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| --- | --- | --- | --- | --- | --- |
| Proband ID | 1 | 2 | 3 | 4 | 5 |
| Genomic coordinates (chr8) | g.101932970G>C | g.101960961C>T | g.101936511G>A | g.101961078C>A | g.101932970G>GAT |
| HGVS coding level (NM\_145690.2) | c.689C>G | c.157G>A | c.434C>T | c.40G​>T | c.688\_689​insAT |
| HGVS protein level | p.Ser230Trp | p.Gly53Arg | p.Ser145Leu | p.Glu14Ter | p.Ser230TyrfsTer44 |
| GERP RS score | 4.60 | 5.64 | 4.84 | 5.30 | 4.60 |
| CADD (v1.4) score | 33 | 32 | 33 | 38 | 35 |
| Zygosity | 1/0 | 1/0 | 1/0 | 1/0 | 1/0 |
| Inheritance | de novo | de novo | de novo | de novo | de novo |
| Gender | Male | Female | Female | Female | Male |
| Age at last examination (years) | 9 | 12 | 17 | 9 | 17 |
| OFC at last evaluation | NR | normal | NR | NR | <-2.5 SD |
| Height | short stature, <5%ile | short stature | -1 SD | +1 SD | 0 SD |
| Weight | <5%ile | low | +2.5 SD |  |  |
| Developmental Delay | yes | yes | yes | yes | yes |
| Motor abilities | motor delay, walked at 4 years | motor delay | mild motor delay | motor delay |  |
| Cognitive abilities |  |  | ID | ID (IQ 55) |  |
| Verbal abilities | almost no speech | almost no speech | NR | speech and language delay | almost no speech |
| Behavior | autistic-like behavior, self-injurious behavior, irritability | NR | NR | obsessive, no concentration, fear, difficulty in making contact | autism, behavioral problems, sleep issues |
| Facial dysmorphism | ptosis, proptosis with long philtrum, thick lips, high forehead, triangular facies, coarse/curly hair, sparse eyebrows | triangular face, mild ptosis | NR | abundant hair, coarse facial features, frontal bossing | NR |
| Skeletal Anomalies | mild scoliosis | NR | NR | NR | NR |
| Cardiac | congenital mild pulmonic stenosis | NR | NR | no | no |
| Seizures | yes | yes, began around age 10-11 years | yes, generalized tonic-clonic epilepsy until age 14 | NR | NR |
| Other | Born at 33 weeks gestation, FTT, GE reflux, G-tube, otitis media with myringotomy tube, hypogammoglobinemia and lymphopenia, hyperkeratosis pilaris, dry skin | feeding difficulties, hyperventilation when anxious | NR | Short fifth digits, sleep issues | Born at 30 weeks gestation due to non-immune hydrops fetalis and HELLP in mother; retinopathy and cataracts |
| Other testing | Negative panels for RASopathies | Negative RASopathy panel | NR | NR | NR |
| Other variants of interest | SCN8A NM\_001177984.2:c.4022\_4023insGA (p.G1341fs) | NR | NR | MAP3K14 NM\_001001671.3:c.522\_523​insC (p.THr175HisfsTer19) | NR |