

Etude Gangliosides MONDOR

Techniques

| | Test 1 | Test 2 | Test 3 | Test 4 |
|--------------------------|----------------------------------|----------------------------------|----------------------------------|-------------------------|
| | Generic Assays | Zentec | D-Tek | Bühlmann |
| | Dot-line | Dot spot | Dot-Line | Elisa |
| Support | membrane | membrane | membrane | Elisa microplate |
| Gangliosides | 11 | 9 | 11 | 5 |
| Sulfatide | yes | yes | yes | no |
| Asialo GM1 | no | no | no | yes |
| Galacto-cerebroside | no | no | no | no |
| Spectrum of glycolipids | | | | |
| | GM1 | GM1 | GM1 | GM1 |
| | GM2 | GM2 | GM2 | GM2 |
| | GM3 | GM3 | GM3 | GD1a |
| | GM4 | GD1a | GM4 | GD1b |
| | GD1a | GD1b | GD1a | GQ1b |
| | GD1b | GD3 | GD1b | |
| | GD2 | GT1a | GD2 | |
| | GD3 | GT1b | GD3 | |
| | GT1a | GQ1b | GT1a | |
| | GT1b | | GT1b | |
| | GQ1b | | GQ1b | |
| Final sera dilution | 1/101 | 1/501 | 1/101 | 1/50 |
| Temperature | + 4° C | + 4° C | + 4° C | + 4° C |
| Serum incubation (h) | 2 | 2 | 2 | 2 |
| Conjugate incubation (h) | 1 | 1 | 1 | 2 |
| Conjugate label | Peroxydase de Raifort | Peroxydase de Raifort | Phosphatase Alcaline | Peroxydase de Raifort |
| Time (h) | 3 | <4 | <4 | 5 |
| Evaluation | Eye reading Semi quantitative | Eye reading Semi quantitative | Eye reading Semi quantitative | Cut-off quantitative |

Critères entre diagnostic clinique et profil des Ac anti-gangliosides

| <u>Tableaux aigus</u> | typique | possible | impossible |
|--|---------------------------|-----------------------------------|------------|
| AMAN (acute axonal motor neuropathy) | IgG GM1 | IgG GD1a/b, IgM GM1/GD1a/b | autres |
| AMAN forme paraparétique | IgG GD1a | IgG GM1, IgM GM1/GD1a/b | autres |
| Syndrome de Guillain-Barré démyélinisant "classique » ou AIDP | Aucun | IgM GM2 | autres |
| Syndrome de Guillain-Barré ataxiant | IgG GD1b | IgG GD3, IgM GD1b/GD3 | autres |
| Déficit pharyngo-cervico-brachial | IgG GT1a | IgG GQ1b | autres |
| Diplégie faciale et paresthésies | Aucun | Aucun | autres |
| Syndrome de Miller-Fisher | IgG GQ1b | IgG GD1b/GT1a, IgM GQ1b/GD1b/GT1a | autres |
| <u>Tableaux chroniques</u> | | | |
| Neuropathie motrice multifocale avec blocs de conduction | IgM GM1 | IgM GD1a | autres |
| Neuropathie sensitivo-motrice multifocale avec blocs | Aucun | Aucun | autres |
| Polyradiculoneuropathie inflammatoire démyélinisante chronique | Aucun | Aucun | autres |
| CANOMAD | IgM GD1b, GD3, GT1b, GQ1b | Aucun | autres |

Critères entre diagnostic clinique et profil des Ac anti-gangliosides

Table 3. Recommended and minimal criteria used in the study for a positive anti-ganglioside antibody finding in a patient serum.

| Clinical diagnosis | Recommended positivity criteria ^{1, 2, 9, 12, 23} | Minimal positivity criteria * |
|---|--|---|
| GBS with ophthalmoplegia | GQ1b and additional GD1a/GT1a/GT1b IgG | GQ1b IgG GT1a/GT1b = atteinte bulbaire ? |
| GBS: AMAN | GMI, GD1a and additional GT1a/ GT1b IgG | GMI and GD1a IgG |
| GBS: AMSAN | GMI, GD1a, GD1b and GT1a/GT1b IgG | GMI, GD1a, and GD1b IgG Atteinte sensitive ? |
| MFS | GT1a and GQ1b IgG | GQ1b IgG |
| GBS post CMV infection | GM2 IgM | GM2 IgM |
| M-IgM chronic motor peripheral neuropathy | GMI and GD1b | High levels of GMI with M-IgM |
| CANOMAD | GD1b, GD3, GT1b, GQ1b IgM ² | High levels of antibodies: GQ1b IgM and one disialylated ganglioside between: GD1b, GD2, GD3, GT1a , GT1b IgM |
| MMN | GMI and GD1b IgM | GMI IgM |

* - Minimal criteria are based on basic profiles including 5 immunodominant gangliosides present on each test, AMAN - acute motor axonal neuropathy, AMSAN - acute motor and sensory axonal neuropathy, CANOMAD - Chronic Ataxic Neuropathy Ophthalmoplegia, IgM-paraprotein cold Agglutinins Disialosyl antibodies, CMV - cytomegalovirus, GBS - Guillain-Barré syndrome, M-IgM - Monoclonal IgM antibodies, MMN - multifocal motor neuropathy with conduction blocks.

