

Donor Availability: Related and Unrelated

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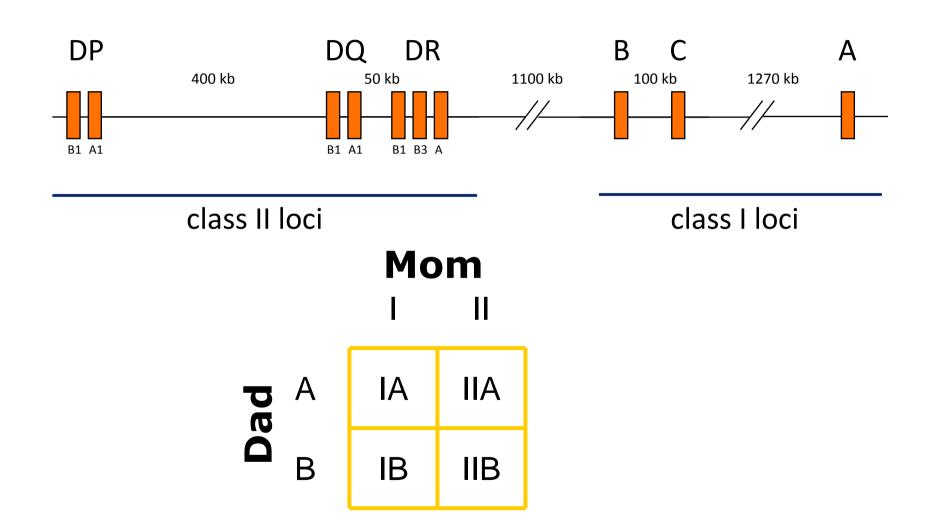
National Marrow Donor Program

November 11, 2011

What Does "Availability" Mean?

- A "matching" HLA type
- Is in the correct age range
 - Related donors: 6 mos 75+ years
 - Unrelated donors: Birth or 18 60 years
- Can be located: "contactable"
- Is willing to donate: "interested"
- Is healthy enough to donate: "suitable"
- Has no disease that will be transmitted infectious or genetic
- Is not pregnant or "temporarily" unable

The HLA System on Chromosome #6



Likelihood of a Sibling Match

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Match			hΛ	\sim
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$$1 - (0.75)^{n}$$

$$1 - (0.75)^1 = 0.25$$

$$1 - (0.75)^2 = 0.44$$

$$1 - (0.75)^4 = 0.68$$

$$1 - (0.75)^7 = 0.87$$

USA:

$$1 - (0.75)^{1.3} = 0.30$$

General Approach to Finding a Donor

- HLA type willing family members
 - Full siblings
 - Parents (to identify haplotypes)
- If unrelated donor is considered -
 - Search for adult donor
 - Search for umbilical cord blood unit(s)
- Consider haploidentical donors

Worldwide Unrelated Donor Pool

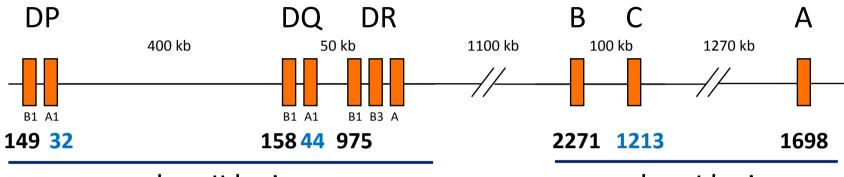
- World Marrow Donor Association (WMDA) surveyed 79 registries and 156 cord blood banks in 2010
 - 15.6 million HLA-A, B and DR-typed adults
 - 592,000 umbilical cord blood units
- Most are accessible through the Bone Marrow Donors Worldwide (BMDW)
 - www.bmdw.org

Asian Donors and Cord Blood Units

- Estimated 2.6 million adult donors in Western Pacific and South-East Asia Regions
- Estimated 213,000 Cord Blood Units
- Actual matching likelihoods are unknown
 - What percent of a population will find a suitable unrelated HLA match?

Huge Numbers of HLA Alleles

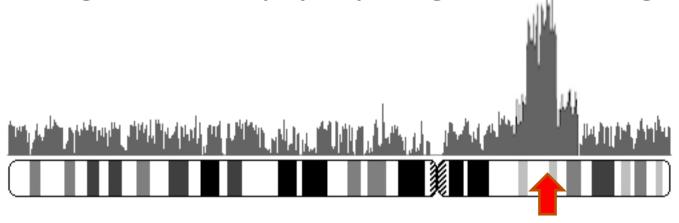
Most human genes have only a few (5-10) variants (alleles)



