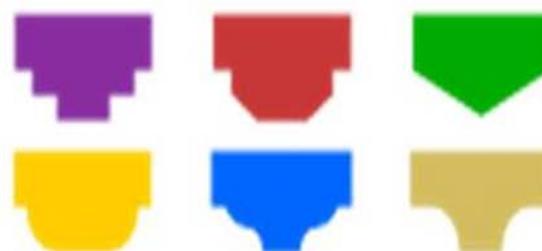


TOOLS BIOINFORMATICI E RISORSE WEB IN ISTOCOMPATIBILITÀ

Donata Mininni
AOU Policlinico - Bari

Antigeni (Ag)

- Tutte le molecole in grado di attivare il sistema immunitario sono detti **antigeni**
- Normalmente gli **Ag** sono **sostanze estranee all'organismo ad alto peso molecolare** quali proteine e lipopolisaccaridi
- Non sono **antigeni molecole a basso peso molecolare**, anche se estranee, come per esempio alcuni farmaci disaccaridi ecc...

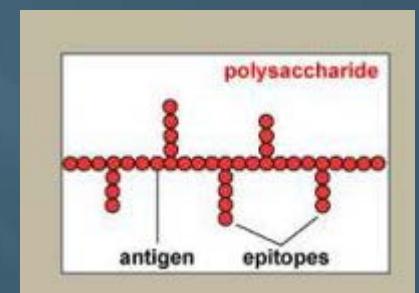
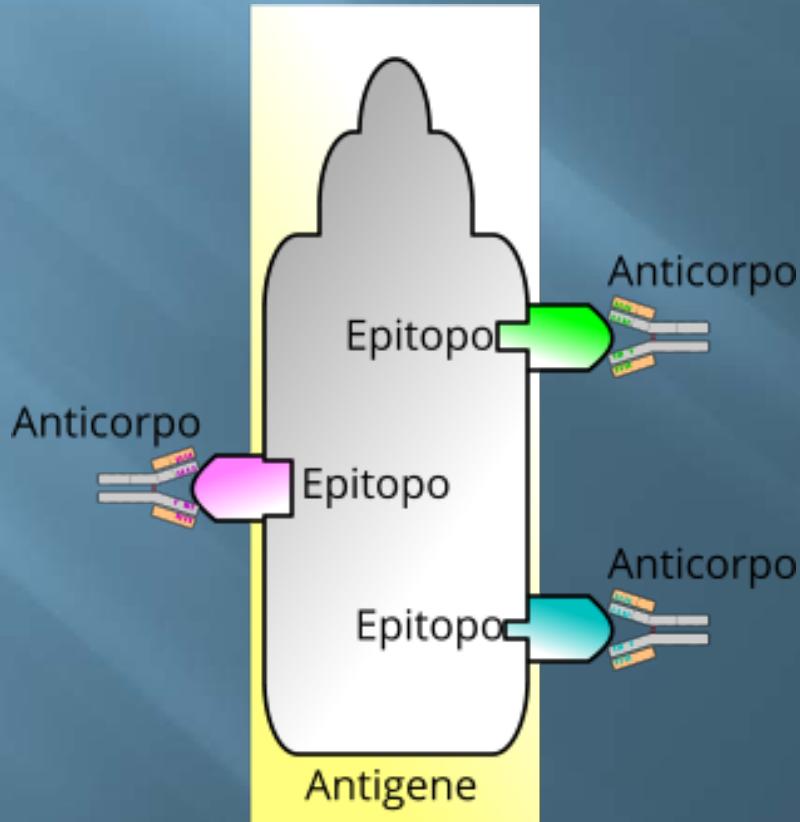


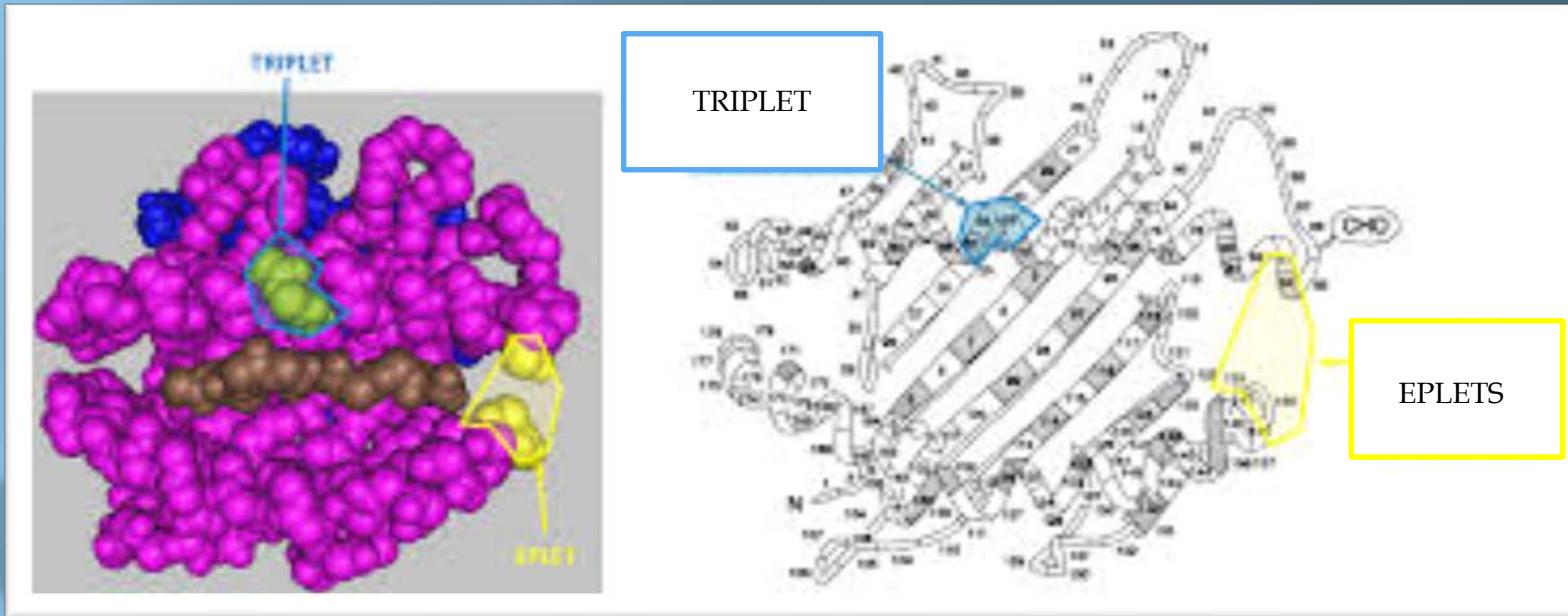
Essi si possono trovare sulla superficie delle cellule oppure si possono trovare liberi (**antigeni circolanti**)

La capacità di un **Ag** di combinarsi con un **Ab** riflette la sua **antigenicità**

EPITOPO

Un **epitopo** o **determinante antigenico** è una frazione dell'antigene capace di scatenare una reazione immunitaria una volta che un anticorpo opera un riconoscimento su di esso. Gli epitopi sono specifici per un solo tipo di anticorpo anche se, in alcuni casi, è possibile la crossreazione tra un epitopo e più tipi di anticorpo.





