

# Calling All Super Heroes

## Search Strategies for Efficient Donor Identification

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# Disclosures

The following faculty and planning committee staff have the following financial disclosures:

Name	Institution	Disclosure
Kelly Lazration	NMDP/Be The Match	None
Janelle Olson	NMDP/Be The Match	None
Kelly Buck	NMDP/Be The Match	None
Jane Kempenich	NMDP/Be The Match	None
Bernadette Anton	NMDP/Be The Match	None
Robin Gach	Mayo Clinic	None

# Learning Objectives

At the conclusion of this session, attendees will be able to:

- ✓ Integrate HLA/search strategy tools and resources into a comprehensive roadmap for navigating patient searches,
- ✓ Evaluate patient searches for more efficient cell source identification, and
- ✓ Determine how HapLogic matching predictions can be optimally used to identify HLA matched donors and cords.



HLA typing  
haplotypes  
disease stage  
7/8  
search assistance funds  
antibodies  
SSA  
10/10  
match  
BMDW  
HaploStats  
donors  
insurance  
HR  
turnaround time  
high resolution  
CT  
alleles  
nonpermissive  
cord blood units  
transplant  
Form 117  
FastTrack  
8/8  
DPB1  
mismatch  
TNC  
traxis  
Coop  
allele reveal  
workup  
Form 22  
permissive  
CD34  
9/10  
donor center



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## TYPE PATIENT

**Consider:**

High Resolution HLA Typing,  
Antibody Testing, DPB1 Typing



- ✓ High resolution HLA typing
- ✓ Antibody testing
- ✓ DPB1 typing

- ✓ Search prognosis tool
- ✓ Related Donor Services

# Search Prognosis Tool

- ✓ HLA genotype frequency (GF) gives a measure of the commonality of a patient's HLA type
- ✓ An NMDP study demonstrated that GF can be predictive of search prognosis, or the ability to identify a 10/10 and/or 9/10 URD in the Be The Match Registry

*Wadsworth, K. et. al. Bone Marrow Transplant 2016 Nov;51(11):1476-81*



*Grab your cape.*

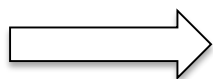


# Genotype Frequency as a Search Prognosis Tool

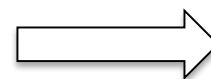
## Patient HLA Type

HLA-A	HLA-B	HLA-C	HLA-DRB1	HLA-DQB1
11:01	18:01	07:01	01:01	05:01
68:02	35:01	04:01	14:01	05:03

+ Patient Ethnicity



Genotype  
Frequency  
Tool



Good

- $\geq 3$  10/10 donors

Fair

- 1-2 10/10s or
- No 10/10s and  $\geq 3$  9/10 donors

Poor

- No 10/10s and  $< 3$  9/10 donors



# Online Search Prognosis Tool

## Patient HLA Typing:

### HLA Table Input

Example: 01:02 or 07:CGNF

[See example typing.](#)

HLA-A	<input type="text" value="11:01"/>	<input type="text" value="68:02"/>
HLA-B	<input type="text" value="18:01"/>	<input type="text" value="35:01"/>
HLA-C	<input type="text" value="04:01"/>	<input type="text" value="07:01"/>
HLA-DRB1	<input type="text" value="01:01"/>	<input type="text" value="14:01"/>
HLA-DQB1	<input type="text" value="05:01"/>	<input type="text" value="05:03"/>

### Self Identified Race and Ethnicity

SIRE (Self Identified Race and Ethnicity)

Self Reported Race is based on the input selection.

White

### Genotype Frequency

Genotype Frequency Information

This frequency is calculated for the most probable haplotype pair based on imputation of the typing.

6.669e-08

### SIRE Search Prognosis

Search Prognosis Information

Using the Patient's Genotype and SIRE, we calculate the frequency and use a model to predict how this patient would fare during the search process.

Fair



# Benefits of Search Prognosis Tool

- ✓ Allows physicians/TC/HLA labs to get an early indication of likely URD search outcome
- ✓ Helps decide when to engage HLA expertise earlier in the search process
- ✓ Guides early clinical consideration of non-fully matched URD alternative stem cell source options

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## ENTER TYPING & VIEW RESULTS

**Verify:**  
HLA Typing Entry & With Lab, If Applicable



*C-B & DRB1-DQB1 Associations  
Allele & Haplotype Frequencies  
Haplostats & IMGT*

- ✓ Verify HLA typing entry
  - Confirm with laboratory?
- ✓ Transplant timeframe
- ✓ View potential donor list in Traxis

- ✓ HLA Analysis Tools
  - B-C, DRB1-DQB1 associations
  - Allele and haplotype frequencies
  - HaploStats and IMGT

# Patient HLA Entry in Traxis: Case Study example 1

Initial Patient HLA Entry in Traxis:

A	B	C	DRB1	DQB1
01:01	08:01	07:01	03:01	02:01
03:01	45:01	08:02	13:02	06:09

Potential 10/10 donor list:

A	B	C	DRB1	DQB1
P P 99	P P 99	1	P P 99	A P 88
P P 99	P P 99	1	P P 99	86
P P 99	P P 99	1	P+ P+ 99	86

Corrected Patient HLA Entry in Traxis:

A	B	C	DRB1	DQB1
01:01	08:01	07:01	03:01	02:01
03:01	45:01	06:02	13:02	06:09

Potential 10/10 donor list:

A	B	C	DRB1	DQB1
A A 99	A A 99	A A 99	A A 99	A A 99
P P 99	P P 99	P P 99	A A 99	P A 99
P P 99	P P 99	P P 99	A A 99	A A 99

# Patient HLA Entry in Traxis: Case Study example 2

Initial Patient HLA Entry in Traxis:

