

Supplementary Table S1

Table S1. Baseline characteristics of donor patients

Variables	Pa1-LCλ	Pa2-LCκ	Pa3-LCκ
Clinical characteristics			
Age, years	81	66	68
Sex	Male	Male	Male
Time to diagnosis, months	38,4	9,6	9,3
Cardiovascular characteristics			
NYHA	III	II	III
Biological variables			
NT-proBNP, pg/mL (reference <1800pg/ml)	7137	620	14647
Troponin T HS, ng/mL (reference <14pg/ml)	51	40	51
Amyloid characteristics			
Light chain subtype	Lambda	Kappa	Kappa
European stage	IIIa	II	IIIb
Medullary plasma cells, %	11	30	18
dFLC, mg/L	217	NA	411
Ratio κ/λ	0.07	NA	14.40
Affected organs	Heart	Heart	Heart
Echocardiographic characteristics			
LVEF, % (normal : >50%)	62	51	43
IVST, mm (normal range : 6-11mm)	20	16	15
GLS, % (normal < -18%)	-10.3	-13.2	-7.1

NYHA: New York Heart Association; NT-proBNP: N-terminal pro-B-type natriuretic peptide; dFLC: free light-chain difference (dFLC = kappa-lambda); LVEF: left ventricular ejection fraction; IVST: interventricular septum thickness; GLS: global longitudinal strain,

Supplementary Figure S1

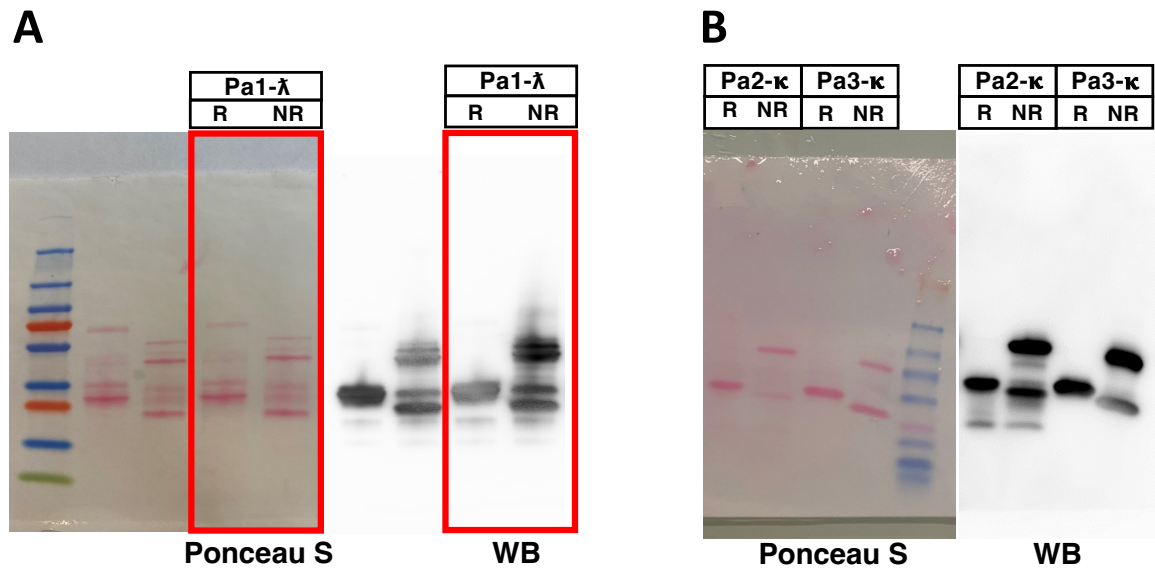


Figure S1. Full-length membranes corresponding to light chain (LC) purification data shown in Figure 1B and C. **Ponceau S**, staining showing total sample proteins ; **WB**, anti-human LC western blots.

(A), Pa1- λ LC; red rectangles indicate the data corresponding to the LC preparation used in this study.

(B), Pa2- κ and Pa3- κ LC.

