

NOV 18 1998

510k Summary of Safety and Effectiveness

K983740

A summary of 510(k) safety and effectiveness information is being submitted in accordance with the requirements of SMDA 1990 and 21 CFR 807.92.

The assigned 510(k) number is:

Applicant Information:

Date Prepared: Oct 20, 1998
Name: Columbia Bioscience, Inc.
Address: 8775 M Centre Park Drive, #559
Columbia, MD 21045

Contact Person: Norman Jenkins
PhoneNumber: 410-995-0450
Fax Number: 410-995-0448

Device Information:

Trade Name:  anti-PR-3 IgG ELISA Kit
Common Name: PR-3 IgG EIA Test
Classification Name: PR-3 Serological Reagent

Bivalent Device:
ELIAS PR-3 Kit

Device Description: The  PR-3 IgG ELISA Kit is an enzyme-linked immunosorbent assay (ELISA) for the detection of IgG to Proteinase-3 in human serum.

Intended Use: The  PR-3 IgG kit is an Enzyme-Linked Immunosorbent Assay (ELISA) For the qualitative detection and semi-quantitation of antibodies against the Proteinase-3 antigen in serum as an aid in the diagnosis of Wegeners granulomatosis. The test can be performed either manually or in conjunction with the Mago Plus automated EIA processor.

Principle of Procedure:

PR-3 antigen is bound to microwells. Diluted patient sera, Calibrators and controls are placed in the microwells and incubated. Anti-PR-3 IgG antibodies, if present, will bind to the antigen forming antigen-antibody complexes. Residual sample is eliminated by aspirating and washing. Conjugate (horseradish peroxidase-labeled anti-human IgG) is added and will bind to these complexes. Unbound conjugate is removed by aspiration and washing. Substrate is then added and incubated. In the presence of bound enzyme the substrate is converted to an end product. The absorbance of this end product can be read spectrophotometrically at 450 nm (reference 600-630 nm) and is directly proportional to the concentration of IgG antibodies to PR-3 present in the sample.

Performance Characteristics

A. Relative Sensitivity and Specificity

Frozen retrospective sera from two hundred and fifty-six patients were tested on the Is-anti-PR-3 IgG Test Kit and another commercially available ELISA for anti-PR-3 antibodies. Based on the results of this testing, the relative sensitivity and specificity were calculated. The results obtained are shown in Table 2:

TABLE 2
Is-anti-PR-3 IgG

		POSITIVE	EQUIVOCAL	NEGATIVE
Other ELISA	POSITIVE	57	0	0
	EQUIVOCAL*	1	0	0
	NEGATIVE	4	7	187

Relative Sensitivity	$57/57 = 100.0\%$	<i>95% CI</i> 93.7-100%
Relative Specificity	$187/191 = 97.9\%$	94.7- 99.4%
Overall Agreement	$244/248 = 98.4\%$	95.9- 99.6%

* Equivocal results were excluded from calculations.

NOTE : Please be advised that "relative" refers to the comparison of the assay's results to that of a similar assay. There was not an attempt to correlate the assay's results with disease presence or absence. No judgment can be made on the comparison's accuracy to predict disease.

B. Clinical Sensitivity and Specificity Using Characterized Sera

A total of 256 frozen retrospective, clinically characterized sera were assayed using the Is-anti-PR-3 IgG Test Kit. The results obtained are shown in Table 3.

TABLE 3

<u>Patient Group:</u>	<u>Positive</u>	<u>Equivocal*</u>	<u>Negative</u>	<u>Total</u>
Normals	3	5	168	176
Wegener's Granulomatosis	38	0	2	40
Microscopic Polyangiitis	21	2	17	40

Clinical Specificity:

Normals	$168/171 = 98.3\%$	<u>95% CI</u> 95.0-99.6
---------	--------------------	----------------------------

Clinical Sensitivity:

Wegener's Granulomatosis	$38/40 = 95.0\%$	<u>95% CI</u> 83.1-91.4
--------------------------	------------------	----------------------------

Microscopic Polyangiitis	$21/38 = 55.3\%$	38.3-71.4
--------------------------	------------------	-----------

* Equivocal results were excluded from calculations.

C. Precision

To determine the precision of the Is-anti-PR-3 IgG Test Kit, four positive and two negative sera were assayed ten times each in three different runs at two different sites. The intra- and interassay precision obtained at each site is shown in Tables 4 and 5.