# Likely 10/10s

A	B	C	DRB1	DQB1
02:01 29:02	07:02 58:01	07:19 07:27	08:04 15:01	04:02 06:02

0

**Corrected** Patient HLA Entry in Traxis:

# Likely 10/10s

A	B	C	DRB1	DQB1
02:01 29:02	07:02 58:01	07:01 07:02	08:04 15:01	04:02 06:02

221

Locus	IMGT/HLA 3.9.0 Allele Name	CWD 2.0.0 Category
C	C*07:19	WD
C	C*07:27:01	WD

## IMGT Ambiguous Allele Combinations

*Ambiguous typing combinations over exon 2 + 3*

Combination 1

C\*07:01:01G+C\*07:02:01G

Combination 2

C\*07:19+C\*07:27:01



# Patient HLA Entry in Traxis: Case Study example 3

Initial Patient HLA Entry in Traxis:

A	B	C	DRB1	DQB1	# Likely 10/10s	HLA Typing				
01:01	08:01	07:01	03:01	02:01	0	Uncommon B/C association (08:01-06:02) AND homozygous B				
68:01		06:02	10:01	05:01		A	B	C	DRB1	DQB1
						P P 99	P P 99	1	P A 99	  99

**Corrected** Patient HLA Entry in Traxis:

A	B	C	DRB1	DQB1	# Likely 10/10s	HLA Typing					Outcome
01:01	08:01	07:01	03:01	02:01	130	Common B/C association (37:01-06:02)					10/10 Transplant
68:01	37:01	06:02	10:01	05:01		A	B	C	DRB1	DQB1	
						A A 99	A A 99	A A 99	A A 99	A A 99	



Patient transplanted with 10/10 donor

Grab your cape.



## 2

## ENTER TYPING & VIEW RESULTS

**Verify:**  
HLA Typing Entry & With Lab, If Applicable



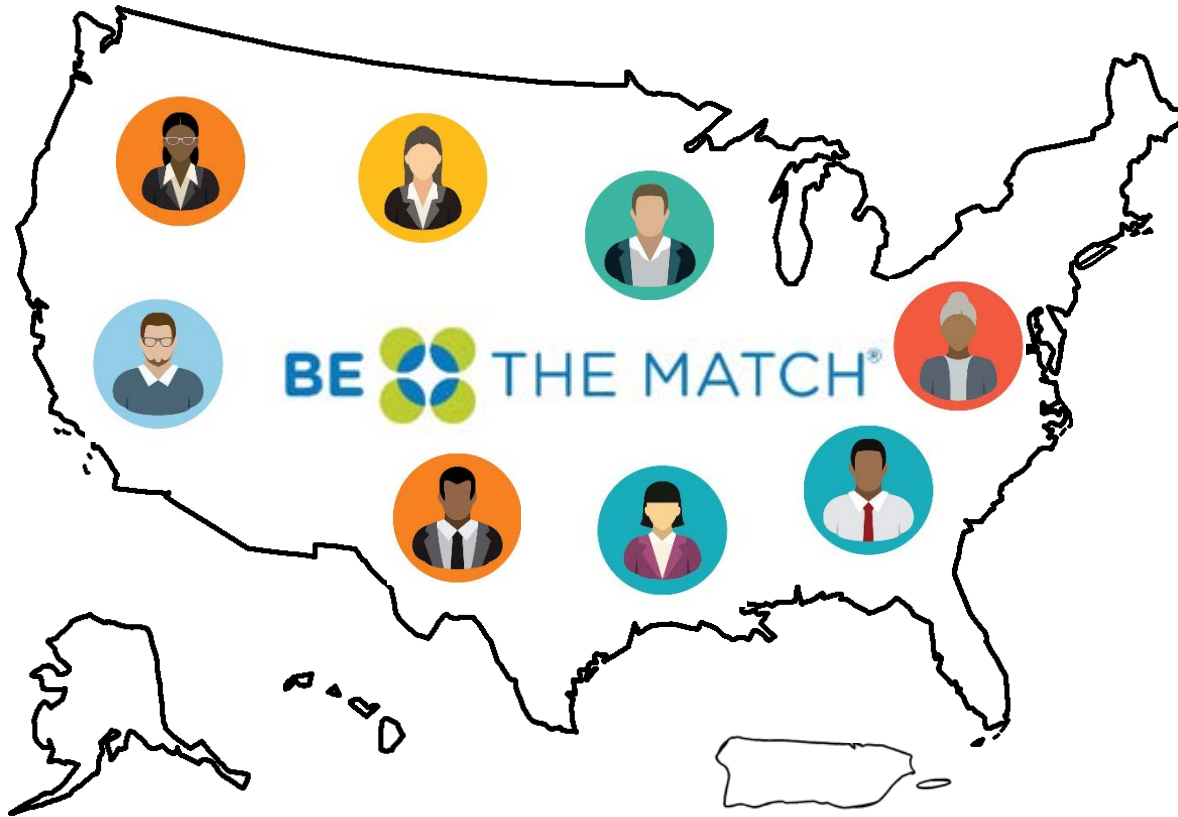
*C-B & DRB1-DQB1 Associations  
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- ✓ Verify HLA typing entry
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- ✓ View potential donor list in Traxis

- ✓ HLA Analysis Tools
  - B-C, DRB1-DQB1 associations
  - Allele and haplotype frequencies
  - HaploStats and IMGT