L'immunogenicità degli antigeni HLA è determinata da brevi sequenze continue e discontinue di aa che formano le regioni accessibili dagli Ab all'interno di ciascun allele HLA noto come EPITOPO.

Ogni allele HLA è considerato come una combinazione di epitopi distinti noti come Triplets (*sequenze amminoacidiche continue*) o Eplets (*sequenze aminoacidiche contigue strettamente localizzate*)

Immunogenicità

Immunogenicità è la capacità di un Ag di indurre una risposta immune

Un Ag può essere **antigenico** ma non **immunogeno**, cioè si lega all'Ab ma non induce la produzione di Ab da parte della cellula B

**Caratteri che condizionano l'immunogenicità di
un antigene**

PESO MOLECOLARE

SOLUBILITÀ

CONFORMAZIONE SPAZIALE

NATURA CHIMICA

DISPONIBILITÀ DEGLI EPITOPI

Nella compatibilità HLA tra don/ric il disallineamento nelle molecole di Classe I e II determina la presenza del Mismatch (MM)

**ESISTE UNA
GERARCHIA DEI MM
ANTIGENICI CHE
PROVOCANO
RIGETTO**



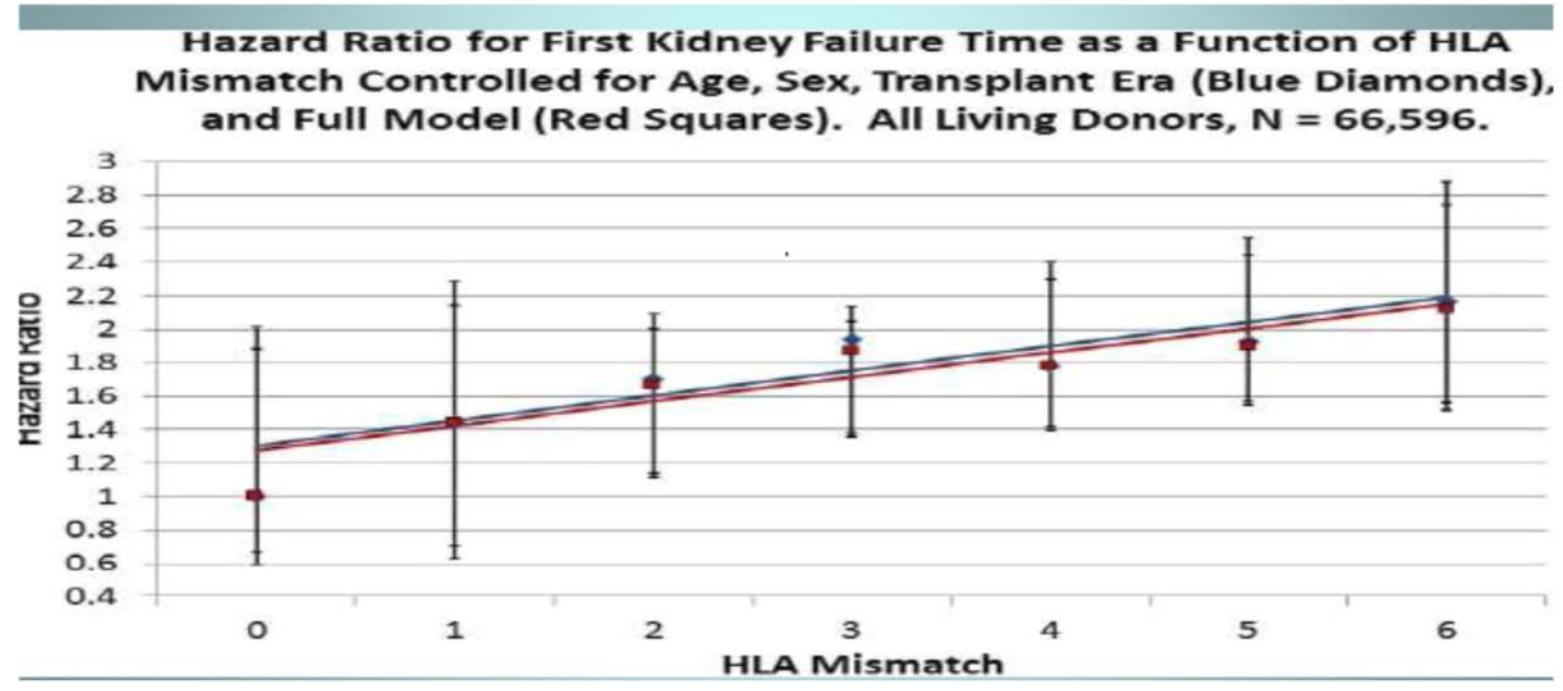
MM accettabile :Antigene HLA che non provoca reazione immunologica energica

MM inaccettabile (immunogenico) : provoca reazione immunologica energica

L'esito di un trapianto d'organo solido dipende anche dalla presenza o assenza di MM

Tx immunogenicity

Williams, Transplant direct, 2017



La forza di immunogenicità puo essere calcolata con
la potenzialità di produrre HLA-Abs

