Supplementary Online Content

- Freeman JQ, Huo D, Shubeck SP, et al. Trends and disparities in the use of immunotherapy for triple-negative breast cancer in the US. *JAMA Netw Open*. 2025;8(2):e2460243. doi:10.1001/jamanetworkopen.2024.60243
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This supplementary material has been provided by the authors to give readers additional information about their work.

eTable 1. Overall Characteristics of Triple-Negative Breast Cancer in the 2019-2021 National Cancer Database

	National Cancer	National Cancer	Overall
	Database, 2019-2021	Database, 2021	
	Stage IV	Stage II-III	Stage II, III and IV
Characteristic	(n=3069)	(n=7655)	(N=10724)
	No. (%)	No. (%)	No. (%)
Age at diagnosis			
Mean (SD)	59.8 (13.7)	54.5 (13.5)	56.1 (13.7)
Median (IQR)	60.0 (51.0, 70.0)	55.0 (44.0, 65.0)	56.0 (46.0, 66.0)
Race/Ethnicity			
Asian or Pacific Islander	97 (3.2)	376 (4.9)	473 (4.4)
Black	848 (27.6)	1721 (22.5)	2569 (24.0)
Hispanic	241 (7.9)	740 (9.7)	981 (9.1)
White	1828 (59.6)	4637 (60.6)	6465 (60.3)
Other	55 (1.8)	181 (2.4)	236 (2.2)
Histologic type			
Ductal	2279 (74.3)	7041 (92.0)	9320 (86.9)
Ductal and lobular	36 (1.2)	61 (0.8)	97 (0.9)
Inflammatory breast cancer	362 (11.8)	198 (2.6)	560 (5.2)
Lobular	90 (2.9)	75 (1.0)	165 (1.5)
Metaplastic breast cancer	72 (2.3)	177 (2.3)	249 (2.3)
Other	230 (7.5)	103 (1.3)	333 (3.1)
Tumor grade			
1	18 (0.7)	47 (0.6)	65 (0.6)
2	459 (17.7)	1127 (15.0)	1586 (15.7)
3	2113 (81.6)	6349 (84.4)	8462 (83.7)
Charlson-Deyo Comorbidity Index			
0	2427 (79.1)	6320 (82.6)	8747 (81.6)
1	442 (14.4)	964 (12.6)	1406 (13.1)
≥2	200 (6.5)	371 (4.8)	571 (5.3)
Percent no high school degree quartiles a			

651 (25.2)	1291 (20.6)	1942 (22.0)
()	` ,	` '
737 (28.5)	1588 (25.4)	2325 (26.3)
660 (25.5)	1791 (28.6)	2451 (27.7)
538 (20.8)	1582 (25.3)	2120 (24.0)
617 (23.9)	1103 (17.7)	1720 (19.5)
582 (22.6)	1284 (20.6)	1866 (21.1)
543 (21.0)	1421 (22.8)	1964 (22.3)
838 (32.5)	2435 (39.0)	3273 (37.1)
475 (15.5)	941 (12.3)	1416 (13.2)
1160 (37.8)	1995 (26.1)	3155 (29.4)
69 (2.2)	166 (2.2)	235 (2.2)
1212 (39.5)	4376 (57.2)	5588 (52.1)
153 (5.0)	177 (2.3)	330 (3.1)
2616 (86.7)	6548 (87.4)	9164 (87.2)
45 (1.5)	89 (1.2)	134 (1.3)
357 (11.8)	853 (11.4)	1210 (11.5)
1023 (36.3)	2178 (33.9)	3201 (34.6)
209 (7.4)	386 (6.0)	595 (6.4)
1033 (36.7)	2349 (36.5)	3382 (36.6)
550 (19.5)	1515 (23.6)	2065 (22.3)
	660 (25.5) 538 (20.8) 617 (23.9) 582 (22.6) 543 (21.0) 838 (32.5) 475 (15.5) 1160 (37.8) 69 (2.2) 1212 (39.5) 153 (5.0) 2616 (86.7) 45 (1.5) 357 (11.8) 1023 (36.3) 209 (7.4) 1033 (36.7) 550 (19.5)	660 (25.5) 1791 (28.6) 538 (20.8) 1582 (25.3) 617 (23.9) 1103 (17.7) 582 (22.6) 1284 (20.6) 543 (21.0) 1421 (22.8) 838 (32.5) 2435 (39.0) 475 (15.5) 941 (12.3) 1160 (37.8) 1995 (26.1) 69 (2.2) 166 (2.2) 1212 (39.5) 4376 (57.2) 153 (5.0) 177 (2.3) 2616 (86.7) 6548 (87.4) 45 (1.5) 89 (1.2) 357 (11.8) 853 (11.4) 1023 (36.3) 2178 (33.9) 209 (7.4) 386 (6.0) 1033 (36.7) 2349 (36.5)