# NMDP HLA Allele and Haplotype Frequencies



- ✓ Based on the HLA typing of donors on the Be The Match registry
- ✓ HLA allele/haplotype frequencies differ among different ethnic groups
- ✓ Helps identify donor populations that carry HLA alleles/haplotypes of interest
- ✓ Form the basis of the HapLogic matching algorithm

# NMDP Haplotype Frequencies, by the numbers

**6.59**  
**million**  
**subjects**

**6 HLA loci**

HLA-A, -B, -C, -DRB1,  
-DQB1, -DRB3/4/5

**23 ETHNIC GROUPS**



# Accessing NMDP Allele and Haplotype Frequencies

Clinicians Network Payer **Bioinformatics** Search Bioinformatics

NATIONAL MARROW DONOR PROGRAM® BE THE MATCH

HLA Resources ▾ Search

Haplotype Frequencies Allele Frequencies

High Resolution HLA Alleles and Haplotypes in the US Population

HLA-A	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-B	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-DRB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-C	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-DQB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-A~B	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-A~B~DRB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-A~B~DRB1~DQB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-A~C~B	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-A~C~B~DRB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-A~C~B~DRB1~DQB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-B~DRB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-C~B	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-C~B~DRB1~DQB1	<a href="#">XLSX</a>	<a href="#">XLS</a>
HLA-DRB1~DQB1	<a href="#">XLSX</a>	<a href="#">XLS</a>

HLA Resources ▾ Search

Haplotype Frequencies Allele Frequencies

A-B-DRB1 224 Haplotype Frequencies

High-Resolution HLA Alleles and Haplotypes in the US Population

Jewish High-Resolution Haplotype Frequencies

Be The Match Registry Haplotype Frequencies

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race/ethnic categories

Allele Frequencies

Immunology. Available

# HLA-C~B Association Table

C	B	AFA_freq	AFA_rank	API_freq	API_rank	CAU_freq	CAU_rank	HIS_freq	HIS_rank
C*04:01g	B*53:01	0.10790	1	0.00079	102	0.00363	44	0.01530	16
C*02:02g	B*15:03g	0.06076	2	0.00012	165	0.00076	66	0.01129	26
C*07:02g	B*07:02g	0.05984	3	0.02923	12	0.12300	1	0.05652	2
C*04:01g	B*35:01g	0.05482	4	0.03018	11	0.05588	4	0.06563	1
C*17:01g	B*42:01	0.05262	5	0.00012	168	0.00032	92	0.00628	45
C*06:02g	B*58:02	0.04008	6	0.00007	212	0.00013	132	0.00365	63
C*16:01	B*45:01g	0.03735	7	0.00041	121	0.00072	69	0.00920	32
C*04:01g	B*44:03	0.03164	8	0.00199	73	0.01567	18	0.01478	18
C*03:04g	B*15:10	0.03088	9	0.00027	140	0.00031	95	0.00445	55
C*07:01g	B*49:01	0.02804	10	0.00287	61	0.01611	17	0.02362	9

Code	NMDP Ethnic Group
AFA	African American
API	Asian or Pacific Islander
CAU	Caucasian
HIS	Hispanic



Grab your cape.



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## DETERMINE SEARCH STRATEGY



Search Strategy Advice (SSA)

Well Typed Donors  
>90% Likelihood to Match*Matched or Mismatched  
(10/10, 8/8, 9/10, 7/8)*FastTrack  
SearchConsider:  
Proceeding Directly  
to WorkupEvaluate Donor  
Age & DPB1  
Matching (1, 2)Potential Donors Available,  
Matching is UncertainEvaluate Matching Predictions  
of Donors on the Potential List

Search BMDW &amp; Coop Lists



Haplostats

Are there 8/8 matched or C\*03:03/03:04  
permissively mismatched donors? (3, 4)Are there unidirectional  
mismatched donors? (5)

Evaluate Donor Age &amp; DPB1 Matching (1, 2)

Cord Blood Unit  
SearchCord Blood  
Consultation  
Service

Allele Reveal

Matching at HLA-C,  
Antibodies

Criteria:

TNC/kg

CD34/kg

x/8 Matching

RBC Depletion

Total Frozen Volume (6)

# Search Strategy Advice (SSA)

Provides a prioritized list of potential matches to aid in the selection of donors and CBUs by TCs

- ✓ **Make donor/cord blood unit recommendations based on TC criteria**
- ✓ Advisors with HLA expertise use patient HLA typing along with knowledge of haplotypes and linkage frequencies to develop a search strategy
- ✓ Search Be The Match registry and all global registries
- ✓ Turnaround time within 3-5 days
- ✓ Free service that requires no commitment
  - Request for some or all patients

**Well Typed Donors**  
**>90% Likelihood to Match**

*Matched or Mismatched*  
*(10/10, 8/8, 9/10, 7/8)*



**FastTrack  
Search**

**Consider:**  
**Proceeding Directly  
to Workup**



**Evaluate Donor  
Age & DPB1  
Matching (1, 2)**

**>90% likelihood to match**

- ✓ FastTrack Search
- ✓ Proceed directly to workup
- ✓ Evaluate donor age and DPB1 matching

**Grab your cape.**



# CT at WU: Recommendations for donor typing

High-resolution, well-typed donors that are likely to match can CT at WU.

