

Appendix Table 1. Summary of clinical features observed in the 14 South African patients and scored according to the established guidelines.[1]

| | 1 | 2 | 3 | 4 | 5 | 6 | 7 | 8 | 9 | 10 | 11 | 12 | 13 | 14 | |
|--------------------------------------|---|---|---|---|---|---|---|---|---|----|----|----|----|----|-------|
| <i>Growth</i> | | | | | | | | | | | | | | | |
| Short stature | + | . | + | + | + | + | + | + | + | + | + | + | . | + | 12/14 |
| Microcephaly | + | + | + | + | + | - | + | + | + | + | + | + | + | + | 13/14 |
| Prenatal growth retardation (<2 SD) | . | . | + | . | - | . | . | . | . | + | - | . | + | + | 4/14 |
| Postnatal growth retardation (<2 SD) | + | - | + | . | + | + | + | + | + | + | + | + | . | + | 11/14 |
| <i>Craniofacial features</i> | | | | | | | | | | | | | | | |
| Brachycephaly | + | . | - | . | . | . | + | . | . | . | . | + | . | + | 4/14 |
| Low anterior hairline | . | . | + | . | . | + | . | . | . | + | + | . | . | . | 4/14 |
| Arched, thick eyebrows | . | . | + | . | + | + | . | + | . | + | + | . | . | + | 7/14 |
| Synophrys | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 14/14 |
| Long/curly eyelashes | + | + | + | + | + | + | + | + | + | + | + | + | + | + | 14/14 |
| Depressed nasal bridge | + | + | + | + | + | . | . | + | + | + | + | - | + | + | 11/14 |
| Anteverted nostrils | + | + | + | + | . | + | + | + | . | + | . | + | . | + | 10/14 |
| Short nose | . | . | + | + | . | + | . | + | . | + | + | + | . | + | 8/14 |
| Broad nasal tip | . | . | + | . | . | . | . | + | . | + | + | . | . | + | 5/14 |
| Long, smooth philtrum | + | . | - | . | + | . | + | + | + | + | + | + | . | + | 9/14 |

| | | | | | | | | | | | | | | | |
|------------------------------------|---|---|---|---|---|---|---|---|---|---|---|---|---|---|-------|
| Thin upper vermillion | . | . | - | . | . | . | . | + | . | + | . | . | . | + | 3/14 |
| Downturned corners of the mouth | + | . | + | . | + | . | . | - | + | + | . | + | + | . | 7/14 |
| Highly arched palate/cleft | . | . | - | . | + | . | . | . | . | + | + | + | . | + | 5/14 |
| Widely spaced teeth | . | . | + | + | . | . | . | . | + | . | . | - | . | . | 3/14 |
| Micrognathia | . | . | . | . | + | . | . | + | . | + | . | . | . | . | 3/14 |
| Low- set and malformed ears | . | . | + | . | . | + | + | + | . | + | + | + | + | + | 9/14 |
| <i>Trunk and limbs</i> | | | | | | | | | | | | | | | |
| Oligodactyly and adactyly (hands) | + | . | + | . | + | - | . | - | + | + | . | + | + | + | 8/14 |
| Small hands | + | . | + | . | + | . | . | + | + | + | . | + | + | + | 9/14 |
| Proximally placed thumbs | . | . | + | . | - | . | . | + | . | + | . | . | . | + | 4/14 |
| Clinodactyly or short fifth finger | . | . | + | + | - | . | + | + | . | + | + | + | . | + | 8/14 |
| Small feet | . | . | + | + | - | - | . | - | . | . | . | . | . | . | 2/14 |
| Hirsutism | . | + | + | + | + | + | + | + | + | + | + | + | . | + | 12/14 |
| Cardiovascular anomalies | . | . | + | . | + | - | . | . | + | . | + | + | . | + | 6/14 |
| Vertebral anomalies | . | . | + | . | . | . | . | - | . | . | . | . | . | . | 1/14 |
| Diaphragmatic hernia | . | . | - | . | . | . | . | - | . | + | . | . | . | . | 1/14 |
| GOR | . | + | . | . | + | . | . | - | . | . | . | + | . | . | 3/14 |

| | | | | | | | | | | | | | | | |
|---------------------------------------|------|---|-------------|---|------|---|----|----|----|----|----|----|--------------|-------|------|
| Cryptorchidism | . | . | - | . | . | . | . | + | . | . | + | . | . | . | 2/14 |
| <i>Cognition and behaviour</i> | | | | | | | | | | | | | | | |
| Intellectual disability (any degree) | + | + | + | + | + | + | + | + | + | . | . | . | . | . | 9/14 |
| autism spectrum disorder | . | . | . | . | . | . | . | . | . | . | . | . | . | . | . |
| Self-injurious behaviour | . | . | . | . | . | . | . | . | . | . | . | . | . | . | . |
| Stereotypic movements | . | . | . | . | . | . | . | . | . | . | . | . | . | . | . |
| CNS Abnormality | . | . | . | . | . | . | . | . | . | + | + | . | . | . | 2/14 |
| Other | SNHL | | UGT anomaly | | Died | | | | | SB | | | Died 1 month | Renal | |
| Score | 12 | 7 | 13 | 8 | 11 | 7 | 11 | 14 | 11 | 17 | 10 | 13 | 7 | 16 | |
| Cardinal features | 4 | 2 | 3 | 2 | 3 | 2 | 3 | 4 | 3 | 6 | 3 | 4 | 3 | 5 | |
| Class | 1 | 3 | 1 | 3 | 1 | 3 | 1 | 1 | 1 | 1 | 2 | 1 | 3 | 1 | |
| Mutation identified | + | - | + | + | - | + | - | - | + | + | + | - | + | + | 9/14 |

+ assessed and present

- assessed but not present

. not assessed