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**Overview****Useful For**

Detecting low level (minimal residual disease) myeloma cells after therapy to confirm remission has been achieved

**Highlights**

This is a high-sensitivity flow cytometry test for detection of minimal residual myeloma cells, post treatment.

It uses adopted EuroFlow guidelines and Cytognos software.

It has a sensitivity of  $10^{-5}$  or better, depending on the antigenic profile of abnormal plasma cells.

**Method Name**

Only orderable as a reflex. For more information see MSMRD / Myeloma Stratification and Risk-Adapted Therapy with Reflex to Minimal Residual Disease, Bone Marrow

Immunophenotyping

**NY State Available**

Yes

**Specimen****Specimen Type**

Bone Marrow

**Specimen Required**

Only orderable as a reflex. For more information see MSMRD / Myeloma Stratification and Risk-Adapted Therapy with Reflex to Minimal Residual Disease, Bone Marrow

**Specimen Type:** Redirected bone marrow

**Container/Tube:**

**Preferred:** Yellow top (ACD solution A or B)

**Acceptable:** Lavender top (EDTA)

**Specimen Volume:** 4 mL

**Specimen Minimum Volume**

2 mL

**Reject Due To**

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All specimens will be evaluated at Mayo Clinic Laboratories for test suitability.

**Specimen Stability Information**

Specimen Type	Temperature	Time	Special Container
Bone Marrow	Ambient (preferred)	72 hours	
	Refrigerated	72 hours	

**Clinical & Interpretive****Clinical Information**

Multiple myeloma is an incurable malignant neoplasm of plasma cells. One of the best prognostic factors in multiple myeloma is the level of minimal residual disease post chemotherapy or autologous stem cell transplantation. The greater depth of the response (less malignant cells present), the longer time to progression and overall survival.(1)

**Reference Values**

Only orderable as a reflex. For more information see MSMRD / Myeloma Stratification and Risk-Adapted Therapy with Reflex to Minimal Residual Disease, Bone Marrow

An interpretive report will be provided.

**Interpretation**

The interpretation of the test is done by evaluating automated and manually gated populations to isolate abnormal plasma cells. If there is an abnormal plasma cell population (cluster of 20 cells or more), then the result is minimal residual disease (MRD)-positive, with the percentage of abnormal plasma cells out of total analyzed events. If no abnormal population is found, then the result will be interpreted as MRD-negative.

This test will be processed as a laboratory consultation. An interpretation of the immunophenotypic findings and correlation with the previous patient history will be provided by a hematopathologist for every case.

**Cautions**

There are situations in which current gating strategies are insufficient to identify abnormal plasma cells. This can occur if the abnormal plasma cells do not phenotypically differ from normal plasma cells. In addition, in patients who have undergone therapeutic antibody treatment (anti-CD38, for example), decreased antigen expression on plasma cells may interfere with the gating strategy.

**Clinical Reference**

1. Martinez-Lopez J, Lahuerta JJ, Pepin F, et al. Prognostic value of deep sequencing method for minimal residual disease detection in multiple myeloma. *Blood*. 2014 May;123(20):3073-3079
2. Rawstron AC, Child JA, de Tute RM, et al. Minimal residual disease assessed by multiparameter flow cytometry in multiple myeloma: impact on outcome in the medical research council myeloma IX Study. *J Clin Oncol*. 2013;31(20):2540-2547
3. Roschewski M, Stetler-Stevenson M, Yuan C, et al. Minimal residual disease: What are the minimum requirements? *J Clin Oncol*. 2014 Feb 10;32(5):475-476

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4. Stetler-Stevenson M, Paiva B, Stoolman L, et al. Consensus guidelines for myeloma minimal residual disease sample staining and data acquisition. *Cytometry B Clin Cytom.* 2016;90(1):26-30 doi: 10.1002/cyto.b.21249
  5. Callander NS, Baljevic M, Adekola K, et al. NCCN Guidelines Insights: Multiple Myeloma, Version 3.2022. *J Natl Compr Canc Netw.* 2022;20(1):8-19. doi:10.6004/jnccn.2022.0002

## Performance

### Method Description

Flow cytometric immunophenotyping for minimal residual disease (MRD) of bone marrow is performed using the following antibodies:

Tube 1: CD138, CD27, CD38, CD56, CD45, CD19, CD117, and CD81.

Tube 2: CD138, CD27, CD38, CD56, CD45, CD19, cyKappa, and cyLambda.

Abnormal plasma cell populations are detected through demonstrating CD38 (multi-epitope) and CD138 positivity along with immunoglobulin light chain restriction (ie, the presence of either predominately kappa or lambda immunoglobulin light chains) and abnormality of CD56, CD117, CD27, CD81, CD19 and/or CD45 expression.

The sensitivity of this assay is conservatively estimated to be 0.001% ( $1 \times 10^{-5}$ ) with a minimum number of  $2 \times 10^{-6}$  total events collected, and an abnormal plasma cell immunophenotype detected in a cluster of at least 20 cells and can be as high as 0.0002% ( $2 \times 10^{-6}$ ). The sensitivity of the assay will be lower in samples with less than  $2 \times 10^{-6}$  total events acquired. The validated limit of detection (sensitivity) meets the current National Comprehensive Cancer Network, International Myeloma Working Group, and EuroFlow guidelines for MRD assessment by flow cytometry in multiple myeloma. The percentage of clonal plasma cells estimated by flow cytometry is affected by specimen processing and antigen loss with specimen aging. MRD reporting is affected by sample volume and cellularity. (Unpublished Mayo method)

### PDF Report

No

### Day(s) Performed

Preanalytical processing: Monday through Saturday

Results reported: Monday through Friday

### Report Available

2 to 4 days

### Specimen Retention Time

14 days

### Performing Laboratory Location

Rochester

## Fees & Codes

### Fees

- Authorized users can sign in to [Test Prices](#) for detailed fee information.
- Clients without access to Test Prices can contact [Customer Service](#) 24 hours a day, seven days a week.
- Prospective clients should contact their account representative. For assistance, contact [Customer Service](#).

### Test Classification

This test was developed and its performance characteristics determined by Mayo Clinic in a manner consistent with CLIA requirements. It has not been cleared or approved by the US Food and Drug Administration.

### CPT Code Information

88184-Flow Cytometry; first cell surface, cytoplasmic or nuclear marker

88185 x 9-Flow Cytometry; additional cell surface, cytoplasmic or nuclear marker

88188-Flow Cytometry Interpretation, 9 to 15 Markers

### LOINC® Information

Test ID	Test Order Name	Order LOINC® Value
MRDMR	Multiple Myeloma MRD by Flow, BM	93022-2

Result ID	Test Result Name	Result LOINC® Value
CK146	% Minimal Residual Disease (MRD)	93021-4
CK147	% Normal Plasma Cells (of total PC)	93020-6
CK148	Non-Aggregate Events	38257-2
CK149	Total Plasma Cell Events	93019-8
CK150	Poly PC Events	93018-0
CK151	Abnormal PC Events	93017-2
615796	% B-cell Precursors	101131-1
615797	% Mast Cells	101130-3
616082	Validated Assay Sensitivity	101129-5
616083	Lower Limit of Quantitation (LLOQ)	87706-8
615798	Patient / Sample Theoretical LOQ	101128-7
614590	Final Diagnosis	74226-2