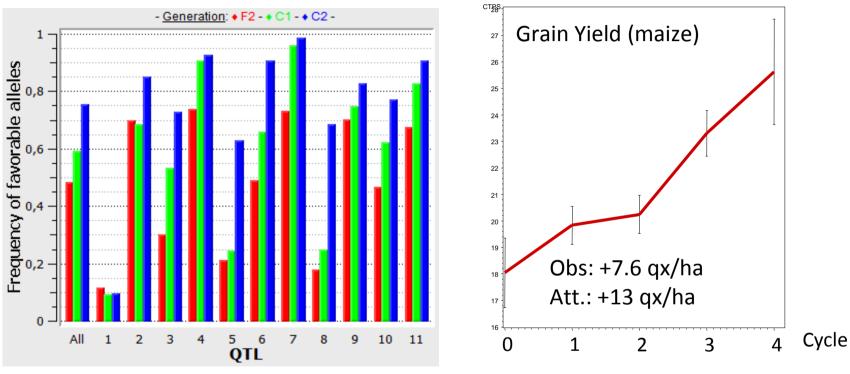
<u>Step3</u>: Identification of crosses to be made among selected individuals

List of crosses	List	1 Complete	Trunc	MS C	rosses				\$	Lis	t5 Complete	ocs	Crosse				
List of Globbes	List1_Complete_Trunc_MS_Crosses						<u> </u>	List5_Complete_QCS_Crosses									
List1_Complete_Trunc_MS_Cross List2_Better_Half_Crosses List3_UC_Crosses		Ind	p1	p2	MS	Weight	UC 🛧	QTL1			Ind	p1	p2	MS	Weight	UC ^	QTL1
		B8xB158	B8	B158	0.8195	0.8195	10,25	0.0000		1	B8xB158	B8	B158	0.8195	0.6935	10,25	0.0000
List4_Truncation_Crosses List5_Complete_QCS_Crosses	2	B8xB28	B8	B28	0.8053	0.8053	10,2071	0.0000		2	B8xB28	B8	B28	0.8053	0.6814	10,2071	0.0000
	3	B8xB13	B8	B13	0.7988	0.7988	10,2071	0.0000		3	B8xB13	B8	B13	0.7988	0.6759	10,2071	0.0000
	4	B8xB38	B8	B38	0.7930	0.7930	10,116	0.0000		4	B8xB38	B8	B38	0.7930	0.6710	10,116	0.0000
	5	B8xB293	B8	B293	0.7817	0.7817	10,116	0.0000		5	B8xB40	B8	B40	0.7885	0.6672	10,116	0.0000
	6	B8xB246	B8	B246	0.7835	0.7835	10,116	0.0000		6	B158xB28	B158	B28	0.7882	0.6670	10,116	0.0000
	7	B8xB242	B8	B242	0.7836	0.7836	10,116	0.0000		7	B8xB242	B8	B242	0.7836	0.6630	10,116	0.0000
Save	8	B158xB28	B158	B28	0.7882	0.7882	10,116	0.0000		8	B158xB13	B158	B13	0.7817	0.6614	10,116	0.0000
Add	9	B158xB13	B158	B13	0.7817	0.7817	10,116	0.0000		9	B8xB124	B8	B124	0.7795	0.6966	10,116	0.2406
Remove	10	B8xB40	B8	B40	0.7885	0.7885	10,116	0.0000		10	B158xB38	B158	B38	0.7759	0.6565	10	0.0000
	11	B28xB13	B28	B13	0.7675	0.7675	10	0.0000	-	11	B158xB40	B158	B40	0.7714	0.6527	10	0.0000
Reset		00.5				15	A	Þ			301				A		Þ

