

Supplemental Material

Rare variants in the *FBNI* gene are associated with sporadic dilated cardiomyopathy in a Chinese Han population

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Table S1. Rare variants of *FBNI* identified in DCM cases and controls.

HGVS.c (NM_000138)	Consequence	MAF in gnomAD exome (EAS)	MAF in ExAC (EAS)	MAF in 1,000 Genomes (EAS)	VEST3 score	REVEL score	CADD phred	MutationTaster pred	ACMG	Pathogenicity	NO. in cases (n = 53)	NO. in controls (n = 9)
c. C13T	missense	5.82E-05	0.0001	ND	0.18	0.119	24.6	N	PM1+PM2+BP1	Uncertain significance	1	0
c. G100C	missense	5.80E-05	ND	ND	0.147	0.103	19.93	N	PM1+PM2+BP1	Uncertain significance	0	1
c. A149T	missense	ND	ND	ND	0.48	0.278	14.91	N	PM1+PM2+BP1	Uncertain significance	2	0
c. T605C	missense	0.0006	ND	ND	0.853	0.829	26.9	D	PM2+PP3+BP1	Uncertain significance	1	0
c. C657G	missense	0.0002	0.0001	ND	0.941	0.741	23.9	D	PM2+PP3+BP1	Uncertain significance	1	0
c. A913G	missense	0.0003	0.0001	ND	0.106	0.22	11	N	PM2+BP1	Uncertain significance	3	1
c. T1217A	missense	ND	ND	ND	0.195	0.132	11	N	PM2+BP1	Uncertain significance	5	0
c. T1841C	missense	0.0002	0.0002	ND	0.837	0.961	25.1	D	PM2+PP3+BP1	Uncertain significance	1	0
c. C1858T	missense	0.0001	ND	ND	0.402	0.224	20.1	D	PM2+BP1	Uncertain significance	1	1
c. A2011G	missense	ND	ND	ND	0.037	0.31	6.945	N	PM2+BP1	Uncertain significance	1	0
c. A2149T	missense	ND	ND	ND	0.386	0.405	16.23	D	PM2+BP1	Uncertain significance	0	1

HGVS.c (NM_000138)	Consequence	MAF in gnomAD exome (EAS)	MAF in ExAC (EAS)	MAF in 1,000 Genomes (EAS)	VEST3 score	REVEL score	CADD phred	MutationTaster pred	ACMG	Pathogenicity	NO. in cases (n = 53)	NO. in controls (n = 9)
c. G2269C	missense	ND	ND	ND	0.554	0.768	26.2	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G2290A	missense	0.0003	0.0002	ND	0.05	0.259	16.72	N	PM2+BP1	Uncertain significance	1	0
c. A2613C	missense	5.80E-05	ND	ND	0.588	0.652	24	D	PM2+PP3+PP5+BP1	Uncertain significance	1	0
c. 2793dupG	frameshift insertion	ND	ND	ND	NA	NA	NA		PVS1+PM2	Likely pathogenic	1	0
c. G2887C	missense	5.80E-05	0.0001	ND	0.192	0.344	19.9	D	PM2+PP3+BP1	Uncertain significance	1	1
c. A3142G	missense	0.0002	ND	ND	0.348	0.297	23	D	PM2+BP1	Uncertain significance	1	1
c. G3571A	missense	5.81E-05	0	ND	0.105	0.407	24.1	D	PM2+PP3+BP1	Uncertain significance	1	0
c. C3697G	missense	ND	ND	ND	0.35	0.431	20.2	D	PM2+PP3+BP1	Uncertain significance	1	0
c. A3971G	missense	0	0	ND	0.442	0.893	23.4	D	PM2+PP3+BP1	Uncertain significance	1	0
c. T3984G	missense	ND	ND	ND	0.259	0.607	23.1	D	PM2+BP1+BP4	Likely benign	0	1
c. A4325G	missense	ND	ND	ND	0.544	0.484	17.69	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G4406A	missense	0	0	ND	0.85	0.791	33	D	PM2+PM5+PP3+BP1	Uncertain significance	1	0
c. G4750A	missense	0.0002	0.0002	ND	0.71	0.659	28.2	D	PM2+PP3+BS2+BP1	Likely benign	1	0
c. C5210G	missense	ND	ND	ND	0.947	0.927	28.5	D	PM2+PP3+BP1	Uncertain significance	1	0
c. C5509T	missense	0	0	ND	0.803	0.719	21.6	D	PM2+PP3+BP1	Uncertain significance	1	0
c. A5596G	missense	0.0004	0.0004	ND	0.44	0.357	12.48	D	PM2+BP1	Uncertain significance	2	1
c. A5750G	missense	ND	ND	ND	0.186	0.292	15.09	D	PM2+BP1	Uncertain significance	1	0
c. G6174A	missense	ND	ND	ND	0.169	0.24	22.4	D	PM2+BP1	Uncertain significance	1	0
c. A6270C	missense	ND	ND	ND	0.324	0.507	19.47	D	PM2+PP3+BP1	Uncertain significance	1	0
c. A6301G	missense	ND	ND	ND	0.085	0.245	7.934	N	PM2+BP1	Uncertain significance	1	0

