

Supplementary materials: A Matching-Adjusted Indirect Comparison of Pembrolizumab + Chemotherapy vs Nivolumab + Ipilimumab as First-Line Therapies in Patients with PD-L1 TPS $\geq 1\%$ Metastatic NSCLC

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Table S1. Overall survival and progression-free survival in patients with PD-L1 TPS $\geq 1\%$ before matching.

	Outcomes	
	Overall Survival	Progression-Free Survival
ITC HR (95% CI), p-value^a		
Primary approach ^b	0.84 (0.63, 1.11), 0.218	0.58 (0.46, 0.74), <0.001
Secondary approach ^c	0.82 (0.64, 1.04), 0.102	0.62 (0.49, 0.77), <0.001
Number of events, (%)		
Pembrolizumab+chemotherapy	273 (57.6)	356 (75.1)
Nivolumab+ipilimumab	259 (65.4)	289 (73.0)
KN189/407/021G: Chemotherapy ^d	241 (70.5)	310 (90.6)
Checkmate 227: Chemotherapy ^d	299 (75.3)	286 (72.0)
Median Months, (95% CI)		
Pembrolizumab+chemotherapy	22.0 (19.7; 25.2)	9.2 (8.3; 10.9)
Nivolumab+ipilimumab	16.9 (15.0; 20.0)	5.0 (4.0; 6.2)
KN189/407/021G: Chemotherapy ^d	13.6 (10.9; 15.7)	4.9 (4.6; 6.0)
Checkmate 227: Chemotherapy ^d	14.9 (12.5; 16.8)	5.5 (4.8; 5.9)
Landmark rate (%) – Pembrolizumab+chemotherapy vs. Nivolumab+ipilimumab		
6-month	85.4 vs 75.2	70.2 vs 45.6
1-year	69.3 vs 62.4	43.5 vs 32.8
2-year	46.6 vs 39.8	26.7 vs 22.0

- a: Two-sided p-value calculated from the test statistic associated with the ITC estimate and its standard error
- b: Calculated using aggregate data published in the literature for nivolumab+ipilimumab. Bucher methodology using separate study results (estimate and its standard error) with a common control arm
- c: Calculated using pseudo-IPD from CheckMate227 Part 1A using a Cox regression model
- d: Platinum-doublet chemotherapy for KN021G/KN0189/KN407 and CheckMate227 Part 1A
- CI: confidence interval; ITC: indirect treatment comparison; HR: hazard ratio; ITT: Intention-to-Treat; KN021: KEYNOTE-021 Cohort G; KN189: KEYNOTE-189; KN407: KEYNOTE-407; PD-L1: Programmed cell death ligand 1; TPS: tumor proportion score.

Table S2. Landmark analysis of overall survival and progression-free survival for chemotherapy arms from KN021G/KN189/KN407 and CheckMate 227 Part 1a before and after matching.

	Landmark rate 6 month, %	Landmark rate 1-year, %	Landmark rate 2-year, %
Overall survival			
Before Matching	76.5 vs 78.4	53.3 vs 56.3	34.3 vs 33.3
After Matching	76.4 vs 78.4	53.9 vs 56.3	32.4 vs 33.3
Progression-free survival			
Before Matching	44.6 vs 42.6	20.3 vs 18.8	8.0 vs 7.3
After Matching	44.1 vs 42.6	20.2 vs 18.8	4.7 vs 7.3

KN021G: KEYNOTE-021 Cohort G; KN189: KEYNOTE-189; KN407: KEYNOTE-407.

