

# Diagnosis/Monitoring of Glioblastoma & Acute Myeloid Leukemia by (D)-2-Hydroxyglutarate Test

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## Technology description

Diagnosis and monitoring of IDH1/IDH2/IDH3 dependent diseases such as: Glioblastomas, astrocytoma, oligodendrogliomas, oligoastrocytoma, acute myeloid leukaemia (AML), chondrosarcoma, intrahepatic cholangiocarcinoma, angioimmunoblastic T cell lymphoma. The technology describes a patented test, which is a simple and robust enzymatic assay with a readout in 3 hours. The test is suitable for 96-/384-well format, less expensive/time-consuming and high-throughput possible in opposite to established GC-MS test.

## Solutions

Researchers from DKFZ and University Hospital of Heidelberg developed a test for detecting D2HG in diverse samples by measuring the production of the reduced state of the dye. The technique can be used for diagnosis but in addition monitoring a D2HG-associated disease of a patient.

Figure

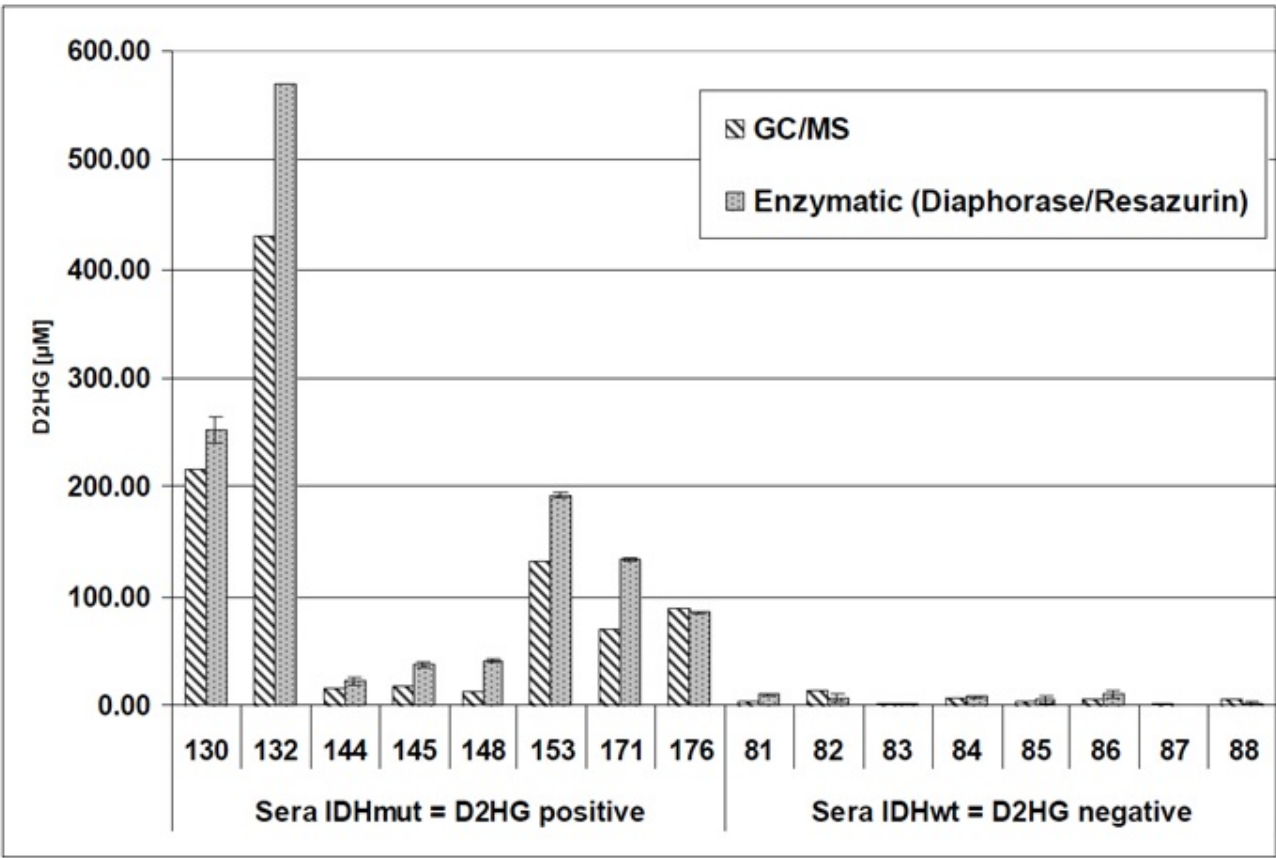


Figure 1: D2HG was diluted in water, blood serum, and urine to get a standard curve (0-375 pmol = 0-15  $\mu\text{M}$ ). Samples were prepared according to the protocol in Figure 2 and assayed with the diaphorase/ resazurin read-out. In the graph, the relative fluorescence (RFU) is plotted against the D2HG concentration