□ In each case: computation of the expected MS of the progeny

Conclusion

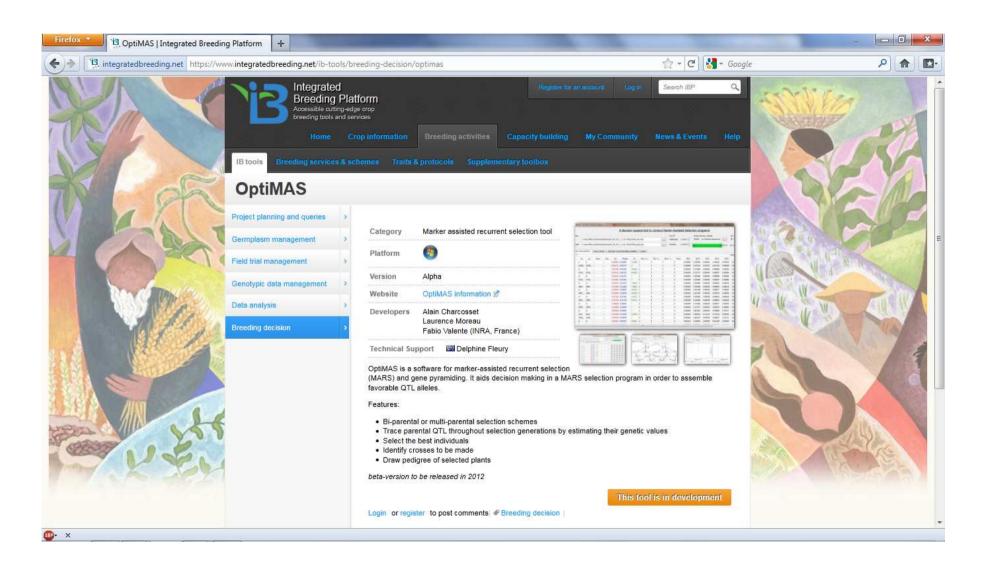
OptiMAS tested on a multiparental connected design (Blanc et al., 2008)



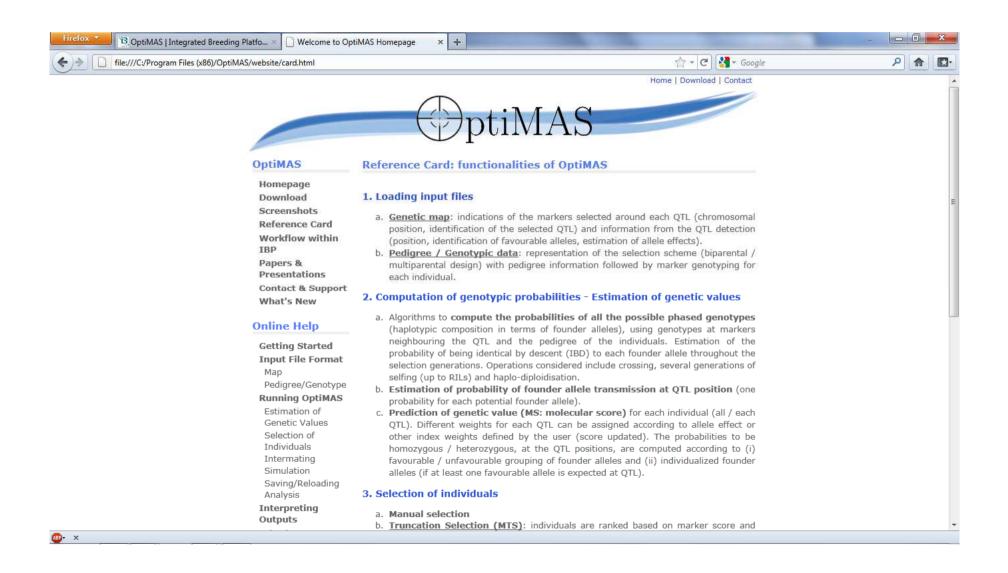
Creation of new genetic material

- Development of 2 version of the tool:
 - Command line: C-ANSI language
 - Graphical Interface (GUI): C++ (Qt & Qwt libraries)
- OptiMAS release: available online soon for Windows & Linux then Mac OSX

Integrated Breeding Platform Website



Tutorial/Documentation



Acknowledgments

- Bill & Melinda Gates Foundation
- Guylaine Blanc & J.B Veyrieras for developing the first prototype of the program
- Nicolas Bardol & Delphine Fleury (GCP) for beta testing



Identification of crosses to be made among selected individuals

- Complete : all possible crosses between the selected individuals.
- Better Half : avoid crosses between the last selected genotypes.
 - → Optimization of selection intensity
- Predefined number of crosses to be made based on :
 - ✓ Molecular score.
 - ✓ Utility criterion [in progress]: expected mean and variance at the next generation (favor heterozygous...).

