CHARLES RIVER LABORATORIES
Preclinical Services

FINAL REPORT
Volume 1 of 12

Guidelines: OECD (451) and JMHLW (Ordinance No. 21)

Testing Facility Study No. EBA00009

Sponsor Ref. No. P00012-04-11

A Two Year Oral (Gavage) Carcinogenicity Study in Rats with BG00012

TESTING FACILITY:

Charles River Laboratories
Preclinical Services
640 North Elizabeth Street
Spencerville, OH 45887

SPONSOR:

Biogen Idec Inc.
14 Cambridge Center
Cambridge, MA 02142

February 28, 2008

1. COMPLIANCE STATEMENT

This study was conducted in compliance with the Good Laboratory Practice (GLP) regulations as described by the FDA (21 CFR Part 58); the Organisation for Economic Cooperation and Development (OECD) Principles of Good Laboratory Practice, C(97)186; and the Japanese Ministry of Health, Labor, and Welfare (MHLW) Ordinance No. 21 with the following exception(s):

Characterization and stability analyses of the bulk test article were conducted in compliance with the Good Manufacturing Practice regulations. Toxicokinetic interpretation was not conducted in compliance with the GLP regulations.

Mark A. Morse 28-FEB-2008

Mark A. Morse, Ph.D., DABT Date
Study Director
Charles River Laboratories
Preclinical Services

2. QUALITY ASSURANCE STATEMENT

This study has been inspected by the Quality Assurance Unit to assure conformance with the Good Laboratory Practice (GLP) regulations promulgated by the FDA (21 CFR Part 58); OECD Principles of Good Laboratory Practice, C(97)186; and the Japanese MHLW Ordinance No. 21. Reports were submitted in accordance with Standard Operating Procedures as follows:

QA INSPECTION DATES

		Date Findings Submitted to:	
Dates of Inspection	Phase(s) Inspected	Study Director	Study Director Management
10/12/04	Animal Receipt Procedure	10/12/04	10/12/04
10/12/04	Protocol Review	10/15/04	10/15/04
10/14/04	Animal Receipt	10/15/04	10/15/04
10/25/04	Animal Identification	10/28/04	10/28/04
10/25/04	Randomization Procedure	10/28/04	10/28/04
10/25/04	Dose Preparation	10/28/04	10/28/04
10/25/04	Analytical Sampling	10/28/04	10/28/04
10/25/04	Test Article Receipt Procedure	11/01/04	11/01/04
10/26/04	Body Weights	10/28/04	10/28/04
10/26/04	Detailed Clinical Observations	10/28/04	10/28/04
10/26/04	Food Consumption	10/28/04	10/28/04
10/26/04	Dosing	10/28/04	10/28/04
11/01/04	Retention Sample	11/01/04	11/01/04
11/04/04	Protocol Amendment Review	02/11/05	02/11/05
01/18/05, 02/16/05, 02/22/05, 09/02/05, 09/12/05, 09/19/05, 10/20/05	Data Audit	11/14/05	11/14/05
03/07/05	Dosing	03/08/05	03/08/05
03/07/05	Dose Preparation	03/08/05	03/08/05
03/07/05	Analytical Sampling	03/08/05	03/08/05
03/08/05	Detailed Clinical Observations	03/08/05	03/08/05
03/15/05	Body Weights	03/15/05	03/15/05
03/15/05	Food Consumption	03/15/05	03/15/05
03/15/05	Protocol Amendment Review	05/27/05	05/27/05
04/05/05	Toxicokinetic Blood Collection	04/05/05	04/05/05
04/05/05	Plasma Processing	04/05/05	04/05/05
04/05/05	Dosing	04/05/05	04/05/05
04/26/05	Protocol Amendment Review	05/27/05	05/27/05
05/23/05	Dose Preparation	05/23/05	05/23/05
05/23/05	Dosing	05/23/05	05/23/05
09/12/05	Dose Preparation	09/13/05	09/13/05
09/12/05	Dosing	09/13/05	09/13/05
09/13/05	Detailed Clinical Observations	09/13/05	09/13/05
09/13/05	Palpable Masses	09/13/05	09/13/05
10/03/05	Bioanalytical Sample Shipping	10/03/05	10/03/05
10/17/05	Protocol Amendment Review	10/17/05	10/17/05
12/29/05	Protocol Amendment Review	05/03/06	05/03/06