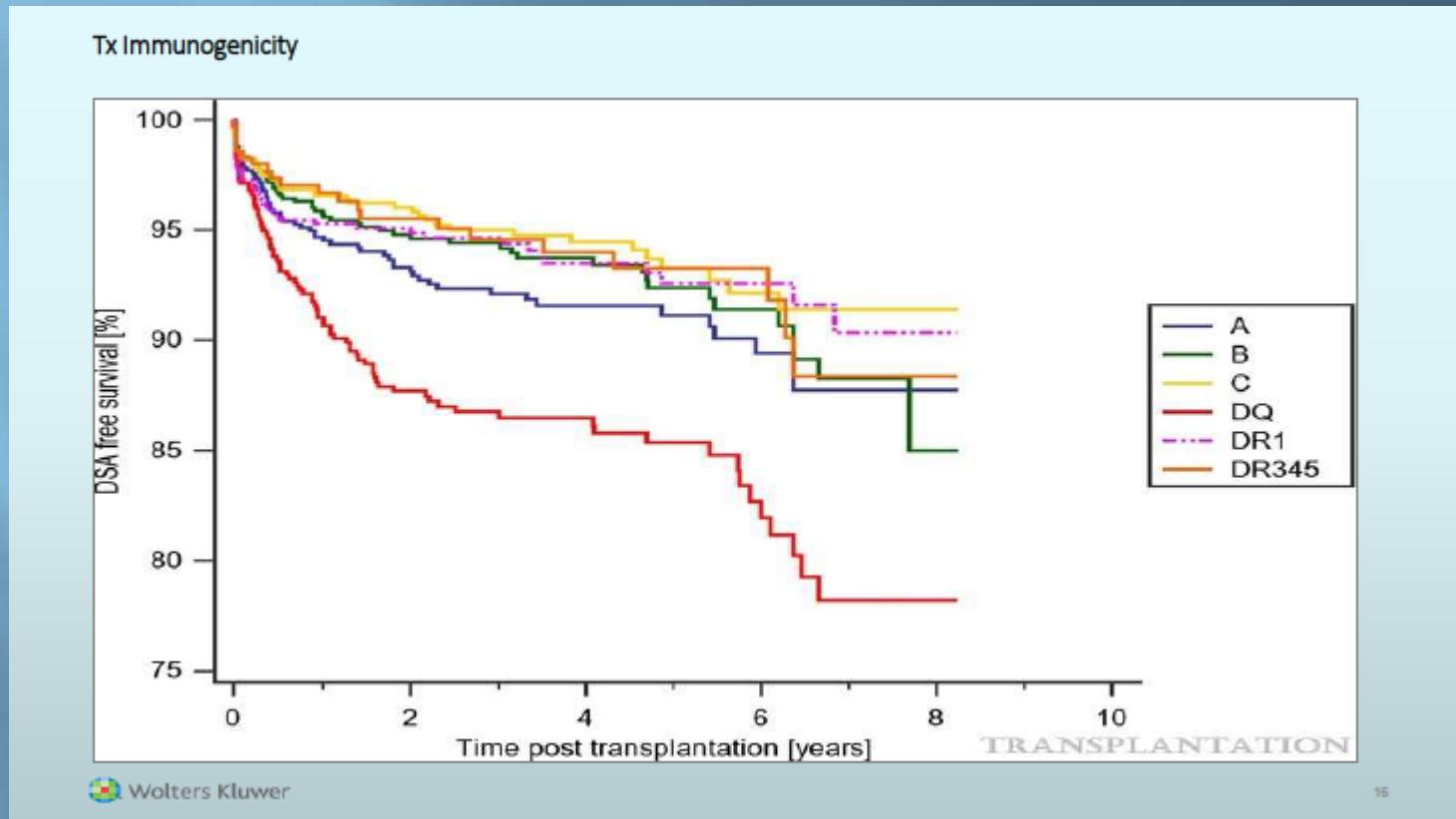


DSA



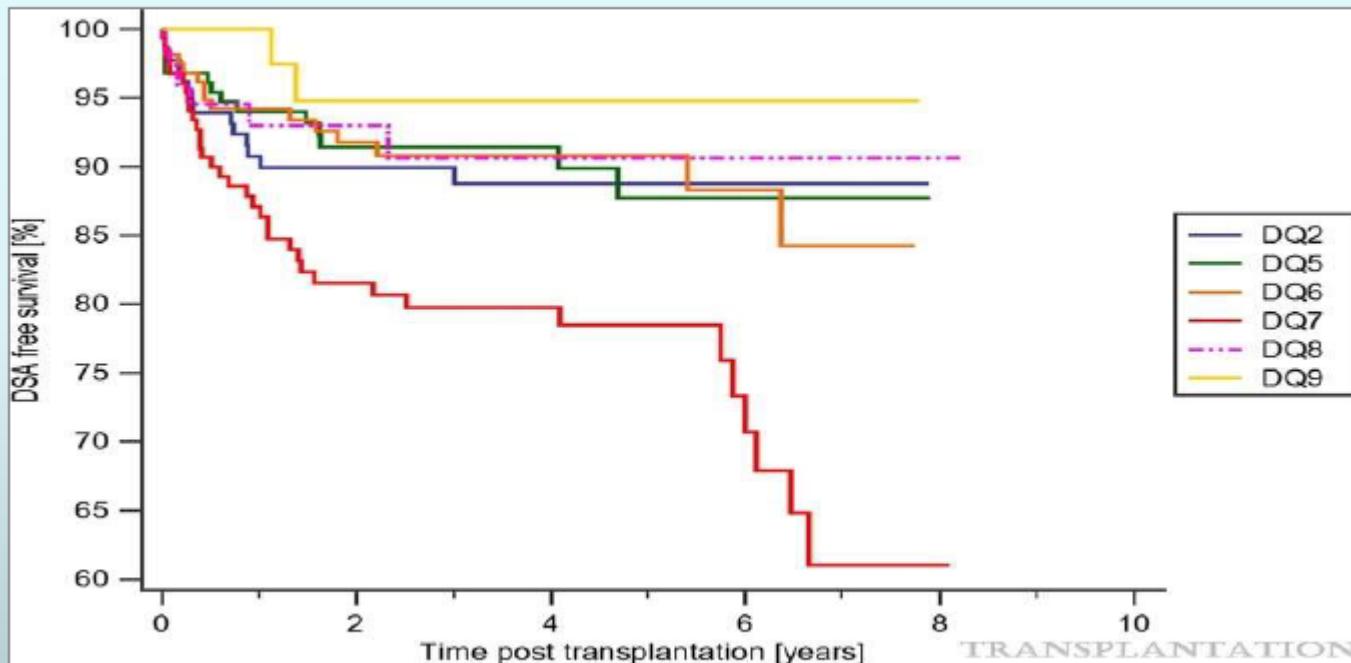
non-DSA

▫ differences in immunogenicity of HLA antigens ?

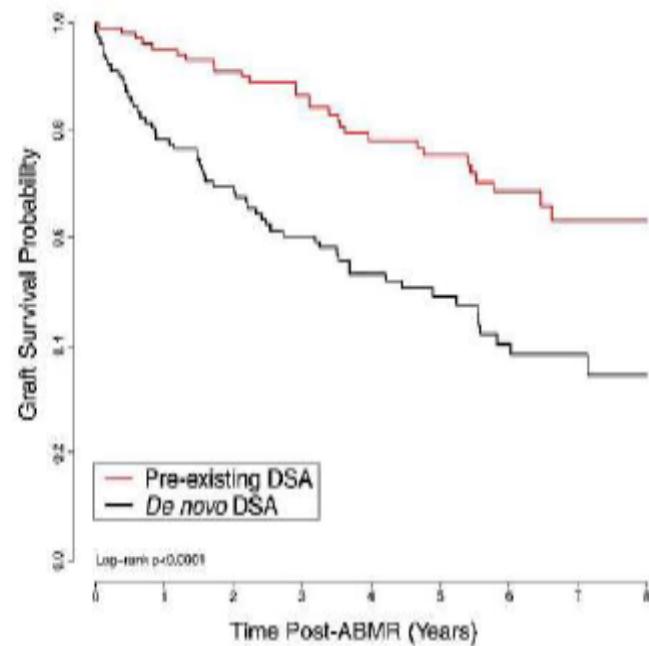


Willicombe M, Transplant,2018

Tx immunogenicity



DSA consideration: inferior graft survival with de novo DSA



N at Risk

| | | | | | | | | | |
|------------------|-----|----|----|----|----|----|----|----|----|
| Pre-existing DSA | 103 | 95 | 87 | 74 | 61 | 49 | 32 | 17 | 11 |
| De novo DSA | 102 | 80 | 70 | 56 | 43 | 31 | 22 | 10 | 4 |

Aubert, JASN 2017

HIERARCHY OF HLA TYPES IN PROVOKING ANTIBODY PRODUCTION FOLLOWING KIDNEY TRANSPLANTATION.