HGVS.c (NM_000138)	Consequence	MAF in gnomAD exome (EAS)	MAF in ExAC (EAS)	MAF in 1,000 Genomes (EAS)	VEST3 score	REVEL score	CADD phred	MutationTaster pred	ACMG	Pathogenicity	NO. in cases (n = 53)	NO. in controls (n = 9)
c. G6725A	missense	0.0001	0.0002	ND	0.432	0.518	28.2	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G6845A	missense	0.0001	0.0002	ND	0.687	0.543	24.1	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G6863C	missense	ND	ND	ND	0.097	0.304	19.8	D	PM2+BP1	Uncertain significance	1	0
c. C7082T	missense	0	ND	ND	0.601	0.611	33	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G7099A	missense	0	0	ND	0.531	0.644	27	D	PM2+PP3+BP1	Uncertain significance	1	0
c. C7193T	missense	0	ND	ND	0.616	0.584	26.2	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G7231A	missense	5.80E-05	0.0001	ND	0.029	0.493	22	D	PM2+BP1	Uncertain significance	1	0
c. G7241A	missense	0.0001	0	ND	0.665	0.45	22.9	D	PM2+BP1	Uncertain significance	1	0
c. C7559T	missense	0.0003	0.0001	ND	0.892	0.656	26.4	D	PM2+PP3+BP1	Uncertain significance	2	0
c. A7598G	missense	0	0	ND	0.042	0.37	11.61	D	PM2+BP1	Uncertain significance	1	0
c. A7807G	missense	ND	ND	ND	0.361	0.586	25.1	D	PM2+PP3+BP1	Uncertain significance	1	0
c. G7816A	missense	0	ND	ND	0.486	0.377	23.9	D	PM2+PP3+BP1	Uncertain significance	1	0
c. T8069G	missense	0.0005	0.0006	ND	0.525	0.373	20.4	D	PM2+BP1	Uncertain significance	1	1
c. G8081A	missense	0	ND	ND	0.801	0.422	23.2	D	PM2+BP1	Uncertain significance	1	0
c. C8306G	missense	ND	ND	ND	0.852	0.697	25.1	D	PM2+PP3+BP1	Uncertain significance	1	0
c. T8507A	missense	ND	ND	ND	0.931	0.732	25.4	D	PM2+PP3+BP1	Uncertain significance	1	0

Abbreviations: HGVS.c: variants using HGVS notation (DNA level); MAF, minor allele frequency; EAS, East Asian; NA, not available; ND, not detected; VEST3 score, VEST(variant effect scoring tool) version 3 score; REVEL score, rare exome variant ensemble learner score; CADD phred, CADD (combined annotation dependent depletion) phred-like score, the deleteriousness threshold recommended by the authors is greater than 20; MutationTaster pred, MutationTaster prediction, which evaluates the pathogenic potential of DNA sequence alterations, D: disease causing - i.e. probably deleterious, N: polymorphism - i.e. probably harmless; ACMG, the American College of Medical Genetics and Genomics; PVS1, PM1, PM2, PM5, PP3, BS2, BP1, and BP4 are part of the 28 criteria in the ACMG guidelines for the clinical interpretation of sequence variations.

Table S2. The deleteriousness prediction for rare missense variants of *FBNI* in DCM cases and controls.

HGVS.c (NM_000138)	HGVS.p	MAF in gnomAD exome (EAS)	VEST3 score	REVEL score	CADD phred	MutationTaster pred	NO. in cases	NO. in controls
c.T605C	p.I202T	0.0006	0.853	0.829	26.9	D	1	0
c.C657G	p.H219Q	0.0002	0.941	0.741	23.9	D	1	0
c.T1841C	p.I614T	0.0002	0.837	0.961	25.1	D	1	0
c.G2269C	p.D757H	ND	0.554	0.768	26.2	D	1	0
c.A2613C	p.L871F	5.80E-05	0.588	0.652	24	D	1	0
c.G4406A	p.R1469H	0	0.85	0.791	33	D	1	0
c.C5210G	P1737R	ND	0.947	0.927	28.5	D	1	0
c.C5509T	P1837S	0	0.803	0.719	21.6	D	1	0
c.G6845A	R2282Q	0.0001	0.687	0.543	24.1	D	1	0
c.C7082T	p.S2361L	0	0.601	0.611	33	D	1	0
c.G7099A	p.G2367R	0	0.531	0.644	27	D	1	0
c.C7193T	p.T2398I	0	0.616	0.584	26.2	D	1	0
c.G7241A	p.R2414Q	0.0001	0.665	0.45	22.9	D	1	0
c.C7559T	p.T2520M	0.0003	0.892	0.656	26.4	D	2	0
c.G8081A	p.R2694Q	0	0.801	0.422	23.2	D	1	0
c.C8306G	p.S2769C	ND	0.852	0.697	25.1	D	1	0
c.T8507A	p.L2836H	ND	0.931	0.732	25.4	D	1	0