• Favor couples that will produce the best individuals as possible in the next generation.

$$U_{\text{couple}} = Mean \text{ of the parental} + i \sigma_{\text{couple}}$$

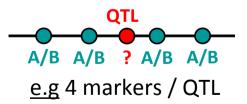
Variance = only heterozygotes genotypes at QTL contribute to variance in the next generation.

Selection of individuals

- Aim: to obtain an ideotype combining all the favorable alleles at QTL positions.
- Prediction of genetic value based on the information from neighboring markers.

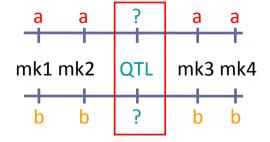
→ Which parental alleles are transmitted at the QTL position ?

□ QTL position rarely located at a marker → QTL alleles are unknown and must be inferred from flanking markers.



Selection of a subset of informative markers to follow the favourable parental alleles (based on haplotypes).

Identification of all the possible phased genotypes for each QTL and computation of their probabilities



Following « phased genotypes »

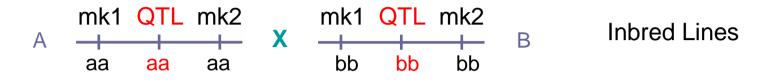
- Information available
 - ✓ Pedigree
 - \checkmark Distance between loci (Haldane \rightarrow recombination rate)
 - Molecular markers (observed genotypes)

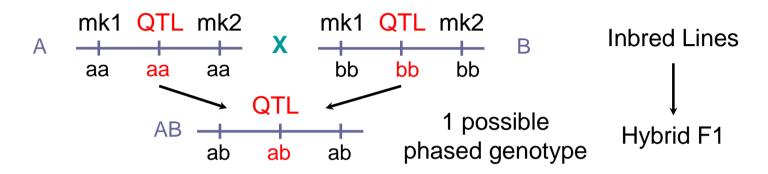
Identification of all the possible phased genotypes for each QTL and computation of their probabilities.

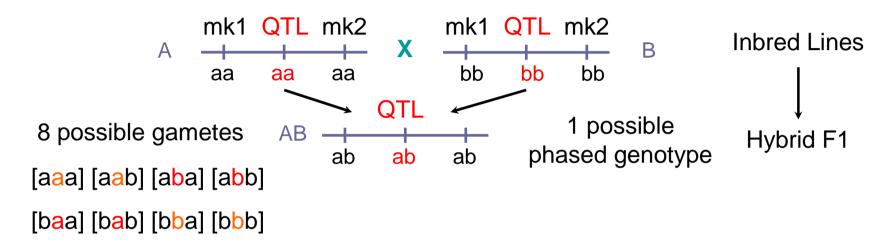


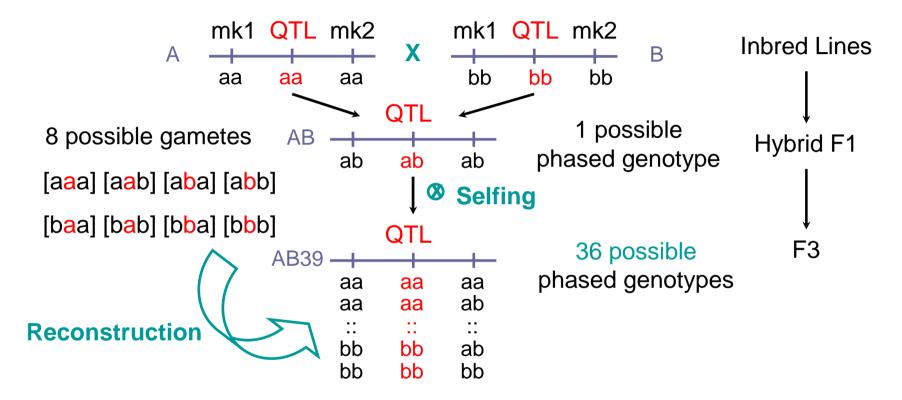
✓ Phase (haplotypes) unknown \rightarrow all possibilities considered