David D. Gae^a, Haibo Sun^a, Owen Buenaventura^a, Kelly J. Cunniffe^a, Gil Da Gente^a, John Roberts^b, Raja Rajalingam^a. ^aImmunogenetics and Transplantation Laboratory, Department of Surgery, University of California San Francisco, San Francisco, CA, United States; ^bTransplant Services, Department of Surgery, University of California San Francisco, San Francisco, CA, United States.

Aim: HLA antigens are diverse in cell surface expression, peptide presentation, binding immune receptors, and in eliciting immune responses. Here we examined the ability of distinct HLA types in inducing antibody (Ab) production following kidney transplantation (Ktx).

Methods: A cohort of 308 candidates (44% female) waiting for 2nd Ktx was studied. Prior to 1st Ktx, all were negative for HLA Abs by One Lambda single antigen bead (OL-SAB) assay. HLA Abs were measured in pre-2nd tx sera using OL-SAB. The strength (mean of MFIs) and frequency of each Abs in pre-2nd tx sera of 308 candidates were determined and plotted.

Results: A positive linear correlation was observed between Ab strengths to frequency with an $r^2 = 0.6$ and $p < 2 \times 10^{-29}$ (see figure). DQ3 is the most frequently and strongly encountered Abs and thus DQ3 is the most aggressive antibody inducer following Ktx. DR53 group of antigens (except DR4) and A2 CREG are the next most aggressive Ab inducers. The hierarchy from the strongest inducer is DQ9 > DQ8 > DQ7 > B76 > DR9 > DQ4 > B45 > DQ6 > B57 > DQ2 > A1 > DQ5 > DR7 > B57 > B58 > A24 > B49 > DR53 > A2 > B82 > B44 > A96 > A23 > A11 > A68. All HLA-Cw antigens and a subset of DPB antigens (DP15, DP13, DP5, DP1, DP23, DP4) induce Ab production at low levels and low frequency. Within Cw types, the most poorly immunogenic represents to be those carrying C1 epitope that binds to KIR2DL2/L3. The moderate inducer includes most HLA-B, HLA-A and DR antigens, and the hierarchy from the strongest inducer within this group is B7 > A10 > B5 > DR52 > DR51 CREG. The B5 CREG was diverse and observed to have four outliers from the linear fit. The strongest to weakest to trigger Abs in B5 CREG is ordered as follows: B76 > B63 > B73 > B62 > B75 > B78 > B46.

Conclusions: Our findings categorize 121 HLA types into three hierarchical sets based on their ability in eliciting Ab production following Ktx – strongly immunogenic, moderately immunogenic, and poorly immunogenic. The strongly immunogenic HLA types should be considered for matching in Ktx, while the poorly immunogenic HLA antigens are tolerable.

- HLA antigen mismatches with highest immunogenicity are called “Taboo” mismatches

- Doxiadis II, lancet,1996;348

- Rene J. Duquesnoy

- Geneugelijk

- J Immunol Res,201

- Slavcev,A

- International J Immunogenetics,2013

- Tx Immunogenicity

- Examples of Taboo Donor/ Recipient combinations

- A2/B44

- A3/DR6

- B7/A1

- B60/DR2

- B60/DR2

- DR5/A2 (worst)

- B44:02/B44:03

RISORSE WEB IN ISTOCOMPATIBILITÀ:

- HLA Matchmaker
- PIRCHE II

[HSCT](#) [SOT](#)[SOT Single Patient](#)[SOT Multi Patient \(CSV\)](#)[SOT Risk Profile](#)[SOT Acceptable Mismatch Profile](#)[Contact](#)[Take the tour](#)

Patient: HLA data is incomplete or contains unknown allele codes.

Population

NMDP EUR haplotypes (2007)

Input Wizard

Paste data here

HLA Data **ID**

dnaser**A***
B*
C*
DRB1*
DQB1*
DPB1*


Donors: Please check HLA data. Allele codes are incomplete or unknown.

Population

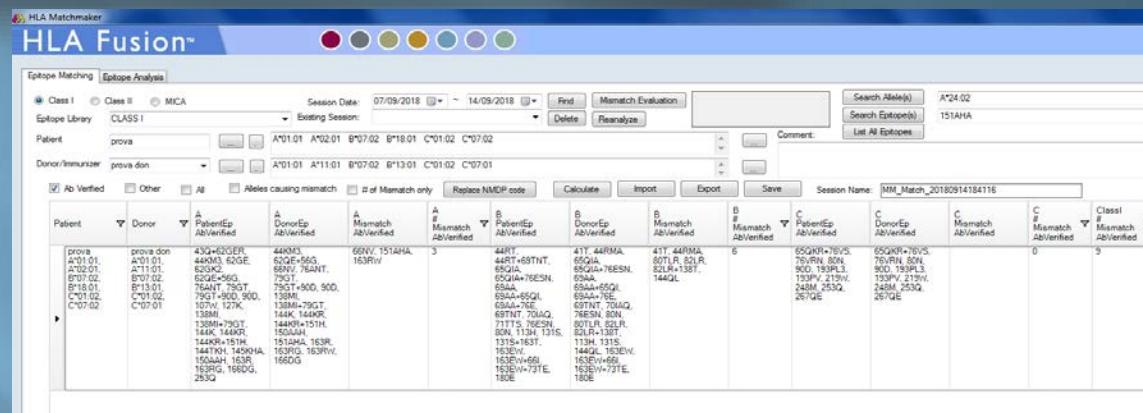
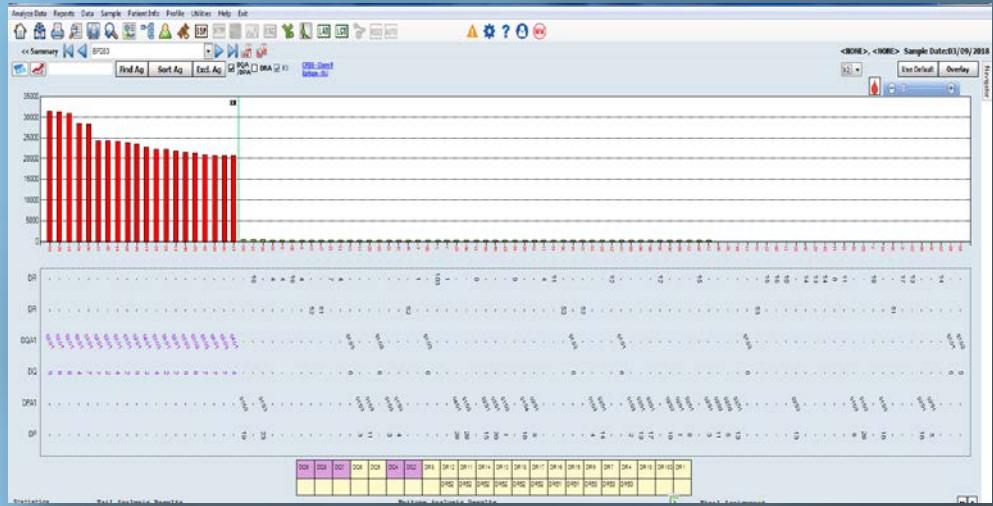
NMDP EUR haplotypes (2007)

