

	<b>Light Microscopy</b>	<b>Mitochondria</b>	<b>Z-line</b>	<b>Disarray</b>	<b>Junctions</b>	<b>Autophagy</b>	<b>Myelin bodies</b>	<b>Contracted/relaxed</b>
24 hours	Sparse organisation with a lack of striations and large disruption	Small with damaged cristae, some intact mitochondria	Some ragged	Some disarray	Ok – gap junction + desmosomes visible	Yes – numerous residual bodies visible	None	Mostly contracted (little relaxed regions)
48 hours	Circular bodies with no sarcomeric structure, large disruption	No intact mitochondria	None visible	Complete disarray	Irregular desmosomes – very short and random	Large areas of residual bodies	None	Contracted – hard to tell
72 hours	Irregular shaped nuclei, lack of striations or regularity	Small irregular mitochondria, lack of integrity/structure, extremely damaged cristae	Not visible	Complete disarray, lack of structural integrity	Very long gap junction, fat desmosomes	Widespread residual bodies	None	Contracted
96 hours	Nuclei near outer edge – no structural integrity, lack of sarcomere/ striations	Groups of very small damaged mitochondria – hard to distinguish	Very thick Z-line and misaligned	Complete disarray	Intact very long gap junction, other short desmosomes visible	Large density of residual bodies	None	Contracted

S4 Table. Morphological alterations of EHT under sorafenib (100  $\mu$ M; 24-96 hours), 1<sup>st</sup> column (red heading): light microscopy observations; 2-9<sup>th</sup> columns (blue headings): Electron microscopy observations.