A	B	C	DRB1	DQB1	A	B	C	DRB1	DQB1
<b>A</b> <b>A</b> 99	<b>A</b> <b>A</b> 99	<b>A</b> <b>A</b> 99	<b>A</b> <b>A</b> 99	<b>A</b> <b>A</b> 99	11:01 24:02	35:01 35:02	04:01 04:01	11:04 11:04	03:01 03:01
A	B	C	DRB1	DQB1	A	B	C	DRB1	DQB1
<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	11:AWFBF 24:AYRZE	35:BCZHB 35:TXYP	04:BCZHM 04:BCZHM	11:AMFTT 11:AMFTT	03:AWFCW 03:AWFCW

CT-WU

A	B	C	DRB1	DQB1	A	B	C	DRB1	DQB1
<b>P</b> <b>P</b> 99	<b>P</b> <b>A</b> 99	99	<b>A</b> <b>A</b> 99	99	11:BMTS 24:BZBD	35:BEZX 35:02		11:04 11:04	

CT



## Potential Donors Available, Matching is Uncertain



Evaluate Matching Predictions  
of Donors on the Potential List



Search BMDW & Coop Lists



Are there 8/8 matched or C\*03:03/03:04  
permissively mismatched donors? (3, 4)



Are there unidirectional  
mismatched donors? (5)



Evaluate Donor Age & DPB1 Matching (1, 2)

## Matching is uncertain

- ✓ Evaluate predictions on the Potential List
  - Should donors be HR typed?

Grab your cape.

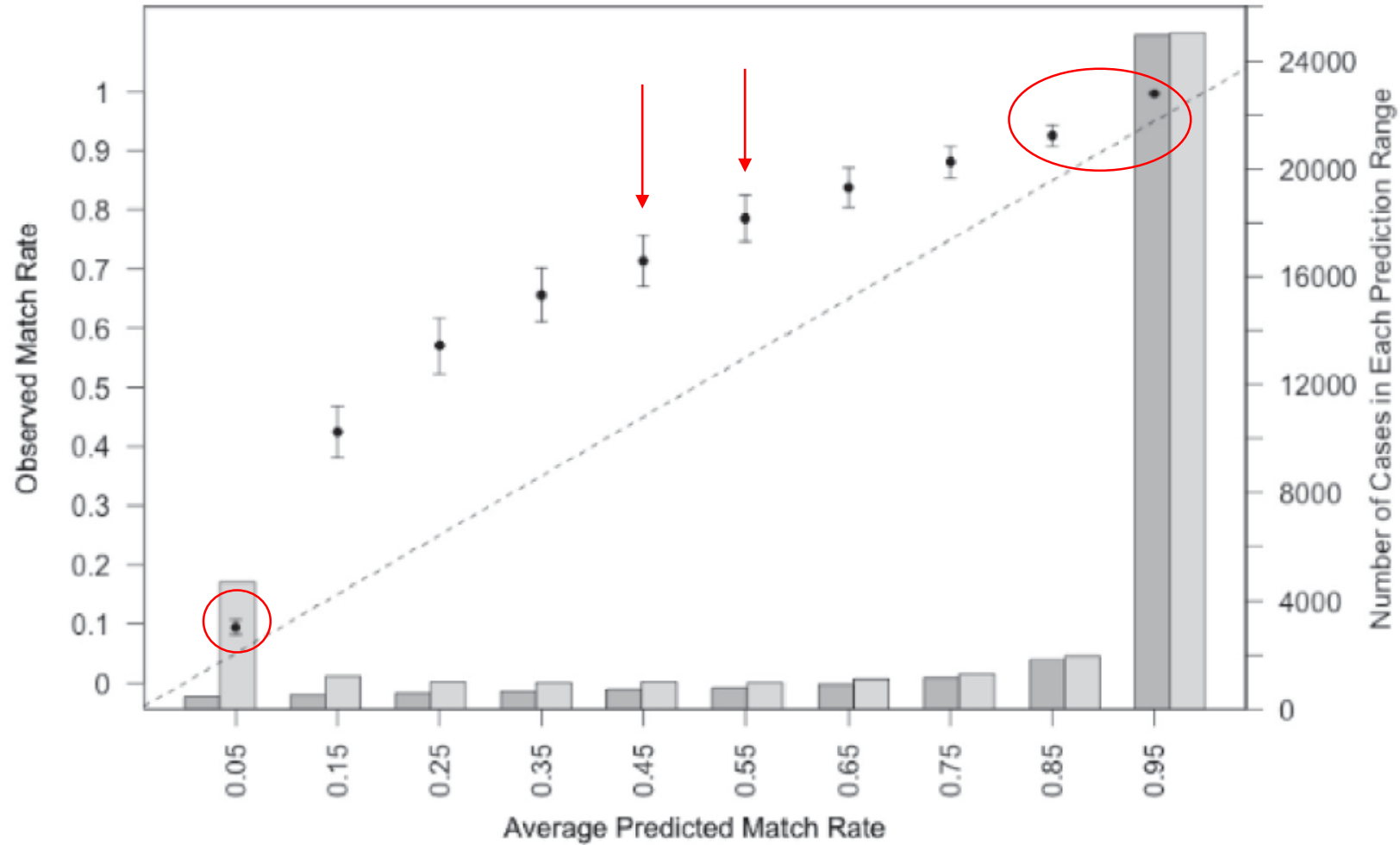


# HapLogic Predictions: Things to Keep in Mind

- ✓ HapLogic prediction of 1% can often be closer to 0%
- ✓ Predictions at low and high ends are most accurate
- ✓ Predictions in intermediate ranges are pessimistic (HapLogic tends to predict lower)



