Overview

Useful For
Distinguishing large deletional hereditary persistence of fetal hemoglobin from other conditions with increased percentage of fetal hemoglobin (Hb F)

Determining the distribution of Hb F within red blood cells

Method Name
Only orderable as a reflex. For more information see:

- HAEV1 / Hemolytic Anemia Evaluation, Blood
- HBEL1 / Hemoglobin Electrophoresis Evaluation, Blood
- MEV1 / Methemoglobinemia Evaluation, Blood
- REVE1 / Erythrocytosis Evaluation, Blood
- THEV1 / Thalassemia and Hemoglobinopathy Evaluation, Blood

Flow Cytometry

NY State Available
Yes

Specimen

Specimen Type
Whole Blood EDTA

Ordering Guidance
This test is for hereditary persistence of fetal hemoglobin only. For testing for possible fetal-maternal bleed, see FMB / Fetomaternal Bleed, Flow Cytometry, Blood.

Specimen Required
Only orderable as a reflex. For more information see:

- HAEV1 / Hemolytic Anemia Evaluation, Blood
- HBEL1 / Hemoglobin Electrophoresis Evaluation, Blood
- MEV1 / Methemoglobinemia Evaluation, Blood
- REVE1 / Erythrocytosis Evaluation, Blood
- THEV1 / Thalassemia and Hemoglobinopathy Evaluation, Blood

Reject Due To
**Test Definition: HPFH**

**Hb F Distribution, B**

<table>
<thead>
<tr>
<th>Gross hemolysis</th>
<th>Reject</th>
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<tbody>
<tr>
<td>Gross lipemia</td>
<td>OK</td>
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**Specimen Stability Information**

<table>
<thead>
<tr>
<th>Specimen Type</th>
<th>Temperature</th>
<th>Time</th>
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<tbody>
<tr>
<td>Whole Blood EDTA</td>
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**Clinical and Interpretive**

**Clinical Information**

More than 75% of the hemoglobin of the newborn is hemoglobin (Hb) F; it diminishes over a period of several months to adult levels, reducing to less than 2% by age 1 and less than 1% by age 2.

Hb F may constitute 90% of the total Hb in patients with beta-thalassemia major or other combinations of beta thalassemia and fetal Hb (hereditary persistence of fetal hemoglobin: HPFH) variants.

Hb F is often mildly to moderately elevated in sickle cell disease, aplastic anemia, acute leukemia, and myeloproliferative disorders such as juvenile myelomonocytic leukemia (JMML), hereditary spherocytosis, and alpha-thalassemia minor. It is commonly increased in hemoglobinopathies associated with hemolysis. Hb F increases to as high as 10% during normal pregnancy. Hb F is also increased due to medications such as hydroxyurea, decitabine, and lenalidomide. Elevation in Hb F has been cited as a discriminator between Diamond-Blackfan congenital pure red cell aplasia (elevated) and transient erythroblastopenia of childhood (normal), but whether this simply reflects the chronicity of anemia inherent to the former condition rather than a specific finding is unclear.

In the common (large deletional) form of the genetic trait HPFH, all of the erythrocytes contain Hb F. When tested by flow cytometry using specificity for Hb F, these HPFH cases display a homocellular distribution pattern of Hb F within the red cell population. Other causes of increased Hb F including delta beta thalassemia, hydroxyurea, and some nondeletional HPFH variants typically display a heterocellular distribution of Hb F within the erythrocytes, reflecting disparate populations of F cells and cells lacking Hb F. Quantification of Hb F percentage should be determined prior to flow cytometry of Hb F red cell distribution to establish the appropriateness of this test. The flow cytometry analysis of elevated Hb F levels is useful when Hb F percentage is 15% to 35% and the clinical differential diagnosis includes large deletional HPFH. Hb F percentages below 15% are likely not due to large deletional HPFH, and causes of Hb F percentages above 35% are better confirmed by molecular and family studies.

**Reference Values**

Only orderable as a reflex. For more information see:

- HAEV1 / Hemolytic Anemia Evaluation, Blood
- HBEL1 / Hemoglobin Electrophoresis Evaluation, Blood
- MEV1 / Methemoglobinemia Evaluation, Blood
- REVE1 / Erythrocytosis Evaluation, Blood
- THEV1 / Thalassemia and Hemoglobinopathy Evaluation, Blood
Test Definition: HPFH
Hb F Distribution, B

Reported as heterocellular or homocellular

Interpretation
Homocellular distribution of fetal hemoglobin (Hb) is found in large deletional hereditary persistence of fetal Hb (HPFH).

Heterocellular distribution is found in delta beta thalassemia, medication induced, and other causes of increased Hb F.

Cautions
When hemoglobin (Hb) F values are above 35%, most specimens show a homocellular pattern; this does not necessarily indicate hereditary persistence of fetal Hb (HPFH). Clinical correlation is needed.

Clinical Reference

Performance

Method Description
This assay uses a flow cytometric method with a monoclonal antibody to hemoglobin (Hb) F. Specimens are analyzed by single-color flow cytometry using fluorescein anti-Hb F. In normal adults, a single peak is seen with minimal fluorescence, which corresponds to Hb A. In neonates, a single peak with bright fluorescence is seen, which corresponds to Hb F. In cases of hereditary persistence of fetal Hb (HPFH) only, a single peak is observed, which has a fluorescence intensity intermediate between the normal Hb A and Hb F peaks. This pattern corresponds to the homocellular (pancellular) pattern obtained by the Kleihauer-Betke (K-B) method. In contrast, specimens from infants, transfused neonates, and cases of beta-thalassemia or delta/beta-thalassemia show both Hb A and Hb F peaks, corresponding to the heterocellular pattern of the K-B method. In patients with Hb S/HPFH, a single peak was observed in contrast to patients with homozygous S in which 2 peaks were observed.(Hoyer JD, Penz CS, Fairbanks VF, Hanson CA, Katzmann JA: Flow cytometric measurement of Hb F in RBC's: Diagnostic usefulness in the distinction of hereditary persistence of fetal hemoglobin (HPFH) and Hb S-HPFH from other conditions with elevated levels of Hb F. Am J Clin Path. 2002; 117(6): 857-863; Stephens AD, Afgangstiniotis M, Baysal E, et al: ICSH recommendations for the measurement of Haemoglobin F Int J Lab Hematol. 2012; 34(1):14-20)

PDF Report
No

Day(s) Performed
Monday through Saturday

**Report Available**
2 to 5 days

**Specimen Retention Time**
1 week

**Performing Laboratory Location**
Rochester

### Fees and 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 Regional Manager. For assistance, contact [Customer Service](#).

**Test Classification**
This test has been modified from the manufacturer's instructions. Its performance characteristics were determined by Mayo Clinic in a manner consistent with CLIA requirements. This test has not been cleared or approved by the US Food and Drug Administration.

**CPT Code Information**
88184

**LOINC® Information**

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