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(57) Abstract: Provided herein are compositions comprising an endoplasmic/sarcoplasmic reticulum calcium channel inhibitor and a gap junction inhibitor for use in a variety of methods, such as methods for treating or preventing necrosis, methods for treating or preventing cell death, methods for treating or preventing tissue damage, or methods for treating or preventing organ damage. The compositions may also comprise a plasma membrane calcium channel inhibitor and/or a calcium chelator. Also provided are compositions comprising a calpain inhibitor and a cathepsin inhibitor for use in a variety of methods. Also provided are compositions comprising calcium chelators, preferably in relatively high doses, for use in a variety of methods.

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