

“MagQu” TDP-43 IMR Reagent

REF MF-TDP-0060



For *In Vitro* Diagnostic & Professional Use

Intended Use

The “MagQu” TDP-43 IMR Reagent is used to quantitatively measure TDP-43 in human fluid specimen, such as plasma. Use “MagQu” TDP-43 IMR Reagent only with the XacPro-S System (MagQu Co., Ltd.).

Summary & Explanation

The TAR DNA binding protein of 43 kDa (TDP-43) is a ubiquitously expressed nuclear protein with roles in transcription and splicing regulation. A hyper-phosphorylated, ubiquitinated and cleaved form of TDP-43, known as pathologic TDP-43 is the major disease protein in ubiquitin-positive, tau and α -synuclein-negative frontotemporal dementia (FTLD-TDP, previously referred to as FTLD-U) and in amyotrophic lateral sclerosis (ALS). Evidences indicates that TDP-43 can be detected in human plasma and CSF and levels are reportedly elevated in cases of ALS, FTLD and Alzheimer’s disease.^{1, 2}

Principles of Test

The “MagQu” TDP-43 IMR Reagent is designed for rapid quantifying TDP-43 by ImmunoMagnetic Reduction (IMR). We conjugate antibody on the surface of around 50 nm-in-diameter Fe₃O₄ magnetic particles. When the antibodies on the surface bind with TDP-43, the magnetic particles form clusters. Therefore, the ac susceptibility (Xac) of magnetic particles would be reduced in the adding ac magnetic field. By measuring the reduction of Xac, TDP-43 can be easily, rapidly and accurately quantified.⁴

Reagents

“MagQu” TDP-43 IMR Reagent.....4 x 1 mL (64 tests)

Storage Conditions & Stability

Storage reagent at 2 ~ 8 °C (35.6 ~ 46.4 °F).

Please eye check whether there is some precipitation in the tube of “MagQu” TDP-43 IMR Reagent by inverting the tube. Do not use the reagent when it has something precipitated.

Please refer to the detail expiration date on the product label.

CAUTION: Do not use reagents beyond the expiration date.

CAUTION: Do not freeze.

Statement of Warnings



BIOHAZARD

All products or objects that come in contact with human or animal body fluids should be handled, before and after cleaning, as if capable of transmitting infectious diseases. Wear facial protection, gloves, and protective clothing.

Safety Data Sheet is available at www.magqu.com.

1. Do not freeze.
2. Please keep away from events with strong magnetism.
3. For *in vitro* diagnostic use only.
4. For professional use only.
5. Do not use the reagent when it has left from 2 to 8 °C (35.6 to

46.4 °F) environment out over 24 hours.

6. Do not use the reagent when it has something precipitated.
7. Immediately after use reagent should be returned to cold storage (2 to 8 °C).
8. Do not use reagents beyond the expiration date printed on the vial.

Reagent Preparation

1. No preparation is necessary.
2. Please use the “MagQu” TDP-43 IMR Reagents at room temperature (15-30°C).

Specimen Collection & Preparation



BIOHAZARD

All products or objects that come in contact with human or animal body fluids should be handled, before and after cleaning, as if capable of transmitting infectious diseases. Wear facial protection, gloves, and protective clothing.

1. **Collection precautions:** Collect all blood samples by wearing protective equipment and following universal precautions for venipuncture.
2. 6 ~ 10 mL of whole blood into a blood collection tubes prepared with EDTA as an anticoagulant (Lavender Top; K2-EDTA or K3-EDTA tube).
NOTE: Please collecting the whole blood following the manual of blood collection tube from manufacturer.
3. Invert the tube smoothly 5-10 times and make sure the whole blood specimen is mix well with EDTA.
4. Centrifuge the blood collection tubes for 15 minutes at 1,500 ~ 2,500 x g at room temperature to separate the plasma from the blood cells with swing-out(bucket) rotor.
5. After centrifugation, the upper layer of plasma sample can be assayed followed by “Procedure”. The plasma sample must be labeled and deep frozen (-80°C) if it is not freshly used. Avoid repeated freezing and thawing.

CAUTION: Precipitant in plasma may interfere the assay.

CAUTION: Use blood collection tubes contain K2-EDTA or K3-EDTA only. The blood collection tubes of difference brands may have a few difference substances that may influence the assay.

Procedure

Material supplied

“MagQu” TDP-43 IMR Reagent

Materials required but not supplied

Magnetic Immunoassay Analyzer (XacPro-S)

Sample testing tubes

Transfer pipettes

1. Allow reagent and sample to reach room temperature before use.
2. Vortex them for about 5 ± 2 seconds.
3. Add 60 μ L of sample into a clear sample testing tube.
4. Add 60 μ L of “MagQu” TDP-43 IMR Reagent to tube.
5. Vortex them for about 5 ± 2 seconds. The rest of “MagQu” TDP-43 IMR Reagent return to 2~8°C.
6. Insert the sample testing tube into the measuring slot of Magnetic Immunoassay Analyzer (XacPro-S).
NOTE: Step 4 to 6 must be done within 20 minutes.
7. Process the measurement and data analysis according to the user’s manual of Magnetic Immunoassay Analyzer (XacPro-S).
8. We suggest retesting sample if error signal (NaN) is displayed of Magnetic Immunoassay Analyzer (XacPro-S).

Performance Characteristics

Precision

The TDP-43 samples were measured in duplicate, twice per day over 20 days. Two different TDP-43 concentrations were used for the tests. The standard deviations of repeatability and within-lab for various TDP-43 concentrations were obtained:

Item tested	Mean of measured TDP-43 concentrations (pg/mL)	Standard deviation (%CV)	
		Repeatability	Within-Lab
pool 1	0.102	0.003 (3.2)	0.002 (2.3)
pool 2	1.018	0.023 (2.2)	0.016 (1.6)

Precision testing was determined according to CLSI/NCCLS document EP5-A3.