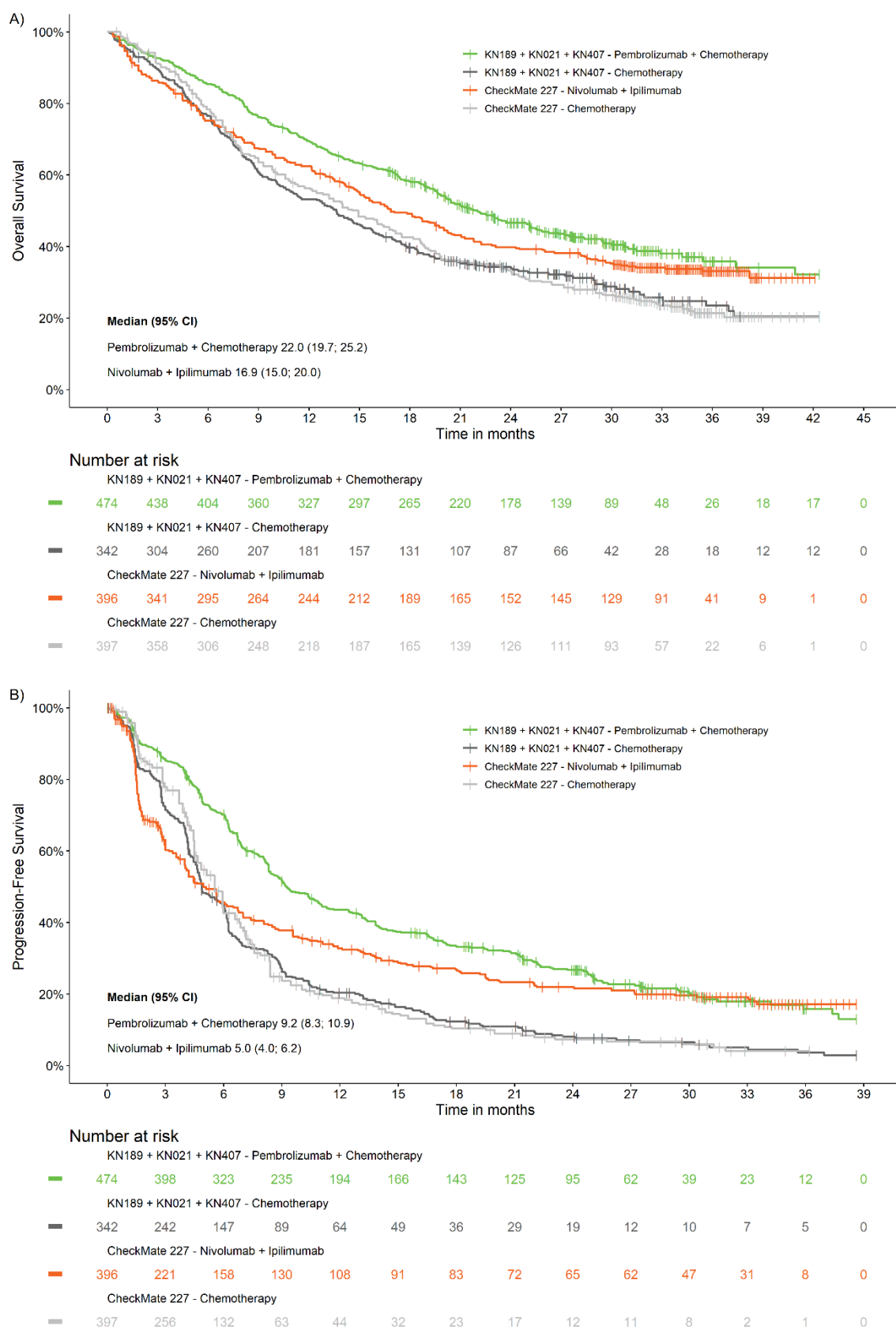


Figure S1. Kaplan–Meier curves before matching adjustment for a) overall survival b) progression-free survival.