class II loci

class I loci

The HLA region is the most polymorphic region of the human genome



July 2011

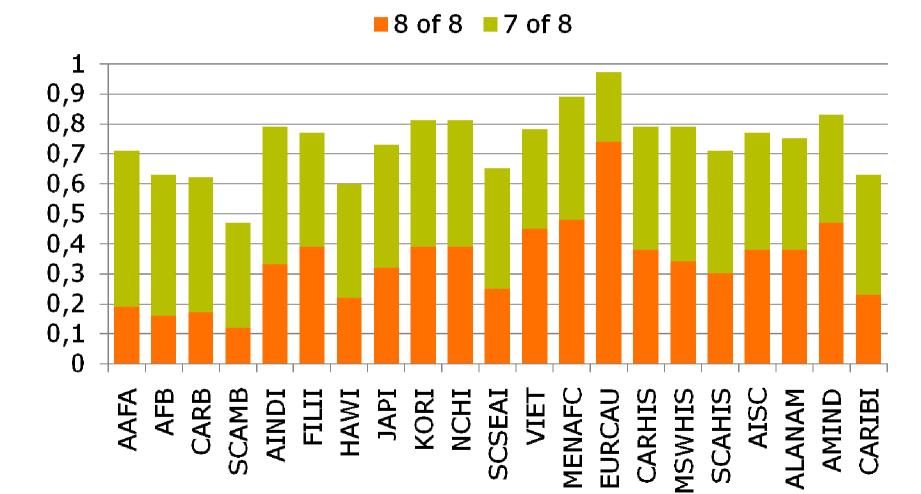
21 US Populations

Race_Code	Description		
AAFA	African American		
AFB	African	AFAA	
CARB	Black Caribbean	AFAA	
SCAMB	Black South or Central America		
AINDI	South Asian		
FILII	Filipino	API	
HAWI	Hawaiian or other Pacific Islander		
JAPI	Japanese		
KORI	Korean		
NCHI	Chinese		
SCSEAI	Other Southeast Asian		
VIET	Vietnamese]	
MENAFC	MidEast/No. Coast of Africa	CAU	
EURCAU	European Caucasian	CAU	
CARHIS	Caribbean Hispanic		
MSWHIS	Mexican or Chicano	HIS	
SCAHIS	South/Cntrl Amer. Hisp.		
AISC	American Indian South or Central American		
ALANAM	Alaska Native or Aleut	NAM	
AMIND	North American Indian		
CARIBI	Caribbean Indian		

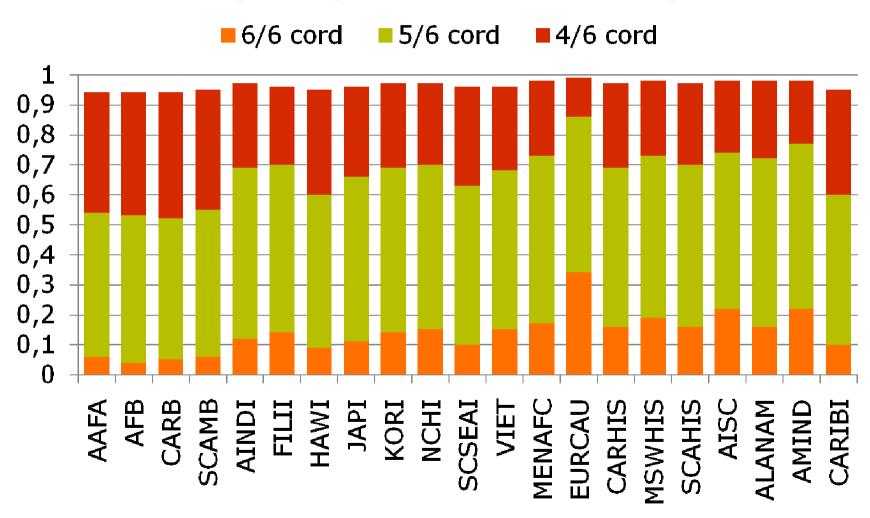
21 US Populations

Race_Code	"Available" Donors in NMDP File		
AAFA	140,277		
AFB	7,982	AFAA	
CARB	9,623		
SCAMB	1,688		
AINDI	73,065		
FILII	19,822		
HAWI	4,146	API	
JAPI	10,479		
KORI	33,625		
NCHI	37,469		
SCSEAI	10,035		
VIET	9,483		
MENAFC	182,211	CALL	
EURCAU	3,216,667	CAU	
CARHIS	52,862		
MSWHIS	136,685	HIS	
SCAHIS	62,844		
AISC	2,514	NAM	
ALANAM	650		
AMIND	17,893	IVAIM	
CARIBI	2,157		

7/8 and 8/8 Allele, Available-Match Rates in the Adult Donor Registry - 2010



Pediatric Cord Match Rates in the Cord Blood Registry, Dose ≥2.5/Kg - 2010



Donor Availability: Summary

- Many factors impact donor availability
- HLA matching is one of the most important
 - Relatively trivial within the family
 - Complex in the unrelated donor setting
 - New techniques are helping to model
 HLA matching with greater accuracy
- Sufficient numbers of ethnically diverse donors and cord blood units are essential