Abbreviations: HGVS.c: variants using HGVS notation (DNA level); HGVS.p: HGVS notation (protein level); MAF, minor allele frequency; EAS, East Asian; VEST3 score, VEST(variant effect scoring tool) version 3 score; REVEL score, rare exome variant ensemble learner score; CADD phred, CADD (combined annotation dependent depletion) phred-like score, the deleteriousness threshold recommended by the authors is greater than 20; MutationTaster pred, MutationTaster prediction, which evaluates the pathogenic potential of DNA sequence alterations, D: disease causing - i.e. probably deleterious.

Table S3. Rare deleterious *FBNI* variants across the DCM cohort.

Subject	Exon	HGVS.c (NM_000138)	HGVS.p	SNP	MAF in gnomAD exome (EAS)	MAF in ExAC (EAS)	MAF in 1,000 Genomes (EAS)	ACMG	Pathogenicity
DCM1	7	c.T605C	p.I202T	rs1300991442	0.0006	ND	ND	PM2+PP3+BP1	Uncertain significance
DCM2	7	c.C657G	p.H219Q	rs774754863	0.0002	0.0001	ND	PM2+PP3+BP1	Uncertain significance
DCM3	16	c.T1841C	p.I614T	rs762427180	0.0002	0.0002	ND	PM2+PP3+BP1	Uncertain significance
DCM4	19	c.G2269C	p.D757H	rs2043616147	ND	ND	ND	PM2+PP3+BP1	Uncertain significance
DCM5	22	c.A2613C	p.L871F	rs770290542	5.80E-05	ND	ND	PM2+PP3+PP5+BP1	Uncertain significance
DCM6	24	c.2793dupG	p.S932fs	NA	ND	ND	ND	PVS1+PM2	Likely pathogenic
DCM7	36	c.G4406A	p.R1469H	rs397515808	0	0	ND	PM2+PM5+PP3+BP1	Uncertain significance
DCM8	42	c.C5210G	P1737R	NA	ND	ND	ND	PM2+PP3+BP1	Uncertain significance
DCM9	45	c.C5509T	P1837S	rs755430984	0	0	ND	PM2+PP3+BP1	Uncertain significance
DCM10	56	c.G6845A	R2282Q	rs759696323	0.0001	0.0002	ND	PM2+PP3+BP1	Uncertain significance
DCM11	58	c.C7082T	p.S2361L	rs397515844	0	ND	ND	PM2+PP3+BP1	Uncertain significance
DCM12	58	c.G7099A	p.G2367R	rs368978109	0	0	ND	PM2+PP3+BP1	Uncertain significance
DCM13	58	c.C7193T	p.T2398I	rs1169351376	0	ND	ND	PM2+PP3+BP1	Uncertain significance
DCM14	59	c.G7241A	p.R2414Q	rs143863014	0.0001	0	ND	PM2+BP1	Uncertain significance
DCM15	61	c.C7559T	p.T2520M	rs763759308	0.0003	0.0001	ND	PM2+PP3+BP1	Uncertain significance
DCM16	61	c.C7559T	p.T2520M	rs763759308	0.0003	0.0001	ND	PM2+PP3+BP1	Uncertain significance
DCM17	65	c.G8081A	p.R2694Q	rs371375126	0	ND	ND	PM2+BP1	Uncertain significance
DCM18	66	c.C8306G	p.S2769C	NA	ND	ND	ND	PM2+PP3+BP1	Uncertain significance
DCM19	66	c.T8507A	p.L2836H	NA	ND	ND	ND	PM2+PP3+BP1	Uncertain significance

Abbreviations: HGVS.c: variants using HGVS notation (DNA level); HGVS.p: HGVS notation (protein level); DCM, dilated cardiomyopathy; MAF, minor allele frequency; EAS, East Asian; NA, not available; ND, not detected; ACMG, the American College of Medical Genetics and

Genomics; PVS1, PM2, PP3, PP5, and BP1 are part of the 28 criteria in the ACMG guidelines for the clinical interpretation of sequence variations.

Table S4. List of 44 DCM-associated genes evaluated by ClinGen and included in this study.

