From: Tull, Lori
Sent: Friday, March 11, 2016 9:28 AM
To: Wouter Van't Hof (wvanthof@clevelandcordblood.org)
Subject: Clinical information request
Attachments: Cleveland Cord Blood BLA_additional clinical data request_March 10
2016.docx

Hi Wouter,

Could you please fill in the tables in the attached document and provide an answer to the
question at the end of the document, with a two-week deadline? All of the additional clinical
information that's requested in the file refers to the original dataset for the BLA. Please make
sure to submit the information as an amendment to the BLA as well as email.

Thanks very much,
Lori Tull
Team Lead
Regulatory Management Staff
Office of Cellular, Tissue, and Gene Therapies
Center for Biologics Evaluation and Research
(240) 402-8361
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Table 1. Demographic Characteristics

Patient Characteristics	Patients Who Received a Suitable Allograft ($\geq 2.5 \times 10^7$ TNC/kg and HLA match $\geq 4/6$)	Patients Who Received $\geq 2.5 \times 10^7$ TNC/kg
Median Age (range)		
Age Category	# (percentage)	
<2yrs		
2- <17yrs		
>17yrs		
Unknown		
Sex		
Male		
Female		
Unknown		
Ethnicity/Race		
White		
African-American		
Hispanic		
Asian		
Other		
Unknown		
Diagnosis		
Hematologic Malignancies		
Inborn Errors of Metabolism		
Immunodeficiency		
Metabolic Disorders		
Bone Marrow Failure		
Hemoglobinopathy		
Others		
Unknown		

Table 2. Graft Failure by HLA Match and TNC Dose

HLA Match	TNC Dose (x 10^7/kg)		
	2.5 to <5	**5 to <10**	**>10**
4/6	# of reported graft failures / # of patients with the given HLA match and TNC dose characteristics (percentage)		
5/6			
6/6			

Table 3. Unit Characteristics and Dose Exposure

Unit Characteristics	Patients Who Received a Suitable Allograft	Patients Who Received $\geq 2.5 \times 10^7$ TNC/kg
Number of Patients		
HLA Match Level		
4/6	# (percentage)	
5/6		
6/6		
Not Reported		

Table 4. Early Mortality (Death $\leq$ Day 100) with Demographic Characteristics

Patient Characteristics	Patients Who Received a Suitable Allograft ($\geq 2.5 \times 10^7$ TNC/kg and HLA match $\geq 4/6$)	Patients Who Received $\geq 2.5 \times 10^7$ TNC/kg
Median Age (range)		
Age Category	# Early Deaths/# of Patients in the given category (percentage)	
<2yrs		
2- <17yrs		
>17yrs		
Unknown		
Gender		
Male		
Female		
Unknown		
Ethnicity/Race		
White		
African-American		
Hispanic		
Asian		
Other		
Unknown		
Diagnosis		
Hematologic Malignancies		
Inborn Errors of Metabolism		
Immunodeficiency		
Metabolic Disorders		
Bone Marrow Failure		
Hemoglobinopathy		
Others		
Unknown		

Table 5. Causes of Death after Transplantation

	Patients Who Received a Suitable Allograft (≥ 2.5 x 10^7 TNC/kg and HLA match $\geq 4/6$)	Patients Who Received ≥ 2.5 x 10^7 TNC/kg
Causes of Death		
Graft failure	# (%)	
Organ failure		
Infection		
GVHD		
Primary disease		
2nd Malignancy		
Prior malignancy		
Hemorrhage		
Pulmonary toxicity		
Unknown		
Other		

Table 6. Incidence of Infusion Reactions

	Patients Who Received at Least a Single Unit N=	Patients Who received ≥ 2.5 x 10^7 TNC/kg N=
# of Infusions with a Reported Infusion Reaction (Percentage of Infusions)	# (%)	
Signs and symptoms reported: Number (%) of patients with this event		
Hypertension	# (%)	
Hypotension		
Hypoxia		
Nausea		
Headache		
Tachycardia		
Vomiting		
Chest Pain		
Fever, Chills		
Rigor mild		
Hives		
Bradycardia		
SOB		
Other		

Questions:
1. With regard to Table 8-3 on Page 7 of Section 31.8, for each of the patients who experienced primary graft failure, please indicate whether the patient received a suitable allograft by providing information about the dose of TNC/kg.