

From: Chunrong Cheng

Through: Jessica Kim, Team Leader

To:

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Subject: Filing of STN 125351/0 for Nycomed's BLA for TachoSil™

Comments to the CBER review committee:

1. This BLA submission provides the raw/tabulation (SDTM) datasets and the analysis (ADaM) datasets for studies -----(b)(4)-----, TC-023-IM, -(b)(4)-----.
2. For studies TC-018-IN and TC-019-IN, only the tabulated datasets were submitted. It could be acceptable since these studies will be used for the evaluation of safety only.
3. The three highlighted studies in the following table are completed. CBER should request the datasets from the sponsor for the evaluation of safety.
4. The SAS analysis programs for the integrated analysis of efficacy and safety were submitted. However, the SAS analysis programs for the individual analysis of studies -----(b)(4)----- were not submitted. CBER should request these datasets from the sponsor. Based on the email communication with the sponsor, these programs are available on request.
5. This BLA is fileable based on the information submitted.

Table 1: Completed Clinical Studies on TachoSil

	Surgical indication	Study Code	Treatments ^a	N	Study Period
Controlled studies providing safety and efficacy data					
1	(b)(4)		TS vs standard surgical treatment	189	2000-2002
2			TS vs standard surgical treatment	301	2006-2007
3	Cardiovascular surgery	TC-023-IM	TS vs hemostatic fleece	120	2006-2007
Controlled studies providing safety data and supportive efficacy data					
4	(b)(4)			121	2001-2002
5				188	2002-2004
6				119	2003-2003
Uncontrolled studies providing additional safety and efficacy ^b data					
7	Liver resection (children)	TC-019-IN	TS (no comparator)	16	2006-2007
8	Various surgeries	TC-018-IN	TS (no comparator)	3098	2005-2008
9	(b)(4)		TS (no comparator)	154	2004-2005
10			TS (no comparator)	616 ^c	2005-2007
11			TS (no comparator)	169	2007-2008

Comments to the sponsor:

1. Please provide SAS XPORT transport files for datasets for studies -----(b)(4)-----
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2. Please provide the corresponding SAS program files using “.sas” for the file
extension for the individual analysis of studies -----(b)(4)-----.