Résultats

| | | Electroclinical diagnosis | GanglioCombi, Bühlmann Lab | | Anti-Gangliosid Dots, D-Tek SA | | Dotzen Ganglio Profile, Zentec SA | |
|------------|--------|--|----------------------------|------------|--------------------------------|-----------------------|-----------------------------------|-----------------|
| | | | IgG | IgM | IgG | IgM | IgG | IgM |
| Patient 1 | F, 57y | AMAN | GM1, GD1a, GD1b | | GM1, GD1a, GD1b, GT1b | | GM1, GD1a, GD1b, GT1b | |
| Patient 2 | F, 38y | AMAN | GM1, GD1b | | none | | GM1 | |
| Patient 3 | M, 28y | AMAN (paraplegic form) | GD1a, GQ1b | | GD1a, GT1a | | GM1, GD1a, GD1b, GT1b | |
| Patient 4 | M, 56y | AMAN (paraplegic form) | GD1a | | GD1a | | GD1a | |
| Patient 5 | M, 23y | AMAN (paraplegic form) | GD1a | GD1a | none | | none | GD1a |
| Patient 6 | M, 70y | AMAN (paraplegic form) | GD1a | | GM1 | | GD1a | |
| Patient 7 | F, 59y | Classical demyelinating GBS | | | | GM2 | | GM2, GD1a |
| Patient 8 | F, 32y | Classical demyelinating GBS | | GM2 | | | | GM2 |
| Patient 9 | M, 79y | Classical demyelinating GBS | GD1b | | | | | GM2 |
| Patient 10 | F, 77y | Classical demyelinating GBS | | | | | | |
| Patient 11 | F, 48y | Ataxic GBS | none | GM1 | none | | none | GM1 |
| Patient 12 | F, 57y | Facial diplegia form of GBS | none | | none | | none | |
| Patient 13 | M, 66y | Miller-Fisher form of GBS | GQ1b | | GQ1b, GT1a | GM1, GD1b, GT1a, GQ1b | GQ1b, GT1a | GM1 |
| Patient 14 | M, 75y | Multifocal motor neuropathy with conduction blocks | | GM1, GD1b | | GM1 | | GM1 |
| Patient 15 | F, 56y | Multifocal motor neuropathy with conduction blocks | | GD1a | | none | | none |
| Patient 16 | F, 59y | Multifocal motor neuropathy with conduction blocks | | none | | none | | none |
| Patient 17 | F, 65y | Multifocal motor neuropathy with conduction blocks | | GD1a | | none | | none |
| Patient 18 | M, 74y | CANOMAD | | GD1a | | GD1a, GD1b | | none |
| Patient 19 | F, 70y | CANOMAD | | GD1b, GQ1b | | GD1b | | GD1b, GD3, GQ1b |
| Patient 20 | M, 52y | Chronic inflammatory demyelinating polyneuropathy | GD1b | GD1b, GD1a | | GM2, GD1b, GT1b | | GD1b |
| Patient 21 | M, 71y | Chronic inflammatory demyelinating polyneuropathy | none | | none | | none | |
| Patient 22 | M, 71y | Chronic inflammatory demyelinating polyneuropathy | none | | none | | none | |

in red: expected Anti-ganglioside Antibodies, in black: possible Anti-ganglioside Antibodies, in blue: unexpected Anti-ganglioside Antibodies

GT1a/GT1b = atteinte bulbaire ?

Atteinte du nerf occulo-moteur ?

Atteinte motrice ?

Résultats

| | Bullman | D-Tek | Zentec |
|--|---------|---------|---------|
| Diagnostic + (expected anti-gangliosides Ab) | 10 | 8 | 8 |
| Diagnostic + (possible anti-gangliosides Ab) | 4 | 1 | 4 |
| Diagnostic + (no anti-gangliosides Ab) | 3 | 3 | 3 |
| Erroneous positivities | 4 | 4 | 5 |
| Erroneous negativities | 1 | 5 | 5 |
| Dg sensitivity/ VPP | 85%/89% | 57%/92% | 70%/88% |