# HapLogic Accuracy



# HapLogic Probability: Case Study example 1

		MCat	Pr(n) of 10 (%)	Pr(n) of 8 (%)	A	B	C	DRB1
Donor 1	Japanese	10/10	10/10=7 9/10=96 8/10=99	8/8=7 7/8=97 6/8=99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	7	<b>P+</b> <b>P+</b> 99
Donor 2	Japanese	10/10	10/10=7 9/10=96 8/10=99	8/8=7 7/8=97 6/8=99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	7	<b>P</b> <b>P</b> 99
Donor 3	Japanese	10/10	10/10=7 9/10=96 8/10=99	8/8=7 7/8=97 6/8=99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	7	<b>P</b> <b>P</b> 99
Donor 4	Japanese	10/10	10/10=7 9/10=96 8/10=99	8/8=7 7/8=97 6/8=99	<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	7	<b>P</b> <b>P</b> 99



# HapLogic Probability: Case Study example 1

		MCat	Pr(n) of 10 (%)	Pr(n) of 8 (%)	A	B	C	DRB1
Donor 2	Japanese	10/10	10/10=99 9/10=99 8/10=99	8/8=99 7/8=99 6/8=99	A A 99	A A 99	A A 99	A A 99
Donor 1	Japanese	9/10	10/10=0 9/10=99 8/10=99	8/8=0 7/8=99 6/8=99	A A 99	A A 99	A M 0	A A 99
Donor 3	Japanese	9/10	10/10=0 9/10=99 8/10=99	8/8=0 7/8=99 6/8=99	P P 99	P P 99	P M 0	A A 99
Donor 4	Japanese	9/10	10/10=0 9/10=99 8/10=99	8/8=0 7/8=99 6/8=99	P P 99	P P 99	P M 0	P A 99

# HapLogic Probability: Case Study example 2

Patient: Asian - Filipino

A	B	C	DRB1	DQB1
24:07 34:01	15:02 40:02	08:01 15:02	12:02 15:02	03:01 05:02

	MCat	Pr(n) of 10 (%)	Pr(n) of 8 (%)	A	B	C	DRB1	DQB1	A	B	C	DRB1	DQB1
Donor 1: Asian - Filipino	10/10	10/10=66 9/10=95 8/10=99	8/8=68 7/8=97 6/8=99	P+ P+ 71	A+ P+ 99	95	A+ A+ 99	92	24:XX 34:XX	15:02 40:BKEZ		12:02 15:02	
Donor 2: Asian - Filipino	10/10	10/10=66 9/10=95 8/10=99	8/8=68 7/8=97 6/8=99	P+ P+ 71	A+ P+ 99	95	A P 99	92	24:XX 34:XX	15:02 40:PJA		12:02 15:AFB	
Donor 3: Asian - Filipino	10/10	10/10=66 9/10=95 8/10=99	8/8=68 7/8=97 6/8=99	P+ P+ 71	A P 99	95	A+ A+ 99	92	24:XX 34:XX	15:02 40:BXZM		12:02 15:02	
Donor 4: Asian - Filipino	10/10	10/10=66 9/10=95 8/10=99	8/8=68 7/8=97 6/8=99	P+ P+ 71	A+ P+ 99	95	A+ A+ 99	92	24:XX 34:XX	15:02 40:BKEZ		12:02 15:02	
Donor 5: Asian - Filipino	10/10	10/10=61 9/10=88 8/10=93	8/8=64 7/8=92 6/8=95	P P 68	P P 94	89	P P 99	90	s24 s34	s75 s61		12:MN 15:ADE	

## Potential Donors Available, Matching is Uncertain



Evaluate Matching Predictions  
of Donors on the Potential List



Search BMDW & Coop Lists



Are there 8/8 matched or C\*03:03/03:04  
permissively mismatched donors? (3, 4)



Are there unidirectional  
mismatched donors? (5)



Evaluate Donor Age & DPB1 Matching (1, 2)

# Matching is uncertain

- ✓ Search BMDW/Coop lists
  - Use HaploStats to evaluate likelihood of matching

Grab your cape.



# HaploStats Overview

*NMDP dataset*

HLA Dataset

NMDP full 2011

*Populations*

Populations

☒ AFA - African American

☐ AAFA - African American

☐ AFB - African

☐ CARB - Caribbean Black

☒ API - Asian or Pacific Islander

☐ AINDI - South Asian Indian

☐ FILII - Filipino

☐ HAWI - Hawaiian or other  
Pacific Islander

☐ JAPI - Japanese

☐ KORI - Korean

☐ NCHI - Chinese

☐ SCSEAI - Southeast Asian

☐ VIET - Vietnamese

☒ CAU - Caucasian

☐ MENAFC - Middle Eastern or  
North Coast of Africa

☐ EURCAU - European Caucasian

☒ HIS - Hispanic

☐ CARHIS - Caribbean Hispanic

☐ MSWHIS - Mexican or Chicano

☐ SCAHIS - South or Central American Hispanic

☒ NAM - Native American

☐ AMIND - North American Indian

☐ CARIBI - Caribbean Indian

Select All

Clear Populations

*Haplotype Loci*

Haplotype Loci

A~C~B~DRB1~DQB1

HLA type

Enter an HLA type:	HLA-A	HLA-B	HLA-C	HLA-DRB1	HLA-DQB1	HLA-DRB3	HLA-DRB4	HLA-DRB5
Type 1	02:01	54:01	01:02	04:05	04:01			
Type 2	24:02	07:02	07:02	01:01	05:01			

SUBMIT QUERY

NATIONAL  
MARROW  
DONOR  
PROGRAM®

BE  THE MATCH®

[www.haplostats.org](http://www.haplostats.org)

*Grab your cape.*



# HaploStats: Case Study example 1

Patient: Asian - Japanese

A	B	C	DRB1
02:01 11:01	35:01 40:06	03:03 08:01	04:05 09:01

NMDP donors:  
unlikely 8/8s

...check BMDW!

