## Supplementary

Table S1 Sample size of patients with TTN mutation in LUAD and LUSC

Therepy types	LU	AD	LUSC		
Therapy types	<i>TTN</i> -MT	<i>TTN</i> -WT	<i>TTN</i> -MT	<i>TTN</i> -WT	
Total	205	358	326	179	
Chemotherapy	71	116	101	42	

TTN, titin; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma; MT, mutant-type; WT, wild-type.

Table S2 Sample size of patients with different types of TTN mutation in LUAD and LUSC

	LUAD				LUSC			
Therapy type	Missense mutation	Nonsense mutation	Splice site	Multi hit	Missense mutation	Nonsense mutation	Splice site	Multi hit
Chemotherapy	62	1	1	7	88	3	0	10

TTN, titin; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma.



**Figure S1** The association between different types of *TTN* mutation and OS in LUAD and LUSC patients on chemotherapy. (A,B) The difference in OS between patients with missense mutation (n=62) of *TTN* and *TTN*-WT (n=116) in LUAD patients treated with chemotherapy, and between patients with missense mutation (n=88) of *TTN* and *TTN*-WT (n=42) in LUSC treated with chemotherapy. (C,D) The difference in OS between patients with multi hit mutation (n=7) of *TTN* and *TTN*-WT (n=116) in LUAD treated with chemotherapy, and between patients with multi hit mutation (n=7) of *TTN* and *TTN*-WT (n=116) in LUAD treated with chemotherapy, and between patients with multi hit mutation (n=7) of *TTN* and *TTN*-WT (n=42) in LUSC treated with chemotherapy, overall survival; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma; WT, wild-type.



**Figure S2** The association between *TTN* mutation and OS in platinum-treated LUAD and LUSC patients on chemotherapy. (A) The difference in OS between patients with *TTN* mutation (n=52) and *TTN*-WT (n=59) in LUAD patients treated with platinum-related drugs. (B) The difference in OS between patients with *TTN* mutation (n=62) and *TTN*-WT (n=23) in LUSC patients treated with platinum-related drugs. *TTN*, titin; OS, overall survival; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma; MT, mutant-type; WT, wild-type.

Table S	3 Sampl	e size of	patients with	n TTN and/or	<b>TP53</b>	mutation i	n LUAD	and LUSC
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Thoropy types	Numbers				
merapy types –	TTN+/TP53+	TTN+/TP53-	TTN-/TP53+		
LUAD					
Total	107	98	84		
Chemotherapy	33	38	30		
LUSC					
Total	218	108	90		
Chemotherapy	65	36	19		

"+" indicates MT; "-" indicates WT. *TTN*, titin; *TP53*, tumor protein 53; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma; MT, mutant-type; WT, wild-type.



**Figure S3** The association between *TTN* and/or *TP53* mutations and OS in LUAD patients treated with chemotherapy. The difference in OS period between TTN mutation and single mutation of TP53 (A), *TTN/TP53* double mutation and single mutation of *TTN* (B) in LUAD patients who were treated with chemotherapy. The difference in OS period between *TTN* mutation and single mutation of *TP53* (C), *TTN/TP53* double mutation and single mutation of *TTN* (D) in LUSC patients who were treated with chemotherapy. "+" indicates MT; "-" indicates WT. *TTN*, titin; *TP53*, tumor protein 53; OS, overall survival; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma; MT, mutant-type; WT, wild-type.



**Figure S4** The difference in OS between *TTN/TP53* double mutation and single mutation of *TP53* in LUSC and LUAD patients treated with chemotherapy. The difference in OS period between *TTN/TP53* double mutation and single mutation of *TP53* in LUAD patients (A) and in LUSC patients (B) who were treated with chemotherapy. "+" indicates MT; "-" indicates WT. OS, overall survival; *TTN*, titin; *TP53*, tumor protein 53; LUAD, lung adenocarcinoma; LUSC, lung squamous carcinoma; MT, mutant-type; WT, wild-type.