Input Wizard

Paste data here

HLA Data **ID**

dnaser**A***
B*
C*
DRB1*
DQB1*
DPB1*
 [Import](#) [Export](#) [Example](#)[Match](#)



- Paziente: donna con GN a depositi mesangiali da IgA
- Stato immunologico pre-TX: PRA pos I e II classe (78% // 68%)
- I° TX : 43aa da donatore cadavere
- I° TX match: A*02, DRB1*07, DRB1*14
- PRA 90% // 97% numerose specificità , NO DSA TX
- Espianto rapido
- Rientro in lista d'attesa
- Stato di Iperimmune → possibilità di offerta donatore da PNI
- Cicli di immunofiltrazione
- II TX : 50 aa da donatore maschio , 31aa
- II° TX match: A*02, B*07, C*15, DRB1*14, DQB1*05
MM: B*15, DRB1*13, DQB1*06

- Presenza di Ab di specificità DPB1* non evidenziabili come DSA
- Post-TX, valutazione Typing donatore per locus DPB1 :
“ DPB1*03:01, DPB1*10:01 ”
- Riscontro delle stesse specificità tra gli A
 - ↓
 - DSA preesistenti
- Formazione di DSA de-Novo : B*15:01, DRB1*13:01, DQB1*06:03 oltre a tante altre specificità con MFI elevato
-

HLA Fusion™

Epitope Matching | Epitope Analysis

Class I Class II MICA

Patient: 12846 DRB1*07:01 DRB1*14:01P DQB1*02:02 DQB1*05:03 DPB1*04:02

Epitope Library: CLASS II 06/08/2017 - 15/09/2018 Find Donor / Immunizer: DON: DRB1*13:01 DRB1*14:01P DQB1*06:03 DQB1*05:03 DPB1*03:01 DPB1*10:01

Not in panel: DRB1*14:01P DQB1*05:03

| Select | Sample ID | Sample Date | Patient | Well Position | Session Name |
|---|--------------|-------------|-----------|-----------------------------------|--------------|
| <input type="checkbox"/> BM757 | set 21, 2017 | 90708 | 44(1,D6) | LS SA II urg270917_20170927_15251 | |
| <input checked="" type="checkbox"/> BM766 | set 21, 2017 | 12846 | 28(1,B4) | LS SA II URG280917_20170929_132 | |
| <input type="checkbox"/> BM768 | set 21, 2017 | 12846 | 28(1,B4) | LS SA II URG280917_20170929_132 | |
| <input type="checkbox"/> BM771 | set 22, 2017 | 90725 | 45(1,E6) | LS SA II urg270917_20170927_15251 | |
| <input type="checkbox"/> BM774 | set 25, 2017 | 1094 | 85(1,E..) | LS SA II urg021017_20171002_14211 | |

Replace NMDP Code: Use Cutoff from LABScreen Analysis!

Data Type: Baseline Cutoff: 1013 1013 1013

Mean of Self (m): 6 18 0 0

SD: 8 0 0 0

m+3SD: 30 18 0 0

Epitope_AbVerified Filter Ep Filter Allele

Show Self Show Imm Show POS Reset Export

Row Count: 6

| Allele | Bead ID | Locus | Raw | Baseline | Ratio | Rxn | Missing EP Def | Cutoff | Self | Imm | Ab Verified | N. Ab Verified | All | N. All | Imm Ab Verified | N. Imm Ab Verified | Imm All | N. Imm All | TP Ab Verified | N. TP Ab Verified | TP All | N. TP All |
|------------|---------|-------|-------|----------|--------|-----|-------------------------------------|--------|--------------------------|-------------------------------------|-------------|----------------|---------------------|--------|-----------------|---------------------|---------------------|------------|----------------|-------------------|--------|-----------|
| DPB1*03:01 | 073 | DP | 12533 | 12450 | 235.98 | POS | <input type="checkbox"/> | 1013 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 84DEAV | 1 | 9YL, 11L, 76V, 8..4 | 84DEAV | 1 | 9YL, 11L, 76V, 8..4 | 9YL, 11L, 76V, 8..4 | 0 | 0 | | | |
| DPB1*03:01 | 072 | DP | 12143 | 12066 | 243.96 | POS | <input type="checkbox"/> | 1013 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 84DEAV | 1 | 9YL, 11L, 76V, 8..4 | 84DEAV | 1 | 9YL, 11L, 76V, 8..4 | 9YL, 11L, 76V, 8..4 | 0 | 0 | | | |
| DPB1*03:01 | 071 | DP | 12041 | 11962 | 236.73 | POS | <input type="checkbox"/> | 1013 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 84DEAV | 1 | 9YL, 11L, 76V, 8..4 | 84DEAV | 1 | 9YL, 11L, 76V, 8..4 | 9YL, 11L, 76V, 8..4 | 0 | 0 | | | |
| DPB1*10:01 | 080 | DP | 4938 | 4843 | 83.72 | POS | <input type="checkbox"/> | 1013 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 84DEAV | 1 | 9H, 11L, 76V, 84..4 | 84DEAV | 1 | 9H, 11L, 76V, 84..4 | 9H, 11L, 76V, 84..4 | 0 | 0 | | | |
| DRB1*13:01 | 022 | DR | 243 | 145 | 4.02 | NEG | <input checked="" type="checkbox"/> | 1013 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |
| DQB1*06:03 | 053 | DQ | 72 | 9 | 1.69 | NEG | <input type="checkbox"/> | 1013 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | 0 | | |