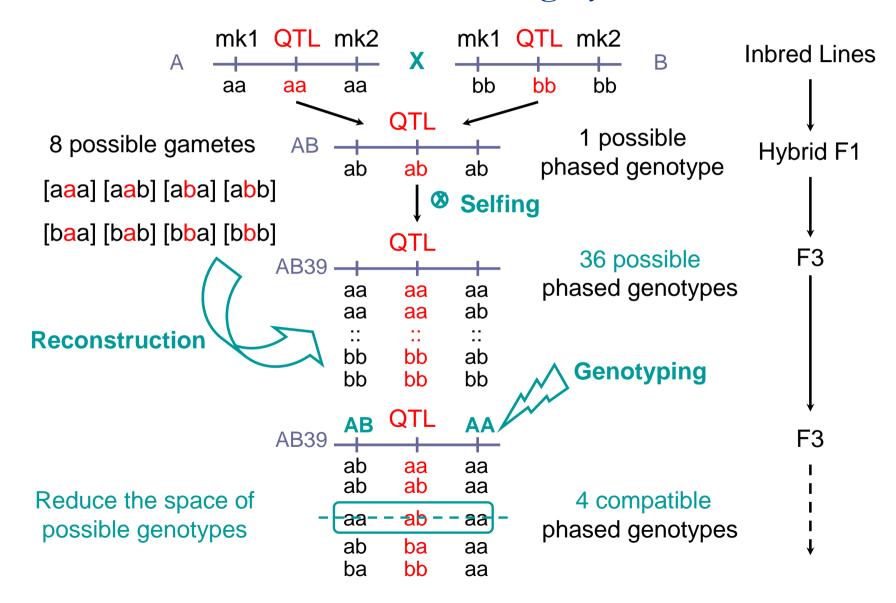
Phased genotype: contains no ambiguity on alleles transmitted from parents and the phase.











Estimation of genetic values

□ _scores: genetic value (molecular score) for each individual

✓ Each QTL: expected dose of favorable allele (0 or 1) for all possible phased genotypes , weighted by the probability of each phased genotype.

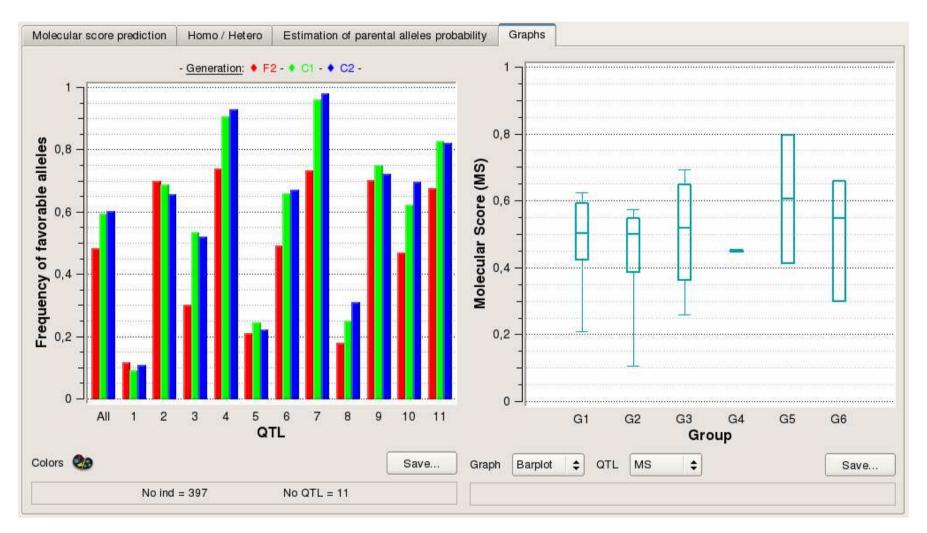
$$\begin{bmatrix}
M_{each} = \frac{\sum \theta_q \cdot p_G}{2} & M_{all} = \frac{\sum M_{each}}{nb_qtl} & 2 \text{ classes:} \\
- Favorable \\
- Not favorable \\
- Not favorable
\end{bmatrix}$$

$$\theta_q : \text{genotype (homo -/+, hetero) at QTL position (0, 1, 2)}$$

$$\theta_G : \text{probability of the current genotype} & Homo - 0 \\
- \text{Option: effect associated with parental allele(s) at QTL}$$

$$= \text{QTL are considered as being independent (unlinked)}$$

<u>Step 1</u>: Computation of genotypic probabilities – Estimation of genetic values (graphs)



□ Mean frequency of favorable alleles at all/each QTL through generation of selection

Intra-group variation (ears)