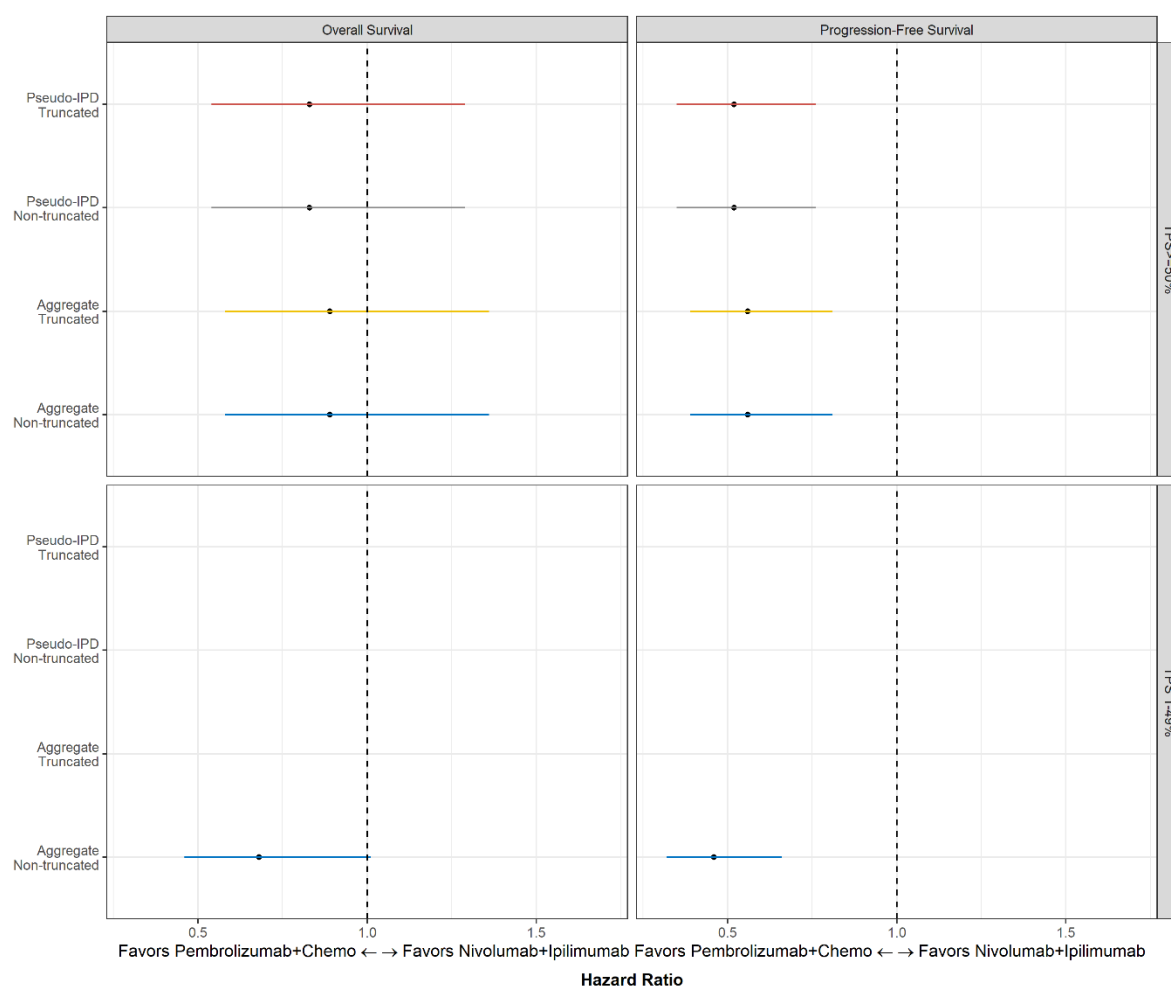


Figure S2. Forest plot of pembrolizumab+chemotherapy versus nivolumab+ipilimumab for overall survival and progression-free survival in patients with PD-L1 TPS 1–49% and $\geq 50\%$ ^a.

^a: The results illustrated with the yellow lines refer to the base case analysis.

IPD: individual patient data; PD-L1: Programmed cell death ligand 1; TPS: tumor proportion score.

Table S3. Pembrolizumab versus nivolumab+ipilimumab for overall survival and progression-free survival in patients with PD-L1 TPS $\geq 1\%$, TPS $\geq 50\%$ and TPS 1–49%.

	Hazard Ratio (95% CI)	
	Before Matching	After Matching
Overall Survival, $\geq 1\%$		
Aggregate, truncated ^a	0.84 (0.63, 1.11)	0.80 (0.59, 1.09)
Pseudo IPD, truncated	0.82 (0.64, 1.04)	0.75 (0.57, 0.99)

Aggregate, non-truncated	0.84 (0.64, 1.11)	0.80 (0.59, 1.09)
Pseudo IPD, non-truncated	0.82 (0.64, 1.04)	0.75 (0.57, 0.99)
Progression-Free Survival (BICR), $\geq 1\%$		
Aggregate, truncated ^a	0.58 (0.46, 0.74)	0.53 (0.41, 0.68)
Pseudo IPD, truncated	0.62 (0.49, 0.77)	0.55 (0.42, 0.71)
Aggregate, non-truncated	0.58 (0.46, 0.74)	0.53 (0.41, 0.68)
Pseudo IPD, non-truncated	0.62 (0.49, 0.78)	0.55 (0.42, 0.71)
Overall Survival, $\geq 50\%$		
Aggregate, truncated ^a	0.90 (0.63, 1.29)	0.89 (0.58, 1.36)
Pseudo IPD, truncated	0.87 (0.60, 1.24)	0.83 (0.54, 1.29)
Aggregate, non-truncated	0.90 (0.63, 1.29)	0.89 (0.58, 1.36)
Pseudo IPD, non-truncated	0.87 (0.60, 1.24)	0.83 (0.54, 1.29)
Progression-Free Survival (BICR), $\geq 50\%$		
Aggregate, truncated ^a	0.63 (0.45, 0.88)	0.56 (0.39, 0.81)
Pseudo IPD, truncated	0.63 (0.45, 0.89)	0.52 (0.35, 0.76)
Aggregate, non-truncated	0.63 (0.45, 0.88)	0.56 (0.39, 0.81)
Pseudo IPD, non-truncated	0.63 (0.45, 0.89)	0.52 (0.35, 0.76)
Overall Survival, 1-49%		
Aggregate, non-truncated	0.71 (0.50, 1.01)	0.68 (0.46, 1.01)
Progression-Free Survival (BICR), 1-49%		
Aggregate, non-truncated	0.49 (0.35, 0.69)	0.46 (0.32, 0.66)

^a: The aggregate/truncated analysis corresponds to the base case analysis and is included for comparison purposes.

BICR: blinded independent committee review; CI: confidence intervals; IPD: individual patient data; PD-L1: Programmed cell death ligand 1; TPS: tumor proportion score.