	ClinGen DCM gene-disease validity classification	Gene symbol
High evidence genes	Moderate	<i>ACTC1</i>
	Moderate	<i>ACTN2</i>
	Definitive	<i>BAG3</i>
	Definitive	<i>DES</i>
	Strong	<i>DSP</i>
	Definitive	<i>FLNC</i>
	Moderate	<i>JPH2</i>
	Definitive	<i>LMNA</i>
	Definitive	<i>MYH7</i>
	Moderate	<i>NEXN</i>
	Definitive	<i>PLN</i>
	Definitive	<i>RBM20</i>
	Definitive	<i>SCN5A</i>
	Definitive	<i>TNNC1</i>
	Moderate	<i>TNNI3</i>
	Definitive	<i>TNNT2</i>
	Moderate	<i>TPM1</i>
	Definitive	<i>TTN</i>
	Moderate	<i>VCL</i>
	Low/variable evidence genes	Limited
Limited		<i>ANKRD1</i>
Limited		<i>CSRP3</i>
Limited		<i>CTF1</i>
Limited		<i>DSG2</i>
Limited		<i>DTNA</i>
Limited		<i>EYA4</i>
Limited		<i>GATAD1</i>
Limited		<i>ILK</i>
Limited		<i>LAMA4</i>
Limited		<i>LDB3</i>
Limited		<i>MYBPC3</i>
Limited		<i>MYH6</i>
Limited		<i>MYL2</i>
Limited		<i>MYPN</i>
Limited		<i>NEBL</i>
Limited		<i>NKX2-5</i>
Limited		<i>OBSCN</i>
Limited		<i>PLEKHM2</i>
Limited		<i>PRDM16</i>

	ClinGen DCM gene-disease validity classification	Gene symbol
Low/variable evidence genes	Limited	<i>PSEN2</i>
	Limited	<i>SGCD</i>
	Limited	<i>TBX20</i>
	Limited	<i>TCAP</i>
	Limited	<i>TNNI3K</i>

Abbreviations: DCM, dilated cardiomyopathy; ClinGen: Clinical Genome Resource.

Table S5. Rare variants identified as pathogenic or likely pathogenic in DCM-associated genes according to ACMG guidelines.