☰ <U.R.P. | "INNO ALLA VITA" 1500 TRAPIANTI DI RENE nella Regione Puglia – Teatro Petruzzelli – 7 luglio 2018 ore 9.00 | Trasmissione del Piano Aziendale per l'incremento della distribuzione diretta del primo ciclo di terapia | Regolar... ▾

HLA Matchmaker

HLA Fusion™

Epitope Matching | Epitope Analysis

Class I Class II MICA

Patient: 12846 A*01:01 A*02:01 B*07:05 B*08:01 C*07:01 C*15:05

Epitope Library: CLASS I 05/08/2017 - 15/09/2018 Find Donor / Immunizer: DON-PARISI: A*02:01 A*02:01 B*07:05 B*15:01 C*05:05 C*07:01

Not in panel: B*07:05 C*07:01 C*15:05

| Select | Sample ID | Sample Date | Patient | Well Position | Session Name |
|---|--------------|-------------|-----------|-----------------------------------|--------------|
| <input type="checkbox"/> BM757 | set 21, 2017 | 90708 | 12(1,D2) | LS SA II urg270917_20170927_15101 | |
| <input checked="" type="checkbox"/> BM766 | set 21, 2017 | 12846 | 10(1,B2) | LS SA II URG280917_20170929_1315 | |
| <input type="checkbox"/> BM768 | set 21, 2017 | 10997 | 48(1,H6) | LSA1_CDOS_N_0617_20180430_13 | |
| <input type="checkbox"/> BM770 | set 21, 2017 | 63741 | 49(1,A..) | LSA1_CDOS_N_0617_20180430_13 | |
| <input type="checkbox"/> BM771 | set 22, 2017 | 90725 | 13(1,E2) | LS SA II urg270917_20170927_15101 | |

Replace NMDP Code: Use Cutoff from LABScreen Analysis!

Data Type: Baseline Cutoff: 1025

Mean of Self (m): 0

SD: 0

m+3SD: 0

Epitope_AbVerified Filter Ep Filter Allele

Show Self Show Imm Show POS Reset Export

Row Count: 2

| Allele | Bead ID | Locus | Raw | Baseline | Ratio | Rxn | Missing EP Def | Cutoff | Self | Imm | All | N. All | Imm All | N. Imm All | TP All | N. TP All |
|---------|---------|-------|-----|----------|-------|-----|--------------------------|--------|-------------------------------------|-------------------------------------|-----|--------|---------|------------|--------|-----------|
| B*15:01 | 040 | B | 132 | 77 | 3.13 | NEG | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 0 | 0 | 0 | 0 | 0 | |
| A*02:01 | 004 | A | 20 | 0 | 0.16 | NEG | <input type="checkbox"/> | 1025 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | 0 | 0 | 0 | 0 | 0 | |

HLA Fusion™

Epitope Matching Epitope Analysis

Class I Class II MICA

Epitope Library CLASS I 06/08/2017 15/09/2018 Find Donor / Immunizer Patient 12846 A*01:01 A*02:01 B*07:05 B*08:01 C*07:01 C*15:05

| Select | Sample ID | Sample Date | Patient | Well Position | Session Name |
|-------------------------------------|-----------|--------------|---------|---------------|----------------------------------|
| <input type="checkbox"/> | BM751 | set 20, 2017 | 90089 | 10(1,02) | LS SA1 urg270917_20170927_15101 |
| <input type="checkbox"/> | BM758 | set 21, 2017 | 90193 | 11(1,C2) | LS SA1 urg270917_20170927_15101 |
| <input type="checkbox"/> | BM757 | set 21, 2017 | 90708 | 12(1,02) | LS SA1 urg270917_20170927_15101 |
| <input checked="" type="checkbox"/> | BM766 | set 21, 2017 | 12846 | 10(1,02) | LS SA1 URG280917_20170929_131517 |
| <input type="checkbox"/> | BM768 | set 21, 2017 | 10997 | 48(1,16) | LSA1_CD05_N_0617_20180430_13 |

Use Cutoff from LABScreen Analysis !

Data Type Baseline Cutoff 1025
Mean of Self (m) 0
SD 0
m+3SD 0

Epitope_AbVerified Filter Ep B*15 Filter Allele
Show Self Show Imm Show POS Reset Export

Row Count: 8

| Allele | Bead ID | Locus | Raw | Baseline | Ratio | Rxn | Missing EP Def | Cutoff | Self | Imm | Ab Verified | N. Ab Verified | Imm Ab Verified | N. Imm Ab Verified | TP Ab Verified | N. TP Ab Verified |
|---------|---------|-------|-------|----------|--------|-----|-------------------------------------|--------|-------------------------------------|-------------------------------------|-------------------------------------|-----------------|-----------------|--------------------|-----------------|-------------------|
| B*15:15 | 046 | B | 10602 | 10278 | 52.56 | POS | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 71SA, 80I, 82LR | 5 | 0 | 71SA, 80I, 82LR | 5 |
| B*15:13 | 045 | B | 8809 | 8837 | 172.01 | POS | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 80I, 80I+69TNT | 6 | 0 | 80I, 80I+69TNT | 6 |
| B*15:12 | 044 | B | 18650 | 1668 | 15.03 | POS | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 163L/S/G | 1 | 0 | 163L/S/G | 1 |
| B*15:03 | 042 | B | 155 | 83 | 2.96 | NEG | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 0 | 0 | 0 | 0 | 0 |
| B*15:01 | 040 | B | 132 | 77 | 3.13 | NEG | <input checked="" type="checkbox"/> | 1025 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input type="checkbox"/> | 0 | 0 | 0 | 0 | 0 |
| B*15:10 | 043 | B | 136 | 71 | 2.85 | NEG | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 0 | 0 | 0 | 0 | 0 |
| B*15:11 | 098 | B | 99 | 21 | 4.78 | NEG | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 0 | 0 | 0 | 0 | 0 |
| B*15:02 | 041 | B | 111 | 6 | 1.55 | NEG | <input type="checkbox"/> | 1025 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 0 | 0 | 0 | 0 | 0 |

HLA Fusion™

Epitope Matching Epitope Analysis

Class I Class II MICA

Epitope Library CLASS I 06/08/2017 15/09/2018 Find Donor / Immunizer Patient 12846 A*01:01 A*02:01 B*07:05 B*08:01 C*07:01 C*15:05

| Select | Sample ID | Sample Date | Patient | Well Position | Session Name |
|-------------------------------------|-----------|--------------|---------|---------------|---------------------------------|
| <input type="checkbox"/> | BM887 | ott 05, 2017 | 89293 | 68(1,09) | LS SA1 urg061017_20171006_13104 |
| <input type="checkbox"/> | BM888 | ott 05, 2017 | 71671 | 69(1,09) | LS SA1 urg061017_20171006_13104 |
| <input type="checkbox"/> | BM892 | ott 05, 2017 | 12846 | 70(1,F9) | LS SA1 urg061017_20171006_13104 |
| <input checked="" type="checkbox"/> | BM892 | ott 05, 2017 | 12846 | 12(1,02) | LS A1 urg121017_20171012_12235 |
| <input type="checkbox"/> | BM893 | ott 05, 2017 | 89472 | 51(1,07) | LSA1_CD05_N_0617_20180430_13 |

Use Cutoff from LABScreen Analysis !