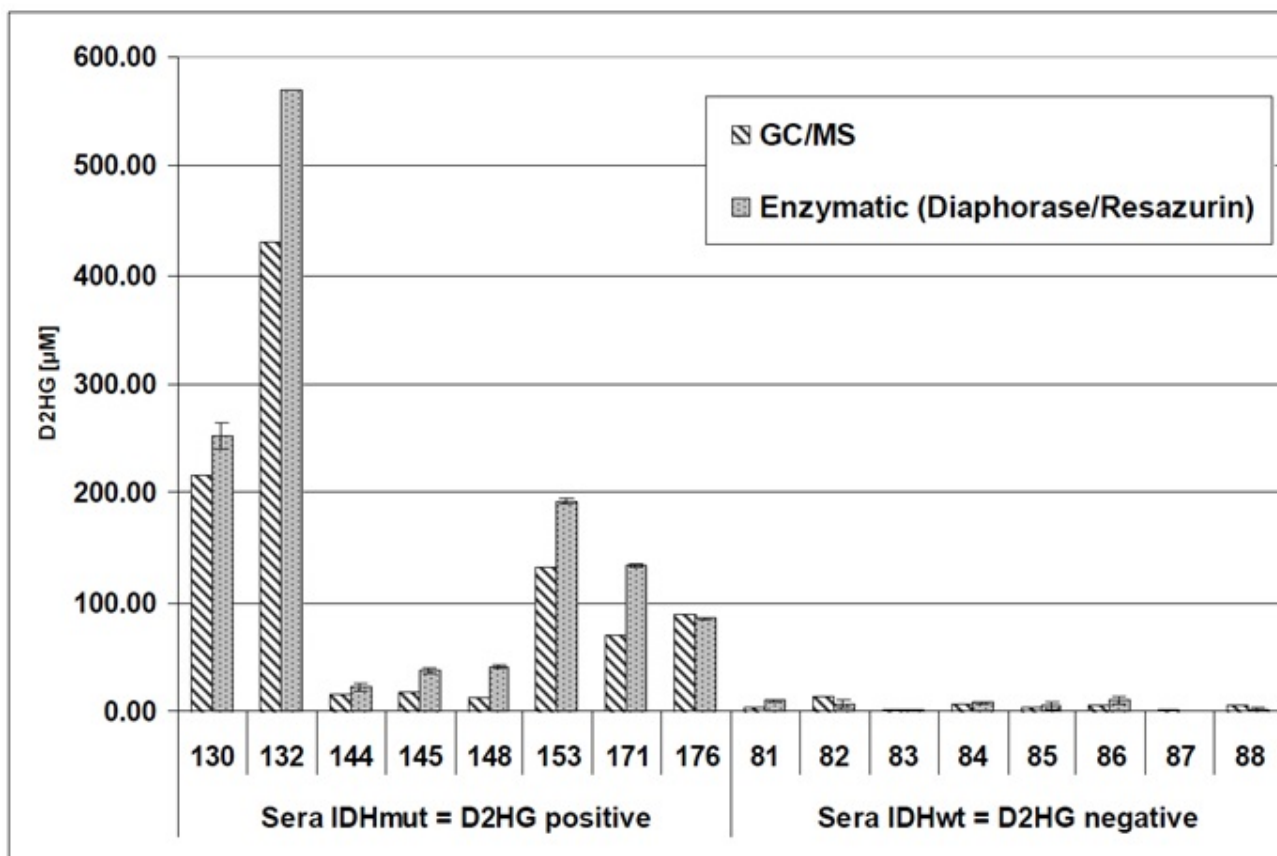


Figure 2: Measurement of sera of patients suffering from acute myeloid leukemia (AML). These patients carry an IDH1 mutation which results in the production of D2HG. Comparison between gas chromatography-mass spectrometry (GC-MS) (single determination) and enzyme assay of the invention using the diaphorase/ resazurin read-out (in triplicate). In comparison, sera of healthy persons who did not carry an IDH1 mutation have been tested. These samples were D2HG negative.

## References

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- "Pan- mutant IDH1 inhibitor BAY 1436032 for effective treatment of IDH1 mutant astrocytoma in vivo." In Acta Neuropathol. 2017 Jan 25. PMID: 28124097 by Stefan Pusch et al.. See at: [www.ncbi.nlm.nih.gov/pubmed/](http://www.ncbi.nlm.nih.gov/pubmed/)
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## Application area

Development and distribution of a simple and robust enzymatic assays for the specific determination of the D2HG. The readout is available already in about 2 to 3 hours. Moreover, it is suitable for 96-well format and can be even further miniaturized to the 1536-well format, thereby allowing for the parallel analysis (high-throughput) of numerous samples at the same time. Beside an already granted license for research use only, we now seek a licensee for the DIAGNOSTIC field.

## Advantages

Diagnosis and monitoring of IDH1/IDH2/IDH3 dependent diseases such as: Glioblastomas, astrocytoma, oligodendrogliomas, oligoastrocytoma, acute myeloid leukemia (AML), chondrosarcoma, intrahepatic cholangiocarcinoma, angioimmunoblastic T cell lymphoma.

Simple and robust enzymatic assay; readout in 3 hours; suitable for 96 up to 1536 well format.

Less expensive/time-consuming and high-throughput possible in opposite to established GC-MS test.

## Institution

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