TABLE 4 : Site #1 - Intra-Assay and Interassay Precision

SERUM	INTRA-ASSAY RUN 1		INTRA-ASSAY RUN 2		INTRA-ASSAY RUN 3		INTERASSAY	
	MEAN EU/ml	CV%	MEAN EU/ml	CV%	MEAN EU/ml	CV%	MEAN EU/ml	CV%
A (POS)	43.3	12.60	43.7	7.34	35.5	9.71	40.8	13.65
B (POS)	65.1	12.86	55.5	8.10	48.8	9.35	56.5	15.86
C (POS)	20.9	14.19	17.8	11.46	13.2	12.18	17.3	22.56
D (POS)	30.6	5.94	27.7	8.11	27.1	6.74	28.4	8.60
E (NEG)	0.5	20.18	0.3	33.99	0.2	57.92	0.3	50.84
F (NEG)	0.5	15.17	0.4	38.33	0.3	43.90	0.4	38.89

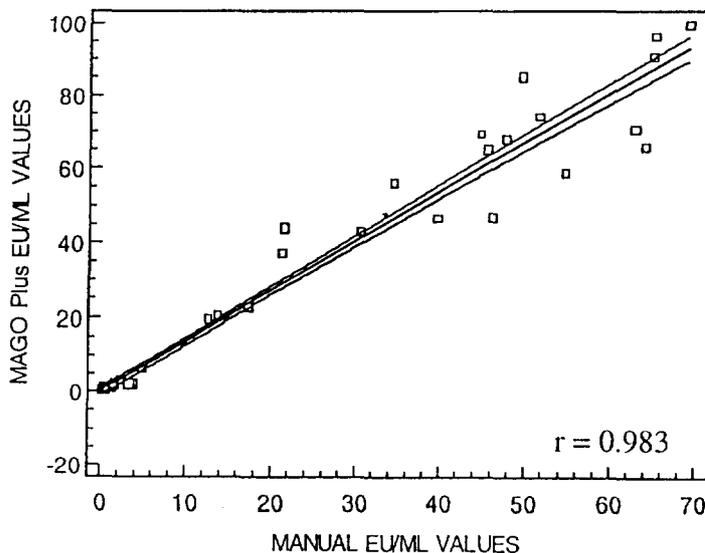
TABLE 5 : Site #2- Intra-Assay and Interassay Precision

SERUM	INTRA-ASSAY RUN 1		INTRA-ASSAY RUN 2		INTRA-ASSAY RUN 3		INTERASSAY	
	MEAN EU/ml	CV%	MEAN EU/ml	CV%	MEAN EU/ml	CV%	MEAN EU/ml	CV%
A (POS)	44.8	8.70	46.4	5.91	48.5	8.01	46.6	8.07
B (POS)	67.3	5.27	66.1	7.52	68.7	8.15	67.4	7.04
C (POS)	15.2	12.28	27.4	8.18	16.4	10.52	19.7	29.87
D (POS)	26.1	10.71	35.1	5.10	28.2	7.80	29.8	15.11
E (NEG)	0.5	29.22	0.4	16.64	0.4	20.20	0.4	31.44
F (NEG)	0.5	21.74	0.5	17.89	0.5	12.95	0.5	17.26

D. Correlation of Manual and MAGO Plus Results

The Is-anti-PR-3 IgG Test Kit has been developed for automated as well as manual use. To demonstrate the equivalence of the manual and MAGO Plus procedures, the results of 90 serum samples tested by both methods were plotted. A scattergram and regression line of the results obtained with 95% confidence intervals is shown in Figure 3. The data indicate good correlation with a Pearson Correlation Coefficient of 0.983.

FIGURE 3 : Manual and MAGO Plus Result Correlation



E. MAGO Plus Precision

The precision of the Is-anti-PR-3 IgG Test Kit when performed on the MAGO Plus Automated EIA Processor was determined by assaying six sera ten times each in three different runs. Table 6 shows the intra- and interassay precision obtained using the MAGO Plus Automated EIA Processor.

TABLE 6 : Site #2- Intra-Assay and Interassay Precision - MAGO Plus

SERUM	INTRA-ASSAY RUN 1		INTRA-ASSAY RUN 2		INTRA-ASSAY RUN 3		INTERASSAY	
	MEAN EU/ml	CV%	MEAN EU/ml	CV%	MEAN EU/ml	CV%	MEAN EU/ml	CV%
A (POS)	34.9	15.47	41.2	11.44	51.8	13.68	42.9	20.55
B (POS)	53.9	17.30	71.9	23.20	80.8	11.19	69.4	23.50
C (POS)	19.5	25.46	25.2	7.59	25.1	7.66	23.3	17.82
D (POS)	32.1	13.83	36.0	5.40	32.7	8.37	33.6	10.61
E (NEG)	0.4	78.26	0.5	27.43	0.5	24.00	0.4	44.80
F (NEG)	0.7	74.05	0.7	23.22	0.7	22.81	0.7	44.89