Data Type Baseline Cutoff 1140
Mean of Self (m) 0
SD 0
m+3SD 0

Epitope_AbVerified Filter Ep B*15 Filter Allele
Show Self Show Imm Show POS Reset Export

Row Count: 8

| Allele | Bead ID | Locus | Raw | Baseline | Ratio | Rxn | Missing EP Def | Cutoff | Self | Imm | Ab Verified | N. Ab Verified | Imm Ab Verified | N. Imm Ab Verified | TP Ab Verified | N. TP Ab Verified | |
|---------|---------|-------|-------|----------|--------|-----|-------------------------------------|--------|-------------------------------------|-------------------------------------|-------------------------------------|-------------------|-----------------|--------------------|----------------|-------------------|---|
| B*15:12 | 044 | B | 20668 | 20471 | 81,81 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input checked="" type="checkbox"/> | 44RMA, 131S, 1..3 | 3 | 44RMA, 131S | 2 | 163L/S/G | 1 |
| B*15:01 | 040 | B | 13146 | 13077 | 165,8 | POS | <input checked="" type="checkbox"/> | 1140 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | 44RMA, 131S, 1..3 | 3 | 44RMA, 131S, 1..3 | 3 | 0 | 0 |
| B*15:03 | 042 | B | 12752 | 12663 | 120,56 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 131S, 163L/W-6..2 | 2 | 131S, 163L/W-6..2 | 2 | 0 | 0 |
| B*15:10 | 043 | B | 10509 | 10425 | 105,49 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 131S, 163L/W-6..2 | 2 | 131S, 163L/W-6..2 | 2 | 0 | 0 |
| B*15:02 | 041 | B | 9402 | 9289 | 68 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 44RMA, 131S, 1..3 | 3 | 44RMA, 131S, 1..3 | 3 | 0 | 0 |
| B*15:13 | 045 | B | 6965 | 6883 | 72,7 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 44RMA, 80I, 80L | 9 | 44RMA, 131S, 1..3 | 3 | 80I, 80I+69TNT | 6 |
| B*15:16 | 046 | B | 7068 | 6768 | 18,06 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 44RMA, 71SA, 8..7 | 7 | 44RMA, 131S | 2 | 71SA, 80I, 82LR | 5 |
| B*15:11 | 098 | B | 5842 | 5749 | 53,02 | POS | <input type="checkbox"/> | 1140 | <input type="checkbox"/> | <input type="checkbox"/> | <input type="checkbox"/> | 44RMA, 131S, 1..3 | 3 | 44RMA, 131S, 1..3 | 3 | 0 | 0 |

PIRCHE®

/ PIRCHE Matching Service / SOT Single Patient

HSCT SOT

SOT Single Patient

SOT Multi Patient (CSV)

SOT Risk Profile

SOT Acceptable Mismatch Profile

Contact

Take the tour

Population NMDP EUR haplotypes (2007)

Input Wizard Paste data here

HLA Data

| ID | A* | B* | C* | DRB1* | DQB1* | DPB1* |
|-----|----------------|----------------|----------------|----------------|----------------|----------------|
| paz | 01:01 02:01 | 07:05 08:01 | 07:01 15:05 | 07:01 14:01 | 02 05:03 | 04:02 04:02 |
| don | 02:01 02:01 | 07:05 15:01 | 07:01 15:05 | 13:01 14:01 | 05:03 06:03 | 03:01 10:01 |

Donors: Please check extracted HLA values for correctness before matching.

Population NMDP EUR haplotypes (2007)

Input Wizard Paste data here

HLA Data

| ID | A* | B* | C* | DRB1* | DQB1* | DPB1* |
|-----|----------------|----------------|----------------|----------------|----------------|----------------|
| paz | 01:01 02:01 | 07:05 08:01 | 07:01 15:05 | 13:01 14:01 | 02 05:03 | 03:01 10:01 |
| don | 02:01 02:01 | 07:05 15:01 | 07:01 15:05 | 13:01 14:01 | 05:03 06:03 | 03:01 10:01 |

Import Export Example Match

PIRCHE® organ transplant report for patient ID: paz

Created by mininnidonata@gmail.com

| Patient / Donor ID | A* | B* | C* | DRB1* | DQB1* | DPB1* | PIRCHE II |
|--------------------|----------------|----------------|----------------|----------------|----------------|----------------|-----------|
| paz | 01:01 02:01 | 07:05 08:01 | 07:01 15:05 | 07:01 14:01 | 02 05:03 | 04:02 04:02 | |
| don | 02:01 02:01 | 07:05 15:01 | 07:01 15:05 | 13:01 14:01 | 05:03 06:03 | 03:01 10:01 | 53.00 |



ent Result

Filter donor id ×

| Patient / Donor ID | A* | B* | C* | DRB1* | DQB1* | DPB1* | PIRCHE II |
|--------------------|------------------------|-------------------------|------------------------|-------------------------------------|-------------------------|-------------------------|----------------|
| paz: | 01:01 02:01 | 07:05 08:01 | 07:01 15:05 | 07:01 14:01 | 02 05:03 | 04:02 04:02 | |
| don: | 02:01 02:01 0:00 | 07:05 15:01 18:00 | 07:01 15:05 0:00 | 13:01 14:01 1:00 | 05:03 06:03 19:00 | 03:01 10:01 15:00 | 53.00 53.00 |
| HLA ID | Presenting Allele | Presented Allele | Core Sequence | Peptide | IC 50 | # | |
| HLA00719 | DRB1*07:01 29:00 | DPB1*03:01 9:00 | ICQVEHTSL VYQLRQEYC | GDVYICQVEHTSLDS ENYYYQLRQEYCYAFN | 522.75 602.38 | 100% 100% | |
| | | CYAFNGTQR | YQLRQEYCYAFNGTQR | 997.27 | 100% | | |

- Paziente maschio con nefropatia
- Età al trapianto da vivente : 49aa
- Donatore : donna 48 aa
- Stato immunologico pre-TX : PRA I Classe Neg
PRA II Classe 5% (Ab anti-DQB1*06) no DSA
- Match TX : DRB1*11:01, DQB1*03:01
- PRA post-TX (1 mese) : I Classe neg
II Classe 14%
- Ab II Classe : DQB1*05:01 (MFI 1100) de novo-DSA
DQA1*01:03 (MFI 2500) non-DSA



Patient: Please check extracted HLA values for correctness before matching.