Supplementary Figure S2

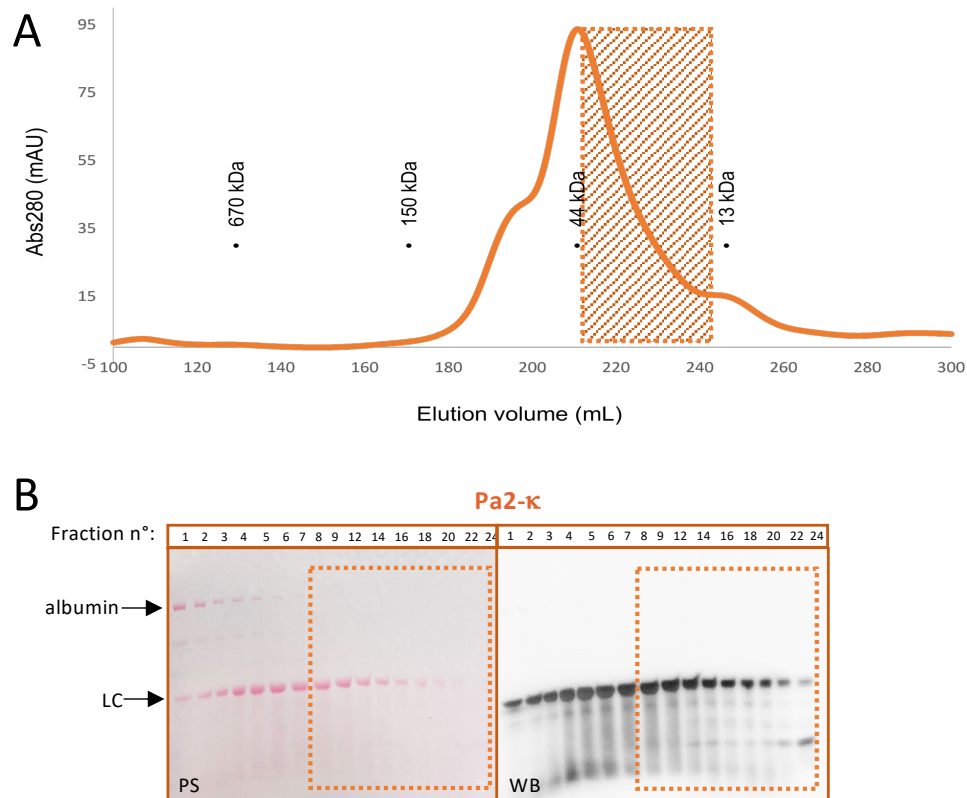


Figure S2. Example of selection of LC fractions devoid of contaminants after purification by size-exclusion chromatography.

(A) Size exclusion chromatogram of Pa2-κ LC from Figure 1; striped rectangle indicates the part of LC-containing peak fractions that have been selected for final concentration and use in experiments.

(B) Ponceau S staining (PS) and western blot (WB, anti-human LC) of fractions corresponding to the LC-containing pic; selected fractions excluding albumin are indicated by dashed rectangles.

Supplementary Figure S3

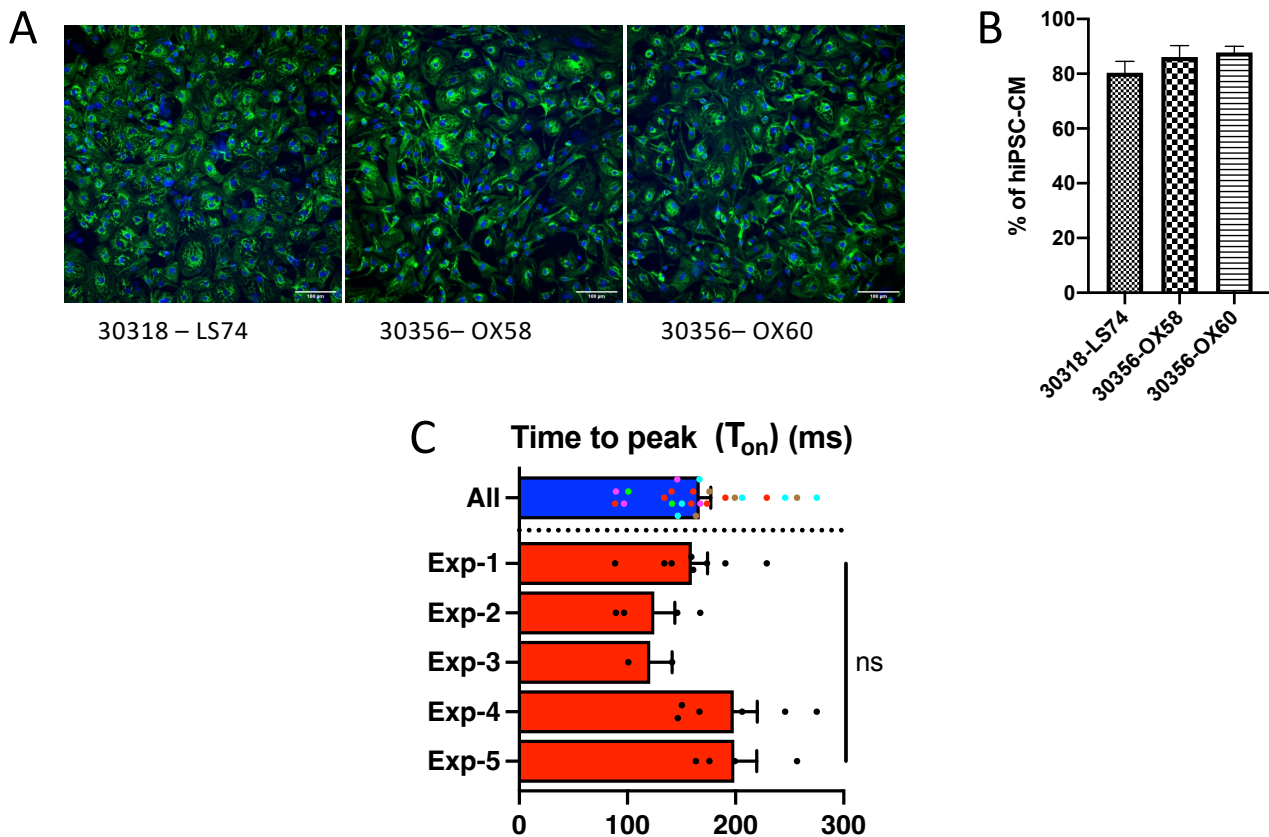


Figure S3. Characterization of hiPSC derived cardiomyocytes (CM) used in this study.

(A) Quality control of differentiation to CMs of 2 hiPSC cell lines: **30318** (1 batch of differentiation: LS74) and **30356** (2 batches of differentiation: OX58 and OX60), as assayed by cardiomyocyte-specific **cardiac troponin T (c-TnT)** immunofluorescence labeling; bars, 100 μ m.

(B) Percentage of **c-TnT** positive cells: CMs exceeded 80% of cells in all cases; n=8 field views for each batch.

(C) Statistical analysis of values for time to peak (T_{on}) of calcium transients in paced non-LC treated (control) spheroids from 5 independent experiments. Each experiment was performed with a newly thawed batch of hiPSC-derived CMs originating from one differentiation. All 3 batches were used through experiments. Kruskal-Wallis test combined with Dunn's multiple comparison tests were applied. No statistically significant differences among untreated spheroids (controls) were found for T_{on} , a parameter which changes after LC treatment.