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr10	121411252	TG	T	frameshift deletion	<i>BAG3</i>	PVS1+PM2	Likely pathogenic	1	0
chr10	121411299	T	TA	frameshift insertion	<i>BAG3</i>	PVS1+PM2	Likely pathogenic	1	0
chr10	121429549	C	T	stop gain	<i>BAG3</i>	PVS1+PP3+PP5	Pathogenic	1	0
chr6	7563990	C	T	stop gain	<i>DSP</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr6	7568096	ACATG	A	frameshift deletion	<i>DSP</i>	PVS1+PM2	Likely pathogenic	1	0
chr6	7577219	C	T	stop gain	<i>DSP</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr6	7580621	C	T	stop gain	<i>DSP</i>	PVS1+PM2+PP5	Pathogenic	2	0
chr6	7581438	AG	A	frameshift deletion	<i>DSP</i>	PVS1+PM2	Likely pathogenic	1	0
chr6	7584541	CTT	C	frameshift deletion	<i>DSP</i>	PVS1+PM2	Likely pathogenic	1	0
chr6	7585369	CAG	C	frameshift deletion	<i>DSP</i>	PVS1+PM2	Likely pathogenic	1	0
chr6	7585826	C	CA	frameshift insertion	<i>DSP</i>	PVS1+PM2	Likely pathogenic	1	0
chr7	128470692	A	T	startloss	<i>FLNC</i>	PVS1+PM2	Likely pathogenic	1	0
chr7	128477557	C	T	stop gain	<i>FLNC</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr7	128477603	G	C	splicing	<i>FLNC</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr7	128480139	A	T	stop gain	<i>FLNC</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr7	128481226	TTC	T	frameshift deletion	<i>FLNC</i>	PVS1+PM2	Likely pathogenic	1	0
chr7	128482356	G	GC	frameshift insertion	<i>FLNC</i>	PVS1+PM2	Likely pathogenic	1	0
chr7	128484099	C	T	stop gain	<i>FLNC</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr7	128488905	CTG	C	frameshift deletion	<i>FLNC</i>	PVS1+PM2	Likely pathogenic	1	0
chr7	128489402	C	T	stop gain	<i>FLNC</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr7	128491547	G	T	stop gain	<i>FLNC</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr7	128494863	TG	T	frameshift deletion	<i>FLNC</i>	PVS1+PM2	Likely pathogenic	1	0
chr20	42788645	GC	G	frameshift deletion	<i>JPH2</i>	PVS1+PM2	Likely pathogenic	1	0
chr20	42788815	G	GCCGA	frameshift insertion	<i>JPH2</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	156084729	GGC	G	frameshift deletion	<i>LMNA</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	156084885	T	C	missense	<i>LMNA</i>	PM1+PM2+PM5+PP3	Likely pathogenic	1	0
chr1	156104248	C	T	missense	<i>LMNA</i>	PM1+PM2+PP3+PP5	Likely pathogenic	1	0
chr1	156104692	C	T	stop gain	<i>LMNA</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr1	156104767	G	C	splicing	<i>LMNA</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr1	156105758	C	T	missense	<i>LMNA</i>	PM1+PM2+PP3+PP5	Likely pathogenic	1	0
chr1	156105800	C	T	missense	<i>LMNA</i>	PM1+PM2+PP3+PP5	Likely pathogenic	1	0
chr1	156106903	G	GATCT	splicing	<i>LMNA</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	156108541	G	GA	frameshift insertion	<i>LMNA</i>	PVS1+PM2	Likely pathogenic	1	0
chr14	23883069	G	A	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23884407	T	C	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23884879	G	C	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	0	1
chr14	23885010	C	T	missense	<i>MYH7</i>	PM1+PM2+PM5+PP2+BP1	Likely pathogenic	1	0
chr14	23886132	C	T	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23886383	G	A	missense	<i>MYH7</i>	PM1+PM2+PP2+PP5+BP1	Likely pathogenic	1	0
chr14	23886440	G	A	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23887584	G	A	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr14	23888742	C	T	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	5	0
chr14	23891462	C	T	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23891500	C	T	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	0	2
chr14	23892874	T	C	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23894061	A	G	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr14	23894084	C	T	missense	<i>MYH7</i>	PM1+PM2+PM5+PP2+PP3+BP1	Likely pathogenic	0	1
chr14	23901959	G	C	missense	<i>MYH7</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr10	112404329	AG	A	frameshift deletion	<i>RBM20</i>	PVS1+PM2	Likely pathogenic	1	0
chr10	112540899	C	T	stop gain	<i>RBM20</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr10	112581063	GA	G	frameshift deletion	<i>RBM20</i>	PVS1+PM2	Likely pathogenic	1	0
chr3	52486161	CA	C	frameshift deletion	<i>TNNC1</i>	PVS1+PM2	Likely pathogenic	0	1
chr1	201328764	C	T	missense	<i>TNNT2</i>	PM1+PM2+PP3+PP5+BP1	Likely pathogenic	1	0
chr1	201333497	G	A	missense	<i>TNNT2</i>	PM1+PM2+PP3+PP5+BP1	Likely pathogenic	1	0
chr15	63362083	C	T	stop gain	<i>TPM1</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179395540	CTG	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	2	0
chr2	179395588	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179396395	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179396467	TA	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179397088	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179399943	CT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179404263	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr2	179404296	G	GA	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179404375	T	TC	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179408130	AG	A	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179411050	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179411339	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP5	Pathogenic	1	0
chr2	179414134	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179414391	G	GT	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179416930	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179417382	A	AT	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179417474	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179418306	CG	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179418850	C	CT	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179419260	ATG	A	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179419385	AC	A	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179421736	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179421743	CA	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179422374	C	CT	splicing	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179422963	C	A	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	2	0
chr2	179424695	CAT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	2	0
chr2	179424731	A	AT	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179425592	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179426257	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr2	179426643	TA	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179427020	G	GA	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179427128	TC	T	stop gain	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179429822	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	4	0
chr2	179429903	TA	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179431835	CA	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	0	1
chr2	179431880	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP5	Pathogenic	1	0
chr2	179432127	GTTCT	G	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179432542	GT	G	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179432880	AG	A	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179433260	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179433839	T	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179434089	CTT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179434925	C	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179435045	CT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179435390	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP5	Pathogenic	1	0
chr2	179436280	G	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179436949	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179437013	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179437449	T	TAAAG	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179438698	G	C	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179439257	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr2	179439773	ACT	A	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179440577	CA	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179440643	T	TA	stop gain	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179440808	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179441449	A	T	stop gain	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179441550	CA	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179443655	G	GT	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179447724	TG	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179449216	CTCCA	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179449972	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	2	0
chr2	179451478	TG	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	2	0
chr2	179452411	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179454531	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179454906	CT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179455623	T	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179456095	A	AATCTC	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179456620	C	T	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179457670	C	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179460312	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179464371	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179468602	C	T	splicing	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179468792	C	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr2	179468831	CACCAT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179468939	CTACTT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179468996	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179469000	C	CTCTG	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179469506	T	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179469581	CT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179470331	A	C	stop gain	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179472209	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179473331	AC	A	splicing	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179474016	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179474816	C	T	splicing	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179476662	C	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179477578	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	2	0
chr2	179480366	A	G	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179481235	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179481609	G	GTACTT	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179481846	C	T	splicing	<i>TTN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr2	179482809	C	G	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179494967	C	T	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	2	0
chr2	179494977	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179495921	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179500166	C	A	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr2	179500879	CT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179501421	AG	A	stop gain	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179604536	TC	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179604706	C	T	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179604886	CT	C	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179606500	TG	T	frameshift deletion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179631340	C	A	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179634405	C	T	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179638282	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179647792	C	G	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179648832	C	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179650739	C	CA	frameshift insertion	<i>TTN</i>	PVS1+PM2	Likely pathogenic	1	0
chr2	179650808	G	A	stop gain	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr2	179659647	A	G	splicing	<i>TTN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr10	92679010	G	GT	frameshift insertion	<i>ANKRD1</i>	PVS1+PM2	Likely pathogenic	2	2
chr11	19207813	G	A	stop gain	<i>CSRP3</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr18	29099830	G	A	missense	<i>DSG2</i>	PM1+PM2+PP3+PP5+BP1	Likely pathogenic	0	1
chr18	29104430	T	A	stop gain	<i>DSG2</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr18	29118742	GA	G	frameshift deletion	<i>DSG2</i>	PVS1+PM2	Likely pathogenic	1	0
chr18	32470299	CT	C	frameshift deletion	<i>DTNA</i>	PVS1+PM2	Likely pathogenic	1	0
chr6	112441484	A	C	splicing	<i>LAMA4</i>	PVS1+PM2+PP3	Pathogenic	0	1
chr6	112452303	A	ATC	frameshift insertion	<i>LAMA4</i>	PVS1+PM2	Likely pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr6	112496514	C	T	splicing	<i>LAMA4</i>	PVS1+PM2+PP3	Pathogenic	0	1
chr11	47353626	G	A	stop gain	<i>MYBPC3</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr11	47354119	TG	T	frameshift deletion	<i>MYBPC3</i>	PVS1+PM2	Likely pathogenic	3	0
chr11	47364572	C	T	missense	<i>MYBPC3</i>	PM1+PM2+PM5+PP3	Likely pathogenic	1	0
chr11	47374196	C	T	startloss	<i>MYBPC3</i>	PVS1+PM2	Likely pathogenic	1	0
chr14	23854251	C	A	splicing	<i>MYH6</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr14	23863118	C	T	splicing	<i>MYH6</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr12	111351128	C	T	missense	<i>MYL2</i>	PM1+PM2+PP2+PP3	Likely pathogenic	1	0
chr10	69957164	C	T	stop gain	<i>MYPN</i>	PVS1+PM2+PP3+PP5	Pathogenic	1	0
chr10	21120237	T	C	splicing	<i>NEBL</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr10	21120430	GCT	G	frameshift deletion	<i>NEBL</i>	PVS1+PM2	Likely pathogenic	0	1
chr10	21129691	G	A	stop gain	<i>NEBL</i>	PVS1+PM2	Likely pathogenic	0	1
chr10	21139402	AT	A	frameshift deletion	<i>NEBL</i>	PVS1+PM2	Likely pathogenic	2	0
chr10	21177032	CTG	C	frameshift deletion	<i>NEBL</i>	PVS1+PM2	Likely pathogenic	0	1
chr5	172660415	CA	C	stop gain	<i>NKX2-5</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228412436	G	C	splicing	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr1	228431123	AG	A	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228433212	C	T	stop gain	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228464876	CGG	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	0	1
chr1	228467521	G	A	splicing	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr1	228469804	C	T	stop gain	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr1	228487176	CAT	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0