Interference (Specificity)

Plasma can contain interfering substances such as hemoglobin, bilirubin, or intra lipid because of common diseases, such as hemolysis, jaundice or hypertriglyceridemia. Other bio-substances that exist naturally in plasma, such as uric acid, rheumatoid factor, or albumin, are also interfering substances. Other interfering substances include drugs or chemicals in medicine that is used to treat inflammatory diseases, viral and bacterial infections, cancers and cardiovascular disease. The level of TDP-43 in each of these pools was then determined and normalized to the level without the respective substances.

Substance	Amount Added	% Recovery (Spike/control x 100)
Hemoglobin	10000 µg/mL	106.2
Conjugated bilirubin	600 µg/mL	105.2
Intra lipid	30000 µg/mL	102.7
Uric acid	200 µg/mL	102.9
Rheumatoid factor	500 IU/mL	101.2
Albumin	60000 µg/mL	107.9
Acetylsalicylic acid	500 µg/mL	103.4
Ascorbic acid	300 µg/mL	97.1
Ampicillin sodium	1000 µg/mL	101.8
Quetiapine fumarate	100 ng/mL	104.2
Galantamine hydrobromide	90 ng/mL	99.0
Rivastigmine hydrogen tartrate	100 ng/mL	95.7
Donepezil hydrochloride	1000 ng/mL	105.7
Memantine hydrochloride	150 ng/mL	110.7

Interference testing was based on the principle of CLSI/NCCLS document EP7.

Analytical Sensitivity

The "MagQu" TDP-43 IMR reagent has an analytical sensitivity of 0.68 fg/mL.

Analytical Measuring Range (AMR)

The analytical measuring range of the reagent is from 0.001 to 100 pg/mL.

Results

By using XacPro-S, we can get two signals: one is the AC signal before the reaction (Xac_0) and the other is the AC signal after reaction (Xac). Then we can have the IMR (%) through two signals by following function:

$$IMR(\%) = \frac{Xac_0 - Xac}{Xac} \times 100$$

IMR (%), as functions of TDP-43 concentration ϕ_{TDP-43} are explored and are found to follow the logistic function:

$$IMR(\%) = \frac{A - B}{1 + \left(\frac{\phi_{TDP-43}}{\phi_o}\right)^\gamma} + B$$

where A, B, ϕ_o , and γ are fitting parameters. For TDP-43, A = 2.88, B = 4.55, $\phi_o = 4.32$, and $\gamma = 0.57$. The concentration of TDP-43 can be obtained by following the equations, and can be converted from IMR(%) by Main-analyzer.

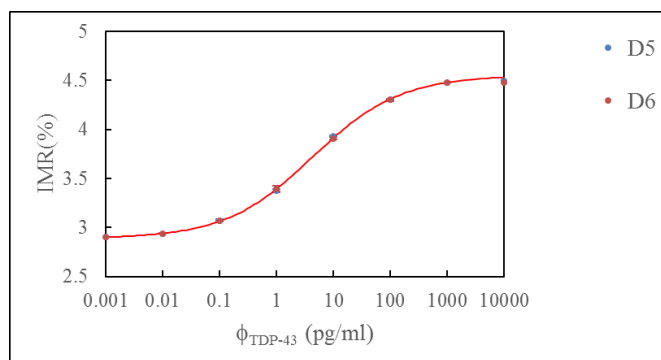


Fig.1 The IMR standard curve of TDP-43
















Limitations

1. The analytical range of reagent is from 0.001 to 100 pg/mL. When the specimen with TDP-43 > 100 pg/mL is to be determined, carry out the following procedures to obtain the accurate concentration. Dilute the specimen, re-assay, and multiply the assayed TDP-43 value by the dilution factor.
2. Reagents should be used before the expiration date printed on the kit label.
3. Data is based upon human plasma sample.
4. Do not use the plasma sample when it has leaved -20°C more than 2 hours or it has something precipitated.
5. Glass testing tubes are single use only.

References

1. Foulds P, McAuley E, Gibbons L, et al. TDP-43 protein in plasma may index TDP-43 brain pathology in Alzheimer's disease and frontotemporal lobar degeneration. *Acta Neuropathol.* 2008;116(2):141-146.
2. Feneberg E, Gray E, Ansoorge O, Talbot K, Turner MR. Towards a TDP-43-Based Biomarker for ALS and FTL. *Mol Neurobiol.* 2018;55(10):7789-7801.

Glossary/symbol definition :

SYMBOL	DESCRIPTION
	Caution, refer to accompanying documents
	Batch code
	Catalogue number
	Content
	Use by Expressed as: CCYY-MM-DD
	Biological risk
	Consult instructions for use.
	Temperature limitation
	Authorized representative in the EU/EC.
	In Vitro diagnostic medical device
	Manufacturer
	Country and date of manufacture
	Do not use if package damaged
	CE MARK = CONFORM WITH EEC DIRECTIVES
	Unique device identifier



Manufacturer

MagQu Co., Ltd.

Rm. 3, 6F., No. 95, Minquan Rd., Xindian Dist., New Taipei City 231625, Taiwan

Tel: +886-2-8667-1897 Fax: +886-2-8667-1809

E-mail: info@magqu.com Website : www.magqu.com

 Authorized representative in the EU/EC

MedNet EC-REP GmbH

Borkstraße 10 48163 Münster Germany

Rev. Jun-07-2022