Dates of Inspection	Phase(s) Inspected	Date Findings Submitted to: Study Director	Date Findings Submitted to: Study Director Management
01/09/06	Dose Preparation	01/09/06	01/09/06
01/09/06	Dosing	01/09/06	01/09/06
01/09/06	Analytical Sampling	01/09/06	01/09/06
02/03/06	Data Audit	02/16/06	02/16/06
03/01/06, 03/02/06	Data Audit	03/02/06	03/02/06
03/13/06	Dose Preparation	03/14/06	03/14/06
03/14/06	Dosing	03/14/06	03/14/06
03/14/06	Food Consumption	03/14/06	03/14/06
04/07/06, 04/10/06, 04/11/06	Data Audit	04/11/06	04/11/06
04/19/06	Sentinel Blood Collection	04/19/06	04/19/06
04/19/06	Serum Processing	04/19/06	04/19/06
05/19/06	Protocol Amendment Review	05/26/06	05/26/06
05/25/06	Data Audit	05/25/06	05/25/06
06/19/06	Dose Preparation	06/19/06	06/19/06
06/19/06	Dosing	06/19/06	06/19/06
06/20/06	Protocol Amendment Review	07/07/06	07/07/06
06/26/06	Analytical Sampling	07/07/06	07/07/06
08/31/06	Data Audit	08/31/06	08/31/06
09/01/06, 09/06/06, 09/07/06, 09/08/06, 09/18/06, 09/19/06	Data Audit	09/19/06	09/19/06
09/07/06	Data Audit	09/07/06	09/07/06
09/08/06	Data Audit	09/08/06	09/08/06
09/11/06	Dose Preparation	09/12/06	09/12/06
09/11/06	Dosing	09/12/06	09/12/06
09/12/06	Detailed Clinical Observations	09/12/06	09/12/06
09/12/06	Palpable Masses	09/12/06	09/12/06
09/14/06	Data Audit	09/15/06	09/15/06
09/18/06	Data Audit	09/18/06	09/18/06
09/19/06, 09/20/06	Data Audit	09/21/06	09/21/06
10/23/06	Protocol Amendment Review	10/23/06	10/23/06
10/24/06	Necropsy	10/27/06	10/27/06
10/25/06	Clinical Pathology Blood Collection	10/27/06	10/27/06
10/26/06	Data Audit	10/27/06	10/27/06
10/27/06	Sentinel Blood Collection	10/27/06	10/27/06
10/27/06	Serum Processing	10/27/06	10/27/06
10/31/06	Data Audit	10/31/06	10/31/06
11/01/06	Data Audit	11/01/06	11/01/06
11/03/06	Data Audit	11/03/06	11/03/06
11/06/06	Data Audit	11/06/06	11/06/06
11/08/06	Protocol Amendment Review	11/15/06	11/15/06
11/15/06, 11/16/06	Data Audit	11/16/06	11/16/06
11/22/06	Trimming	11/28/06	11/28/06
11/27/06, 11/28/06, 11/29/06	Data Audit	11/29/06	11/29/06
11/30/06	Data Audit	11/30/06	11/30/06
12/04/06, 12/05/06	Data Audit	12/05/06	12/05/06

Dates of Inspection	Phase(s) Inspected	Date Findings Submitted to: Study Director	Date Findings Submitted to: Study Director Management
12/05/06	Data Audit	12/08/06	12/08/06
12/06/06, 12/07/06	Data Audit	12/11/06	12/11/06
12/07/06, 12/08/06, 12/11/06	Data Audit	12/12/06	12/12/06
12/21/06	Data Audit	12/21/06	12/21/06
01/09/07	Protocol Amendment Review	01/11/07	01/11/07
01/11/07	Microtomy	01/12/07	01/12/07
01/26/07	Quality Control	01/29/07	01/29/07
04/21/07	Protocol Amendment Review	05/01/07	05/01/07
06/19/07	Protocol Amendment Review	06/19/07	06/19/07
07/06/07, 07/10/07, 07/11/07, 07/12/07, 07/16/07, 07/17/07, 07/18/07, 07/19/07	Draft Report Review	07/20/07	07/20/07
10/16/07	Protocol Amendment Review	10/29/07	10/29/07
01/28/08	Protocol Amendment Review	01/31/08	01/31/08
02/19/08, 02/20/08	Final Report Review	02/20/08	02/20/08

QA statement(s) provided by the following test site(s) have been reviewed:

Test Site(s)	Phase	QA Statement Location
Charles River Laboratories, Preclinical Services	Analytical Chemistry Report	Appendix 3
Charles River Laboratories	Pretest Health Screen (Serology)	Appendix 4
Charles River Laboratories	Sentinel Animals (Serology)	Appendix 5
Charles River Laboratories, Pathology Associates	Histopathology Report	Appendix 13
Charles River Laboratories, Pathology Associates	Histopathology Peer Review	Appendix 13
BioSTAT Consultants, Inc.	Statistical Analysis of Survival and Tumor Incidence	Appendix 14
Charles River Laboratories, Preclinical Services	Bioanalytical Report	Appendix 18
Biogen Idec Inc.	Toxicokinetic Interpretive Report	Appendix 19

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