Chr	Pos	Ref	Alt	Consequence	Gene symbol	ACMG	Pathogenicity	NO. in cases	NO. in controls
chr1	228494102	CAG	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228495865	G	T	stop gain	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr1	228506894	CT	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228521482	G	A	splicing	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	0	1
chr1	228525020	C	G	stop gain	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	0	1
chr1	228525846	G	A	splicing	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	0	1
chr1	228539121	CA	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228547775	CAGAG	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228552733	AG	A	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	0	1
chr1	228553810	C	T	stop gain	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	0	1
chr1	228554527	CAG	C	frameshift deletion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228558395	G	T	stop gain	<i>OBSCN</i>	PVS1+PM2+PP3	Pathogenic	1	0
chr1	228559297	G	GC	frameshift insertion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr1	228560305	C	CAGGT	frameshift insertion	<i>OBSCN</i>	PVS1+PM2	Likely pathogenic	1	0
chr5	155771665	T	A	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr5	155935612	A	G	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	2	0
chr5	156016244	A	C	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	3	1
chr5	156016305	A	C	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr5	156184675	C	T	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	1
chr5	156186300	C	G	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	1	0
chr5	156186354	G	T	missense	<i>SGCD</i>	PM1+PM2+PP2+PP3+BP1	Likely pathogenic	0	1

Abbreviations: Chr, chromosome; Pos, position; Ref, reference allele; Alt, alternative allele.

Table S6. Baseline comparisons among *FBNI* carriers, patients positive for DCM-associated genes, and patients negative for DCM-associated genes.

