

**Supplementary Table 5.****Sub-classification of PDCDs:****Currently known, possible and potential molecular etiologies of impaired pyruvate oxidation**

Enzyme/Complex/Function/Pathway	Gene		
	Known	Possible	Potential
<b>Primary-specific PDCD:</b>			
Pyruvate dehydrogenase complex	<i>PDHA1, PDHB, DLAT, PDHX</i>		
Pyruvate dehydrogenase phosphatase	<i>PDP1</i>	<i>PDP2, PDPR</i>	
Pyruvate dehydrogenase kinase	<i>PDK3</i>	<i>PDK1, PDK2, PDK4</i>	
<b>Primary-generalized PDCD:</b>			
Multiple 2-ketoacid Dehydrogenases	<i>DLD</i>		
Thiamine/thiamine pyrophosphate transporters	<i>SLC25A19, SLC19A2, SLC19A3</i>		
Thiamine pyrophosphokinase	<i>TPK1</i>		
Pyruvate carrier (mitochondrial)	<i>MPC1</i>		
Lipoate metabolism	<i>LIAS, LIPT1, LIPT2, SIRT4</i>		
Fe-S Cluster proteins	<i>BOLA3, NFU1, FDX1L, IBA57, ISCA1, ISCA2, ISCU</i>	<i>GLRX5, NUBPL</i>	<i>NFS1, LYRM4</i>
Protease	<i>LONP1</i>		
Regulation of glutathionylation			<i>GLRX2, LRX2</i>
<b>Secondary PDCD:</b>			
Fatty acid oxidation	<i>ECHS1</i>		
Branched-chain amino acid metabolism	<i>ECHS1, HIBCH</i>		
Tricarboxylic acid (TCA) cycle	<i>SUCLA2, SUCLG1</i>	<i>SUCLG2, SUCLA1</i>	
Phosphoenolpyruvate carboxykinase	<i>PCK2, PCK1</i>		
Bioenergetics/PDC regulation/TCA cycle	<i>TAZ</i>		
<b>Other:</b>			
Pyruvate carboxylase	<i>PC</i>		
	<i>GOT2, KMT2B, SLC25A1</i>	<i>GOT2, FBXL4</i>	