	MCat	Pr(n) of 10 (%)	Pr(n) of 8 (%)	A	B	C	DRB1
Donor 1	10/10	10/10=1 9/10=3 8/10=7	8/8=2 7/8=6 6/8=39	<b>P</b> <b>P</b> 46	<b>P</b> <b>P</b> 8	6	<b>P</b> <b>A</b> 95
Donor 2	10/10	10/10=1 9/10=2 8/10=4	8/8=1 7/8=3 6/8=31	<b>P</b> <b>P</b> 48	<b>P</b> <b>P</b> 7	1	<b>P</b> <b>A</b> 84
Donor 3	10/10	10/10=1 9/10=2 8/10=4	8/8=1 7/8=3 6/8=31	<b>P</b> <b>P</b> 48	<b>P</b> <b>P</b> 7	1	<b>P+</b> <b>P+</b> 84

# HaploStats: Case Study example 1

Patient: Asian - Japanese

A	B	C	DRB1
02:01 11:01	35:01 40:06	03:03 08:01	04:05 09:01

BMDW: 8/8 matched Japan donors

MCat	A	B	C	DRB1	A	B	C	DRB1
10/10	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	02:AAYAC 11:AAYFY	35:AAUKN 40:AAVGN	03:AAMGF 08:AAMTE	04:WNEF 09:WNHB
10/10	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	02:AKHNX 11:AKHXZ	35:AKNNS 40:AKPKC	03:AKFWP 08:AKGGF	04:AFGGU 09:AKKHS
10/10	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	<b>P</b> <b>P</b>	02:WPAD 11:WPGF	35:WJUJ 40:WKPW	03:WGUY 08:WHDJ	04:WNEF 09:WNHB

67% **A\*02:06+A\*11:01** **M**  
 C\*03:03+C\*08:01  
 B\*35:01+B\*40:06  
 DRB1\*04:05+DRB1\*09:01  
 DQB1\*03:03+DQB1\*04:01

99% **A\*02:01+A\*11:01** **P**  
 C\*03:03+C\*08:01  
 B\*35:01+B\*40:06  
 DRB1\*04:05+DRB1\*09:01  
 DQB1\*03:03+DQB1\*04:01



# HaploStats: Case Study example 2

Patient: Asian - Korean

A	B	C	DRB1
02:01 33:03	13:01 58:01	03:02 03:04	01:01 13:02

NMDP Summary Counts:

10/10 = 0

9/10 = 473

...check BMDW!

	MCat	Pr(n) of 10 (%)	Pr(n) of 8 (%)	A	B	C	DRB1
Donor 1	9/10	10/10=0 9/10=98 8/10=99	8/8=0 7/8=99 6/8=99	<b>P</b> <b>P</b> 99	<b>M</b> <b>P</b> 0	<b>P</b> <b>P</b> 99	<b>A</b> <b>A</b> 99
Donor 2	9/10	10/10=0 9/10=98 8/10=99	8/8=0 7/8=99 6/8=99	<b>P</b> <b>P</b> 99	<b>M</b> <b>P</b> 0	<b>P</b> <b>P</b> 99	<b>A</b> <b>A</b> 99
Donor 3	9/10	10/10=0 9/10=98 8/10=99	8/8=0 7/8=99 6/8=99	<b>P</b> <b>P</b> 99	<b>M</b> <b>P</b> 0	<b>P</b> <b>P</b> 99	<b>A</b> <b>A</b> 99

# HaploStats: Case Study example 2

Patient: Asian - Korean

A	B	C	DRB1
02:01 33:03	13:01 58:01	03:02 03:04	01:01 13:02

BMDW: potential 8/8 donors

Demographics <a href="#">Add/Remove Data</a>	MCat	A	B	C	DRB1	DQB1	A	B	C	DRB1	
<a href="#">China-CMDP</a> Donor Count: 1	10/10	<div>P</div> <div>A</div>	<div>P</div> <div>A</div>	<div>P</div> <div>P</div>	<div>A</div> <div>A</div>		02:BGNHR 33:03	13:BFBTV 58:01	03:BFBZN 03:BFBZU	01:01 13:02	99%
<a href="#">Brazil</a> Donor Count: 2	10/10	<div>P</div> <div>P</div>	<div>P</div> <div>P</div>		<div>P</div> <div>P</div>		02:XX 33:XX	13:XX 58:XX		01:XX 13:XX	<1%
<a href="#">China-CMDP</a> Donor Count: 1	10/10	<div>P</div> <div>P</div>	<div>P</div> <div>P</div>		<div>P</div> <div>A</div>		02:BGNHR 33:BFBYR	13:BFBTV 58:AHTPA		01:AWFDC 13:02	96%
<a href="#">Korea-KONOS</a> Donor Count: 1	10/10	<div>P</div> <div>P</div>	<div>P</div> <div>P</div>		<div>P</div> <div>P</div>		s2 s33	s13 s58		s1 s13	82%
<a href="#">Korea-KONOS</a> Donor Count: 1	10/10	<div>P</div> <div>P</div>	<div>P</div> <div>P</div>		<div>A</div> <div>P</div>		s2 s33	s13 s58		01:01 13:AB	82%

## Potential Donors Available, Matching is Uncertain



Evaluate Matching Predictions  
of Donors on the Potential List



Search BMDW & Coop Lists



Are there 8/8 matched or C\*03:03/03:04  
permissively mismatched donors? (3, 4)



Are there unidirectional  
mismatched donors? (5)



Evaluate Donor Age & DPB1 Matching (1, 2)