Population NMDP EUR haplotypes (2007)

Input Wizard Paste data here

HLA Data

| ID | A* | B* | C* | DRB1* | DQB1* | DPB1* |
|---------|-------|-------|-------|-------|-------|-------|
| paz | 01:01 | 27:05 | 02:02 | 11:01 | 03:01 | |
| dna/ser | 26:01 | 40:02 | 06:02 | 11:04 | 03:01 | |

Donors: Please check extracted HLA values for correctness before matching.

Population NMDP EUR haplotypes (2007)

Input Wizard Paste data here

HLA Data

| ID | A* | B* | C* | DRB1* | DQB1* | DPB1* |
|---------|-------|-------|-------|-------|-------|-------|
| don | 03:01 | 14:02 | 07:01 | 01:02 | 03:01 | |
| dna/ser | 68:01 | 49:01 | 08:02 | 11:01 | 05:01 | |

Actions: Import, Export, Example, Match

Patient Result

Filter donor id

Patient / Donor ID

| | A* | B* | C* | DRB1* | DQB1* | DPB1* | PIRCHE II |
|-----|----------------|----------------|----------------|----------------|----------------|-------|-----------|
| paz | 01:01 26:01 | 27:05 40:02 | 02:02 06:02 | 11:01 11:04 | 03:01 03:01 | | |
| don | 03:01 68:01 | 14:02 49:01 | 07:01 08:02 | 01:02 11:01 | 03:01 05:01 | | 191.00 |

The screenshot shows the HLA Matchmaker software interface. At the top, there are tabs for 'Epitope Matching' and 'Epitope Analysis'. Below the tabs, there are buttons for 'Class I', 'Class II', and 'MICA'. The 'Epitope Library' dropdown is set to 'CLASS II'. The 'Session Date' is set to '08/09/2018' to '15/09/2018'. There are buttons for 'Find', 'Mismatch Evaluation', 'Delete', and 'Reanalyze'. On the right, there are buttons for 'Search Allele(s)', 'Search Epitope(s)', and 'List All Epitopes'. A red circle highlights the 'Search Allele(s)' button, which contains the text 'DQB1*06_03'. The main area shows a table with columns: 'Epitope', 'Exposed', 'Antibody Reactive', 'Antibody Verified', 'Allele Matched', and 'Alleles'. The table lists numerous entries, each with a unique identifier (e.g., 3S, 9Y, 13GM, 23R, 26L, 30H, 37YA, 45G, 45GV, 46VY3, 52PQ2, 52PR, 55RPD, 56P, 56PD, 66EV, 67VG, 67VT, 70GT, 74EL, 77T, 85VA, 87F, 116V, 125G, 130R, 135D, 140A2, 167R, 185T) and their corresponding alleles.

HLA Fusion™

Epitope Matching Epitope Analysis

Class I Class II MICA

Epitope Library CLASS II 31/05/2018 15/09/2018 Find Donor / Immunizer donatore SD DRB1*11:01 DRB1*11:04 DOB1*03:01 DQB1*03:01 Not in panel

| Select | Sample ID | Sample Date | Patient | Well Position | Session Name |
|-------------------------------------|-----------|--------------|---------|---------------|---------------------------------|
| <input checked="" type="checkbox"/> | B0614 | mag 21, 2018 | 85676 | 79(1.G..) | LSA2_URG_180618_20180618_1716 |
| <input type="checkbox"/> | B0615 | mag 21, 2018 | 88032 | 80(1.H..) | LSA2_URG_180618_20180618_1716 |
| <input type="checkbox"/> | B0617 | mag 21, 2018 | 1601 | 30(1.F4) | ls2_urg_270618_20180627_131954 |
| <input type="checkbox"/> | B0629 | mag 25, 2018 | 12761 | 70(1.F9) | LSA2_CDPOS_0618_AB_20180817_ |
| <input type="checkbox"/> | B0651 | mag 24, 2018 | 1268 | 40(1.H5) | LSA2_PNI_08062018_20180808_1205 |

Replace NMDP Code Use Cutoff from LABScreen Analysis ! DR DQ DP

Data Type Baseline Cutoff 1016 1016 1016

Mean of Self (n) 0 0 0

SD 0 0 0

m+3SD 0 0 0

Epitope_AbVerified Filter Ep Filter Allele

Show Self Show Imm Show POS Reset Export Row Count: 8

| Allele | Bead ID | Locus | Raw | Baseline | Ratio | Rxn | Missing EP Def | Cutoff | Self | Imm | Ab Verified | N Ab Verified | All | N. All | Imm Ab Verified | N. Imm All | Imm All | N. Imm All | TP Ab Verified | N. TP Ab Verified | TP All | N. TP All |
|------------|---------|-------|------|----------|-------|-----|----------------|--------|-------------------------------------|-------------------------------------|-------------------------------------|---------------|-----|--------|-----------------|------------|---------|------------|----------------|-------------------|--------|-----------|
| DQB1*06:01 | 048 | DQ | 1096 | 1016 | 11.73 | POS | | 1016 | | | <input checked="" type="checkbox"/> | 125SQ | 1 | 125SQ | 1 | 125SQ | 1 | 125SQ | 1 | 0 | 0 | |
| | 003 | DR | 53 | 0 | 0.56 | NEG | | 1016 | | | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |
| DRB1*11:01 | 018 | DR | 86 | 0 | 0.66 | NEG | | 1016 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |
| DQB1*02:01 | 066 | DQ | 126 | 0 | 0.66 | NEG | | 1016 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |
| DQB1*03:01 | 057 | DQ | 93 | 0 | 0.67 | NEG | | 1016 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |
| DQB1*03:01 | 068 | DQ | 48 | 0 | 0.49 | NEG | | 1016 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |
| DQB1*03:01 | 059 | DQ | 44 | 0 | 0.48 | NEG | | 1016 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |
| DQB1*03:01 | 060 | DQ | 41 | 0 | 0.44 | NEG | | 1016 | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | <input checked="" type="checkbox"/> | | 0 | | 0 | | 0 | | 0 | 0 | 0 | |

INTERPRETAZIONE DEL NUMERO DI PIRCHE :



| N° PIRCHE |
|-----------|
| 200 |
| 76 |
| 65 |
| 53 |
| 66 |
| 58 |
| 83 |
| 52 |
| 81 |
| 55 |
| 90 |
| 77 |
| 29 |
| 41 |
| 43 |
| 33 |
| 191 |
| 34 |
| 34 |
| 75 |
| 121 |
| 138 |
| 0 |

