Myeloperoxidase (MPO)

CPT Code 83876 Sample Type EDTA Plasma Order Code C133 Tube Type Lavender Top

MPO levels are associated with an increased risk for:

- Cardiovascular disease
- Myocardial infarction

MPO levels may be measured in:

- Asymptomatic individuals
- Individuals at risk for pre-diabetes/diabetes
- Individuals with CAD

Description

MPO is a white blood cell-derived inflammatory enzyme that measures disease activity from the luminal aspect of the arterial wall.

Briefly, when the artery wall is damaged, or inflamed, MPO is released by invading macrophages where it accumulates¹. MPO mediates the vascular inflammation that propagates plaque formation² and activates protease cascades that are linked to plaque vulnerability³. White blood cell activation in the bloodstream, in response to luminal injury of the artery wall including fissures, erosions or a degrading collagen cap, leads to MPO release in the bloodstream. This combination of detrimental effects demonstrates that MPO is actively involved in the progression of atherosclerosis. The Cleveland HeartLab MPO test measures free MPO in the bloodstream.

Clinical Use

The MPO test may be performed on individuals at intermediate or high risk for developing coronary heart disease who are any age with at least two major risk factors, those \geq 65 years of age with one major risk factor, smokers, those with a fasting blood glucose of \geq 100 mg/ dL, or those who have metabolic syndrome.

Clinical Significance

- Elevated MPO levels predict the risk of heart disease in subgroups otherwise associated with low risk^{4,5}.
- Elevated MPO levels independently predict the risk of future cardiovascular events in patients presenting with an acute coronary syndrome^{6,7}.
- Individuals with elevated MPO levels are more than 2x as likely to experience cardiovascular mortality⁸.
- MPO enhances cardiovascular risk prediction when used independently or alongside standard biomarker testing such as hsCRP⁸.
- MPO levels are not likely to be elevated due to chronic infections or rheumatologic disorders due to the fact that free MPO in the blood is a **specific marker** of vascular inflammation and vulnerable plaque/erosions/fissures.

 The p-ANCA test (anti-MPO antibody test) is not the same as the MPO test performed by Cleveland HeartLab. The p-ANCA test primarily measures the amount of antibodies directed toward the MPO protein whereas the MPO test performed by Cleveland HeartLab directly measures the amount of MPO protein. The p-ANCA test is useful for identifying systemic inflammation and vasculitis. In contrast, the MPO test performed by Cleveland HeartLab is useful for identifying cardiovascular risk.

Sample Type

The MPO test should be performed on an EDTA plasma sample.

Commercial Insurance or Medicare Coverage

Coverage guidelines, also known as NCD (National Coverage Determination) or LCD (Local Coverage Determination) have been established or posted by CMS (Medicare & Medicaid). Guidelines should be reviewed for coverage and limitations. Limited information has been provided by the majority of the larger carriers (Aetna, United HealthCare, Cigna, Blues).

Understanding Medical Necessity

The following ICD-9 codes for MPO are listed as a convenience for the ordering practitioner. The ordering practitioner should report the diagnosis code that best describes the reason for performing the test and provide the 4th and 5th ICD-9 digit as appropriate.

Diagnosis	Diagnosis Code
Diabetes Mellitus Type II or Unspecified, Not Stated as Uncontrolled	250.00
Diabetes Mellitus Type II or Unspecified, Uncontrolled	250.02
Pure Hypercholesterolemia	272.0
Mixed Hyperlipidemia	272.2
Other and Unspecified Hyperlipidemia	272.4
Benign Essential Hypertension	401.1
Unspecified Essential Hypertension	401.9
Coronary Atherosclerosis of Unspecified Type of Vessel, Native or Graft	414.00
Coronary Atherosclerosis of Native Coronary Artery	414.01
Other Abnormal Blood Chemistry	790.6



RELATIVE RISK

MPO (pmol/L)

<420 Low

420 - 479 Moderate

≥480 High

Treatment Considerations

These treatment considerations are for educational purposes only. Specific treatment plans should be provided and reviewed by the treating practitioner.

✓ Assess LDL-C levels.

• If not at goal, consider lipid-lowering therapy, ideally with a statin-based regimen if not contraindicated.

✓ Assess risk for pre-diabetes/diabetes.

- If abnormal fasting glucose or oral glucose tolerance test, consider PPAR agonists, metformin or DPP-IV inhibitor if not contraindicated.
- ✓ Assess the presence of coronary artery disease (CAD) with imaging techniques such as carotid intima media thickness testing (CIMT) or coronary artery calcium scoring.
 - Consider aspirin therapy if not contraindicated.
- Consider clopidogrel if history of CAD (i.e., myocardial infarction or revascularization) and/or a history of cerebrovascular disease (i.e., TIA or stroke).

✓ Assess dental health (periodontal disease).

 Refer to dentist to identify gum disease.
 NOTE: Poor dental health may cause significant inflammation and is associated with the presence of atherosclerosis⁹.

✓ Assess HDL-C levels.

- If not at goal, consider niacin or fenofibrate therapy.
- Assess CoQ10 levels as recent evidence suggests that low ApoA1 and/or HDL-C levels are associated with low CoQ10 levels¹⁰.

✓ Assess smoking habits.

NOTE: Smoking cessation is essential as individuals who smoke are at increased risk of heart disease and blood clots.

✓ Assess blood pressure.

• If not at goal, consider initiating, or titrating, anti-hypertensive therapy.

NOTE: An elevated blood pressure may contribute to endothelial dysfunction and coronary disease formation.

✓ Assess lifestyle habits.

- Consider diet/exercise/weight reduction efforts if appropriate.
- ✓ Assess the presence of inflammatory conditions such as Crohn's disease, rheumatoid arthritis (RA) and systemic lupus erythematosus (SLE).

NOTE: Chronic inflammatory diseases may exhibit elevated MPO values due to increased vascular disease associated with these conditions. For example, RA is associated with a 5x increased risk for myocardial infarction¹¹.

✓ Assess the presence of vasculitis.

NOTE: MPO values may be elevated in individuals with vasculitis as it is characterized by increased vascular inflammation.

✓ Assess the presence of bone marrow dyscrasias.

NOTE: MPO values may be elevated in individuals with chronic lymphocytic leukemia or other leukemias, that cause increased white blood cell destruction.

✓ Assess level of exercise.

NOTE: MPO values may be elevated in marathon runners¹² and extreme athletes and may identify those with increased oxidative stress and basal levels of inflammation.

If asymptomatic, with all of the above factors ruled out, an elevated MPO value may in fact be the patient's baseline. MPO levels should be monitored every 3-6 months.

References

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