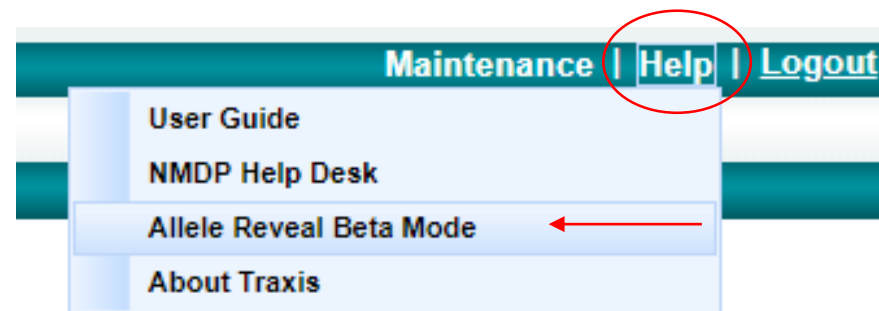
## Matching is uncertain

- ✓ Permissively mismatched donors
  - 8/8 match (DQB1 mismatch)
  - C\*03:03/03:04
- ✓ Unidirectional HvG mismatched 7/8 donors

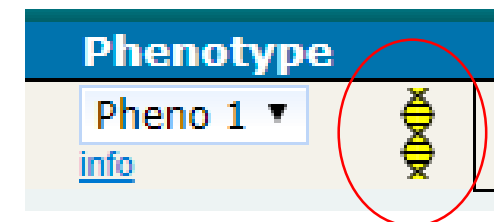
Grab your cape.



# Allele Reveal Overview



- ✓ Allows user to better understand details of the search
- ✓ Displays more comprehensive HLA information for NMDP donors and cord blood units
- ✓ Most likely allele level typing in place of allele codes and at untyped loci, based on the HapLogic algorithm



# Permissive Mismatch: Case Study example

Patient

A	B	C	DRB1	DQB1
02:01 03:01	15:01 35:03	03:03 04:01	04:01 10:01	03:01 05:01

Permissive mismatch: C\*03:03/C\*03:04

	MCat	Pr(n) of 10 (%)	Pr(n) of 8 (%)	A	B	C	DRB1	DQB1
Donor 1: 49 F	10/10	10/10=65 9/10=99 8/10=99	8/8=65 7/8=99 6/8=99	P P 99	P P 99	P P 65	A A 99	P A 99
Donor 2: 31 F	10/10	10/10=59 9/10=99 8/10=99	8/8=59 7/8=99 6/8=99	P P 99	P P 99	59	A A 99	P P 99



Allele Reveal

C Genotype Probabilities		
Genotype	Percent	MatchGrade
C*03:03+C*04:01	65%	A A
C*03:04+C*04:01	35%	L A

## Cord Blood Unit Search



Cord Blood  
Consultation  
Service



Allele Reveal

*Matching at HLA-C,  
Antibodies*



*Criteria:*

TNC/kg

CD34/kg

x/8 Matching

RBC Depletion

Total Frozen Volume (6)

# Cord Blood Unit Search

- ✓ Cord Blood Clinical Consultation service
- ✓ Use Allele Reveal to help predict matching

Grab your cape.



# Allele Reveal: Case study example 1

Patient typing

A	B	C	DRB1
02:01 24:02	07:02 51:01	02:02 07:02	01:01 13:02

Potential 6/8 cord blood unit

A	B	C	DRB1	DQB1	A	B	C	DRB1
<b>P</b> <b>P</b> 99	<b>P</b> <b>P</b> 99	9	<b>M</b> <b>P</b> 0	1	02:XX 24:XX	07:XX 51:AFKE		04:AFMZ 13:XX

*Use Allele Reveal mode in Traxis to avoid two DRB1 mismatches*

Likely double DRB1 mismatch

A	B	C	DRB1
<b>P</b> 02:01g <b>P</b> 24:02g 99 info (d)	<b>P</b> 07:02g <b>P</b> 51:01g 99 info (d)	(14:02g) (07:02g) 9 info	<b>M</b> 04:01 <b>P</b> 13:01g 0 info (d)

Locus Information		
DRB1 Genotype Probabilities		
Genotype	Percent	MatchGrade
DRB1*04:01+DRB1*13:01	76%	M L
DRB1*04:01+DRB1*13:02	21%	M A
DRB1*04:01+DRB1*13:03	3%	M L
DRB1*04:01+DRB1*13:05	1%	M L

# Allele Reveal: Case study example 2

## Patient typing

A	B	C	DRB1
02:01 24:02	07:02 51:01	02:02 07:02	01:01 13:02

## Antibody report

Class I	Class II
A11, 23, 24, 68, 69	DR17, 18
B8, 44, 45, 57, 58, 82	DQ2, 4
Cw5, 14, 15	DP10, 11, 13, 23, 28

## Potential 6/8 cord blood unit

A	B	C	DRB1	DQB1	A	B	C	DRB1
<b>A</b> <b>M</b> 0	<b>A</b> <b>A</b> 99	6	<b>A</b> <b>A</b> 99	92	02:01 01:01	07:02 51:01		01:01 13:02

A	B	C	DRB1
<b>A</b> 02:01g• <b>M</b> 01:01g• 0 <u>info (d)</u>	<b>A</b> 07:02g• <b>A</b> 51:01g• 99 <u>info (d)</u>	(01:02g) (07:02g) 6 <u>info</u>	<b>A</b> 01:01g• <b>A</b> 13:02g• 99 <u>info (d)</u>

Locus Information		
C Genotype Probabilities		
Genotype	Percent	MatchGrade
C*01:02+C*07:02	29%	M A
C*14:02+C*07:02	28%	M A
C*15:02+C*07:02	23%	M A
C*02:02+C*07:02	6%	A A
C*16:02+C*07:02	5%	M A
C*04:01+C*07:02	2%	M A
C*05:01+C*07:02	2%	M A
C*12:03+C*07:02	1%	M A
C*07:02+C*07:02	1%	M A
C*16:01+C*07:02	1%	M A

