Eligible patients include those children with severe metabolic, central nervous system, or chromosomal/genetic conditions who meet all of the following criteria:

1) This condition is **progressive** and **not curable**.

<table>
<thead>
<tr>
<th>Progressive?</th>
<th>Yes</th>
<th>No</th>
</tr>
</thead>
<tbody>
<tr>
<td>Not Curable?</td>
<td>Yes</td>
<td>No</td>
</tr>
</tbody>
</table>

2) There is no effective treatment directed towards the pathology of the child’s disease **OR** there is a treatment, but it is failing in this patient.

3) The condition is known to have/likely to have a genetic or metabolic cause.

4) The condition manifests in neurological (CNS) impairment.

5) There is a good probability that a child with this condition will die before their 20s.

Ideally patients will be referred to the study as early in their diagnosis as possible. The child does not have to be at risk of dying in the near-term. Conditions such as “CP”, which are due to a single, identifiable event (e.g., trauma, infection) are not the subject of this study.

### Metabolic Diseases and Central Nervous System degenerative disorders
- Amino acid metabolism disorders (organic acidemias and others)
- Carbohydrate metabolism disorders
  - Lipid metabolism disorders (e.g., peroxisomal diseases, leukodystrophies, sphingolipidoses)
- Mucopolysaccharidoses
- Spinocerebellar degenerative diseases
- Other metabolic disorders (including severe mitochondrial disease)
- Cerebral degenerations manifest in childhood
  - neuronal ceroid lipofuscinoses, Pelizaeus-Merzbacher disease, Alexander disease, Canavan spongy degeneration, Menke disease, Rett syndrome, and subacute sclerosing panencephalitis
- Severe Neurological Impairment NYD with **ongoing deterioration in health**
  - progressive severe CP and mental handicap with seizures, major functional impairments

### Chromosomal diseases and gene conditions
- Chromosomal anomalies, including aneuploid states, i.e., unbalanced rare chromosome abnormalities
- Gene deletions with progressive deterioration

*Representative diseases include: Trisomy 13; Trisomy 18; Severe deletions and translocations NOS*