	<i>FBNI</i> ⁺	<i>FBNI</i> ⁻ / DCMGenes ⁺	<i>FBNI</i> ⁻ / DCMGenes ⁻	<i>P</i> value	
	(n = 19)	(n = 205)	(n = 835)	<i>FBNI</i> ⁺ vs. <i>FBNI</i> ⁻ / DCMGenes ⁺	<i>FBNI</i> ⁺ vs. <i>FBNI</i> ⁻ / DCMGenes ⁻
Male (%)	12 (63.2)	160 (78.0)	604 (72.3)	0.141	0.378
Age of onset (years)	54.74 ± 19.57	51.12 ± 12.47	52.21 ± 13.95	0.254	0.439
Age at enrollment (years)	57.42 ± 18.63	54.23 ± 12.92	55.09 ± 14.32	0.324	0.486-
NYHA III/IV (%)	14 (73.7)	158 (77.1)	555 (66.5)	0.738	0.509
Smoke (%)	5 (26.3)	89 (43.4)	355 (42.5)	0.149	0.157
Alcohol intake (%)	0 (0.0)	26 (15.1)	112 (16.5)	0.105	0.086
Non-fatal stroke (%)	1 (5.3)	5 (2.4)	41 (4.9)	0.466	0.945
Hypertension (%)	10 (52.6)	72 (35.1)	444 (53.2)	0.13	0.963
Hyperlipidemia (%)	1 (5.3)	14 (6.8)	91 (10.9)	0.794	0.433
Diabetes mellitus (%)	4 (21.1)	34 (16.6)	138 (16.5)	0.62	0.6
Renal insufficiency (%)	6 (33.3)	8 (4.4)	71 (9.5)	<0.001	0.001
Body mass index (kg/m ²)	24.97 ± 3.36	23.73 ± 4.13	23.88 ± 4.59	0.419	0.506
TC (mmol/L)	3.89 ± 1.25	3.82 ± 0.87	3.93 ± 1.01	0.785	0.885
TG (mmol/L)	1.40 ± 0.75	1.39 ± 0.80	1.39 ± 0.98	0.96	0.988
HDL-C (mmol/L)	0.98 ± 0.25	0.93 ± 0.28	0.97 ± 0.34	0.53	0.935
LDL-C (mmol/L)	2.40 ± 1.15	2.43 ± 0.78	2.51 ± 0.83	0.891	0.619
NT-proBNP (pg/mL)	3368.00	3804.00	3841.00	0.969	0.853
	[1533.00, 10404.00]	[1655.00, 7459.50]	[1722.00, 9000.00]		
ALT (U/L)	30.00 [22.00, 45.50]	25.00 [16.25, 43.75]	26.00 [16.00, 48.00]	0.324	0.351

AST (U/L)	27.00 [22.00, 34.50]	27.00 [18.00, 39.00]	27.00 [20.00, 42.00]	0.754	0.935
Cr (μ mol/L)	89.00 [73.00, 127.00]	87.00 [72.00, 102.00]	89.00 [74.00, 113.25]	0.617	0.85
Atrial fibrillation (%)	8 (42.1)	42 (20.5)	187 (22.4)	0.03	0.043
Non-sustained ventricular tachycardia (%)	2 (10.5)	34 (16.6)	109 (13.1)	0.491	0.746
Left bundle branch block (%)	4 (21.1)	12 (5.9)	96 (11.5)	0.014	0.2
Any arrhythmia (%)	12 (63.2)	85 (41.5)	353 (42.3)	0.068	0.069
IVS (mm)	9.44 \pm 1.34	9.37 \pm 1.22	9.61 \pm 1.47	0.818	0.638
LVPW (mm)	9.44 \pm 1.04	9.34 \pm 1.13	9.60 \pm 1.39	0.701	0.629
LVEDD (mm)	65.42 \pm 9.44	66.24 \pm 7.72	66.77 \pm 8.37	0.666	0.488
LAD (mm)	46.16 \pm 8.08	45.65 \pm 7.87	45.66 \pm 7.91	0.788	0.785
LVEF (%)	33.32 \pm 8.35	29.80 \pm 9.67	31.77 \pm 10.08	0.127	0.506
E/A	1.45 \pm 1.00	1.71 \pm 0.90	1.75 \pm 1.67	0.403	0.584
E/e'	24.75 \pm 16.80	21.82 \pm 11.29	22.67 \pm 12.64	0.497	0.646
Aortic root diameter (mm)	27.88 \pm 2.80	31.07 \pm 28.56	29.26 \pm 3.97	0.753	0.327
Proximal ascending aorta diameter (mm)	34.58 \pm 4.46	31.71 \pm 4.96	32.82 \pm 4.78	0.057	0.207
Aortic root dilatation (%)	0 (0.0)	8 (6.2)	41 (7.9)	0.397	0.332
Proximal ascending aorta dilatation (%)	6 (42.9)	31 (25.4)	152 (30.9)	0.165	0.341
Any aorta dilatation (%)	6 (40.0)	37 (23.1)	169 (27.0)	0.147	0.266