Use Allele Reveal to identify units that could carry an antigen to which the patient has a reported antibody





# 4

## SELECT POTENTIAL SOURCES

### Consider:

TC Criteria, Patient Antibodies, Donor Last Contact Date



Search  
Assistance Funds

- ✓ TC criteria
- ✓ Patient anti-HLA antibodies
- ✓ Donor last contact date
- ✓ Apply for Search Assistance Funds

# Search Assistance Funds to the rescue...

SAF is here for your patients **when you can't start a search** because:

- Insurance doesn't cover search costs
- Search coverage is inadequate for what you need to request
- Patient doesn't have insurance yet (waiting for Medicaid approval, for example)
- You're waiting for prior authorization
- Fill in other financial or insurance barrier here \_\_\_\_\_

SAF is a **limited guarantee of approval** for specific NMDP search activities, per NMDP fee schedule

- Can cover Fast Track Service and workup/procurement costs, as well

Responses given within **one business day**

Payment is through **reimbursement** to your TC, on behalf of the patient

Applications must be approved **before requesting sources** (TCC or financial coordinator submits)

**Contact:** patientgrants@nmdp.org or 763-406-8114



5

## REQUEST TYPING

Consider:  
CT vs Customized Typing, Turnaround Time



6

## REQUEST WORKUP OR SHIPMENT

- ✓ Donor typing
  - CT vs. HR screening
- ✓ Select backup donor(s)

- ✓ FastTrack Testing
- ✓ Electronic forms reporting



# NMDP Laboratory Services

## FastTrack Testing

- ✓ Rapid, high quality, cost-effective histocompatibility testing
  - Partnership with an ASHI-accredited contract laboratory
  - Turnaround time of 3 business days
  - Allele level confirmatory testing
  - Additional testing: KIR, CCR5  $\Delta$ 32 mutation, Antibody screening, Crossmatch, Chimerism

## Electronic Forms reporting

- ✓ NMDP Lab Services staff will work with your lab to set it up
  - Elimination of paper Form 117 and/or Form 22
  - More immediate HLA results from Traxis
  - Reduced potential for submission errors



# HLA typing

high resolution  
antibodies  
DPB1

## Traxis

SSA  
10/10, 8/8, 9/10, 7/8  
BMDW/Coop  
CBUs

## Donor/CBU selection

search assistance funds  
HR, CT, WU request  
Forms 117, 22

**Transplant**



*Grab your cape.*



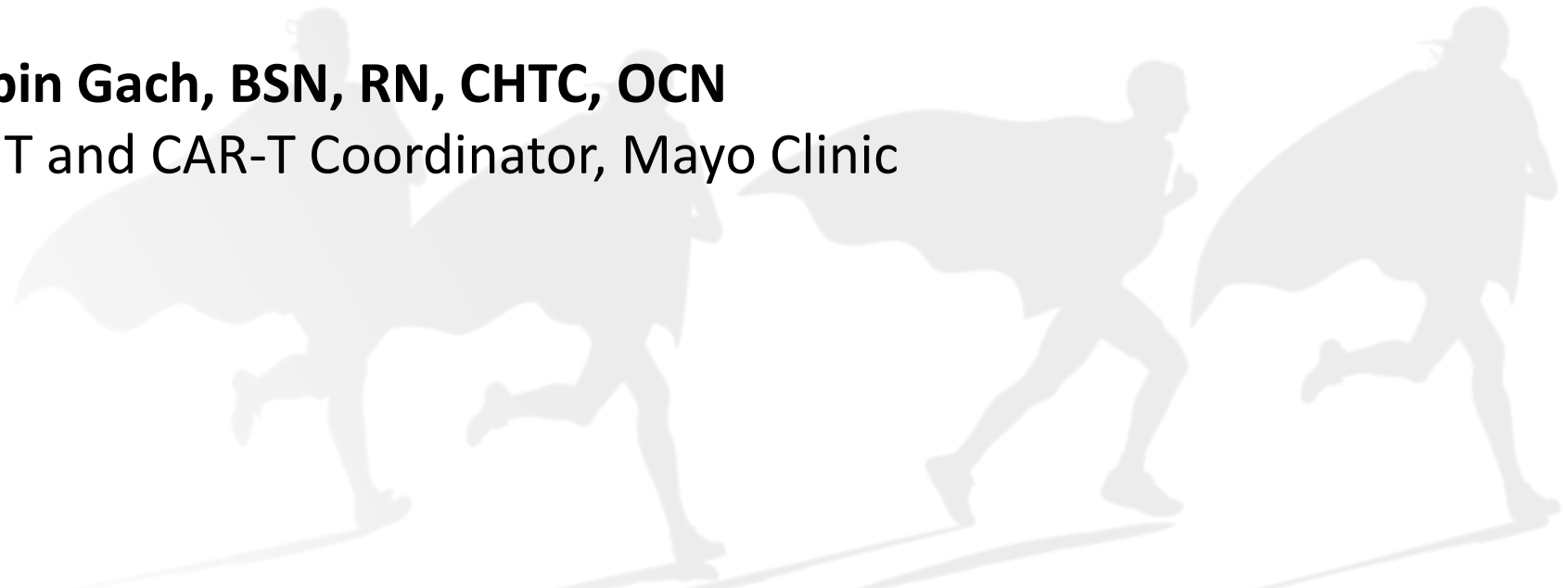
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# ***Thank you!***



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