Characteristics	n= 275
Age, yrs	$61.3 \pm 10.3$
Male	206 (74.9)
Smoking (n=256)	
Current	113 (44.1)
Former	69 (27.0)
Never	74 (28.9)
Stage	
1	82 (29.8)
2	98 (35.6)
3	91 (33.1)
4	4 (1.5)
Lung cancer cell type	
Adenocarcinoma	134 (48.7)
Squamous cell carcinoma	134 (48.7)
Other	7 (2.5)

Table S1. Baseline characteristics of 275 cancer patients<sup>a</sup>

<sup>a</sup> Data are presented as numbers (percentages) unless otherwise indicated. Plus-minus values are means±standard deviation.

	Cytoplasmic expression			Nuclear expression		
Characteristics	Low (0-3) n=126	High (4-9) n=149	<i>P</i> -value	Low (0) n=184	High (≥1) n=91	<i>P</i> -value
Age, yrs	$61.9 \pm 9.8$	$61.4 \pm 10.0$	0.687	$61.7 \pm 9.7$	$61.6 \pm 10.3$	0.970
Male	94 (74.6)	112 (75.2)	0.914	138 (75.0)	68 (74.7)	0.961
Ever smokers	88 (75.2)	94 (67.6)	0.213	124 (72.5)	58 (68.2)	0.477
Size > 3cm	88 (47.6)	97 (52.4)	0.351	124 (67.0)	61 (33.0)	0.904
Stage						
I-II	90 (71.4)	90 (60.4)	0.055	128 (69.6)	52 (57.1)	0.042
III-IV	36 (28.6)	59 (39.6)	0.033	56 (30.4)	39 (42.9)	0.042
Pathology (n=268)						
Adenocarcinoma	49 (40.2)	85 (58.2)	0.002	82 (45.8)	52 (58.4)	0.052
Squamous cell	73 (59.8)	61 (41.8)	0.005	97 (54.2)	37 (41.6)	0.032

Table S2. Clinical and pathological characteristics of patients with high and low cytoplasmic or nuclear expression<sup>a</sup>

<sup>a</sup> Data are presented as numbers (percentages) unless otherwise indicated. Plus-minus values are means±standard deviation.

AIMP2: aminoacyl-tRNA synthetase interacting multi-functional protein 2, AIMP2-DX2: aminoacyl-tRNA synthetase interacting multi-functional protein 2–exon 2 deletion (AIMP2-DX2)

## **Fig. S1. Representative microphotographs of cytoplasmic and nuclear AIMP2-DX2 expression in lung cancer.** Expression of AIMP2-DX2 was evaluated using a scoring system that takes into account the product of

staining intensity and the percentage of positive cells. Staining intensity was classified as 0, 1, 2, and 3 and frequency was classified as 0 (<10%), 1 (10–50%), 2 (51–80%), and 3 (>80%).



Expression	Description			
grade	Intensity	Frequency		
0	negative	< 10%		
1	Weak	10-50 %		
2	Medium	51-79 %		
3	Strong	>80%		

**Fig. S2. Expression of AIMP2-DX2 in mouse and human lung tissue by immunohistochemistry and immunoblotting.** Representative microphotographs showing AIMP2-DX2 expression in (A) 8-week-old wildtype C57BL/6 mice, (B) Kras-LSL G12D and Kras-LSL G12D:p53<sup>fl/fl</sup> lung cancer model mice, (C) normalappearing human tissues adjacent to lung cancer tissues, and (D) different histologic types of human lung cancer. Immunoblots showing AIMP2-DX2 expression in (E) wild-type C57BL/6 mice and Kras-LSL G12D and Kras-LSL G12D:p53<sup>fl/fl</sup> lung cancer model mice and (F) 12 pairs of human lung cancer tissues and adjacent normal-appearing lung tissues. Immunohistochemical analysis of the distribution of AIMP2-DX2 expression in (G) the cytoplasm and (F) the nucleus of 275 human lung cancer tissues. AIMP2-DX2: aminoacyl-tRNA synthetase-interacting multi-functional protein 2–exon 2 deletion



**Fig. S3. Kaplan–Meier curve for overall survival and progression/recurrence-free survival in high and low AIMP2-DX2 expression in the cytoplasm and nucleus.** Samples with high expression of cytoplasmic AIMP2-DX2 include those with 4 or higher product of intensity and frequency of cytoplasmic AIMP2-DX2; samples with low expression include those with 3 or less. Samples with high expression of nuclear AIMP2-DX2 include those with 1 or higher product of intensity; samples with low expression include those with 0. *P*value was obtained by log-rank test. AIMP2-DX2: aminoacyl-tRNA synthetase-interacting multi-functional protein 2–exon 2 deletion.



Expression of	Median		
AIMP2-DX2	OS, months (95% CI)	PFS, months (95% CI)	
Cytoplasmic expression			
Low (0-3)	80.0 (44.9 - 115.1)	149.0	
High (4-9)	73.1 (48.7 - 97.5)	105.5 (20.7 - 190.3)	
Nuclear expression			
Low (0)	79.9 (46.9 - 112.9)	157.2 (106.3 - 208.1)	
High (≥ 1)	73.1 (41.0 - 105.2)	105.5	