DEPARTMENT OF HEALTH & HUMAN SERVICES

Public Health Service

NOV 18 1998

Food and Drug Administration
2098 Gaither Road
Rockville MD 20850

Mr. Norman Jenkins
President
Columbia Bioscience, Inc.
8775 M Centre Park Drive, #559
Columbia, Maryland 21045

Re: K983740
Trade Name: Is-Anti PR-3 IgG ELISA Test System
Regulatory Class: II
Product Code: MOB
Dated: October 21, 1998
Received: October 22, 1998

Dear Mr. Jenkins:

We have reviewed your Section 510(k) notification of intent to market the device referenced above and we have determined the device is substantially equivalent (for the indications for use stated in the enclosure) to legally marketed predicate devices marketed in interstate commerce prior to May 28, 1976, the enactment date of the Medical Device Amendments, or to devices that have been reclassified in accordance with the provisions of the Federal Food, Drug, and Cosmetic Act (Act). You may, therefore, market the device, subject to the general controls provisions of the Act. The general controls provisions of the Act include requirements for annual registration, listing of devices, good manufacturing practice, labeling, and prohibitions against misbranding and adulteration.

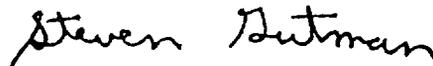
If your device is classified (see above) into either class II (Special Controls) or class III (Pre-market Approval), it may be subject to such additional controls. Existing major regulations affecting your device can be found in the Code of Federal Regulations, Title 21, Parts 800 to 895. A substantially equivalent determination assumes compliance with the Current Good Manufacturing Practice requirements, as set forth in the Quality System Regulation (QS) for Medical Devices: General regulation (21 CFR Part 820) and that, through periodic QS inspections, the Food and Drug Administration (FDA) will verify such assumptions. Failure to comply with the GMP regulation may result in regulatory action. In addition, FDA may publish further announcements concerning your device in the Federal Register. Please note: this response to your premarket notification submission does not affect any obligation you might have under sections 531 through 542 of the Act for devices under the Electronic Product Radiation Control provisions, or other Federal laws or regulations.

Page 2

This letter will allow you to begin marketing your device as described in your 510(k) premarket notification. The FDA finding of substantial equivalence of your device to a legally marketed predicate device results in a classification for your device and thus, permits your device to proceed to the market.

If you desire specific advice for your device on our labeling regulation (21 CFR Part 801 and additionally 809.10 for in vitro diagnostic devices), please contact the Office of Compliance at (301) 594-4588. Additionally, for questions on the promotion and advertising of your device, please contact the Office of Compliance at (301) 594-4639. Also, please note the regulation entitled, "Misbranding by reference to premarket notification" (21 CFR 807.97). Other general information on your responsibilities under the Act may be obtained from the Division of Small Manufacturers Assistance at its toll-free number (800) 638-2041 or (301) 443-6597, or at its internet address "<http://www.fda.gov/cdrh/dsma/dsmamain.html>".

Sincerely yours,



Steven I. Gutman, M.D., M.B.A.
Director
Division of Clinical
Laboratory Devices
Office of Device Evaluation
Center for Devices and
Radiological Health

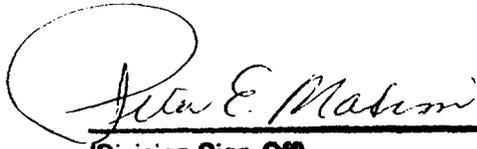
Enclosure

K983740

510(k) Number: ~~Not Known~~

Device Name: ~~PR-3~~ PR-3 IgG ELISA

Indications For Use: The ~~PR-3~~ PR-3 IgG kit is an Enzyme-Linked Immunosorbent Assay (ELISA) For the qualitative detection and semi-quantitation of antibodies against the Proteinase-3 (PR-3) antigen in serum as an aid in the diagnosis of Wegener's granulomatosis. The test can be performed either manually or in conjunction with the Mago Plus automated EIA processor.


(Division Sign-Off)
Division of Clinical Laboratory Devices K983740
510(k) Number _____

PLEASE DO NOT WRITE BELOW THIS LINE-CONTINUE ON ANOTHER PAGE IF NEEDED)

Concurrence of CDRH, Office of Device Evaluation (ODE)

Prescription Use
(Per 21 CFR 801.109)

OR

Over-The Counter Use _____
(Optional Format 1-2-96)