Pacemaker implantation (%)	0 (0.0)	11 (5.4)	43 (5.2)	0.3	0.309
Implantable cardioverter-defibrillator (%)	2 (10.5)	4 (2.0)	13 (1.6)	0.027	0.003
Cardiotonic use (%)	10 (52.6)	102 (49.8)	395 (47.6)	0.81	0.664
Diuretic use (%)	16 (84.2)	173 (84.4)	680 (81.9)	0.984	0.798
ACE inhibitor use (%)	13 (68.4)	152 (74.1)	637 (76.7)	0.588	0.403
Beta-blocker use (%)	8 (42.1)	101 (49.3)	446 (53.7)	0.55	0.315
Aldactone use (%)	17 (89.5)	159 (77.6)	638 (76.9)	0.226	0.196

Abbreviations: DCM, dilated cardiomyopathy; *FBNI*⁺, DCM cases carrying rare deleterious variants in *FBNI*; *FBNI*⁻ / DCMGenes⁺, DCM cases not carrying rare deleterious variants in *FBNI*, but carrying rare deleterious variants in other known DCM-associated genes; *FBNI*⁻ / DCMGenes⁻, DCM cases not carrying rare deleterious variants in either *FBNI* or other known DCM-associated genes; NYHA, New York Heart Association; NT-proBNP, N-terminal pro-B-type natriuretic peptide; TC: total cholesterol; TG: triglycerides; HDL-C: high-density lipoprotein cholesterol; LDL-C: low-density lipoprotein cholesterol; ALT: alanine aminotransferase; AST: aspartate aminotransferase; Cr: creatinine; IVS, interventricular septum; LVPW, left ventricular posterior wall; LVEDD, left ventricular end-diastolic diameter; LAD, left atrial diameter; LVEF, left ventricular ejection fraction.

For non-normally distributed data, values are presented as the median and interquartile range (IQR). All other numerical data, which are normally distributed, are represented as the mean \pm standard deviation (SD). Categorical data are provided as counts (percentages).

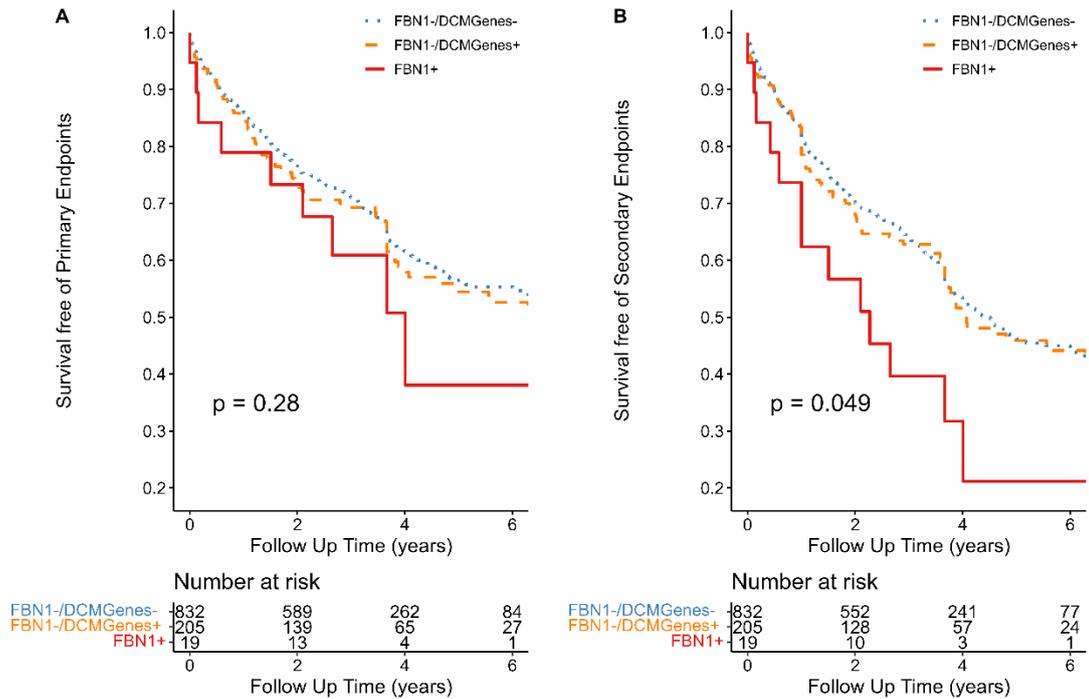


Figure S1. Clinical outcomes in different groups of DCM patients. (A) Kaplan-Meier curve illustrates survival free of primary endpoints, which include cardiac mortality or heart transplantation. (B) The curve represents survival free of secondary endpoints, comprising all-cause mortality or heart failure recurrence. Probability values were calculated using the log-rank test. *FBNI*⁺, DCM cases carrying rare deleterious variants in *FBNI*; *FBNI*⁻ / *DCMGenes*⁺, DCM cases not carrying rare deleterious variants in *FBNI*, but carrying rare deleterious variants in other known DCM-associated genes; *FBNI*⁻ / *DCMGenes*⁻, DCM cases not carrying rare deleterious variants in either *FBNI* or other known DCM-associated genes.