Abbreviations: SD, standard deviation; IQR, interquartile range; No., number.

^a Defined as education attainment for patient residence areas and measured by matching the zip code of the patient recorded at the time of diagnosis against files derived from the 2016 American Community Survey data.

^b Based on the 2016 American Community Survey data, spanning years 2012–2016 and adjusted for 2016 inflation.

^c Measured by matching the state and county FIPS code of the patient recorded at the time of diagnosis against 2013 files published by the United States Department of Agriculture Economic Research Service.

eTable 2. Estimated Prevalence of Immunotherapy Use among Patients With Triple-Negative Breast Cancer in the 2017-2021 National Cancer Database, by Stage and Year of Diagnosis, Overall, and by Race and Ethnicity

		Overall	Asian or Pacific Islander	Black	Hispanic	White	Other
Characteristic		Yes	Yes	Yes	Yes	Yes	Yes
		No. (%)	No. (%)	No. (%)	No. (%)	No. (%)	No. (%)
Stage IV							
	Year of diagnosis						
	2017	54 (5.5)	1 (3.7))	12 (4.6)	4 (5.9)	36 (6.0)	1 (5.0)
	2018	91 (9.2)	5 (17.2)	18 (6.8)	7 (9.3)	59 (9.7)	2 (15.4)
	2019	291 (28.4)	5 (23.8)	67 (22.2)	16 (24.6)	194 (31.4)	9 (42.9)
	2020	337 (32.7)	18 (42.9)	80 (28.5)	27 (31.0)	210 (35.0)	2 (9.5)
	2021	393 (38.8)	13 (38.2)	91 (34.3)	38 (42.7)	245 (40.1)	6 (46.2)
Stage II-III			, ,				
	Year of diagnosis						
	2017	211 (4.2)	11 (5.5)	41 (3.7)	20 (4.4)	137 (4.3)	2 (2.5)
	2018	323 (6.5)	23 (12.2)	51 (4.6)	31 (6.4)	211 (6.8)	7 (8.2)
	2019	345 (6.5)	26 (11.2)	48 (4.1)	33 (6.4)	233 (7.0)	5 (5.4)
	2020	700 (13.0)	40 (16.3)	106 (9.2)	68 (13.2)	473 (14.0)	13 (11.4)
	2021	3662 (48.0)	201 (53.5)	788 (46.0)	358 (48.5)	2228 (48.2)	87 (48.3)

Abbreviation: No., number.

eTable 3. Estimated Prevalence of Immunotherapy Use among Patients With Triple-Negative Breast Cancer, by Stage and Race/Ethnicity

		Receipt of immun	otherapy		No. of days immunother		tial diagnosis and	start of
Characteris	tic	No (chemotherapy alone)	Yes (chemotherapy and immunotherapy)					
		No. (row %)	No. (row %)	P value ^a	Mean (SD)	P value a	Median (IQR)	P value a
	ncer Database, 20						1	
Stage IV	Overall	2048 (66.7)	1021 (33.3)		54.6 (39.0)		46.0 (33.0, 67.0)	
(n=3069)	Race/Ethnicity							
	Asian or			.004	44.0 (18.9)	<.001	41.5 (30.0, 56.5)	.002
	Pacific Islander	61 (62.9)	36 (37.1)					
	Black	610 (71.9)	238 (28.1)		60.4 (34.0)		51.0 (37.0, 77.0)	
	Hispanic	160 (66.4)	81 (33.6)		51.6 (30.0)		44.0 (33.0, 58.0)	
	White	1179 (64.5)	649 (35.5)		52.5 (33.1)		46.0 (31.0, 63.0)	
	Other	38 (69.1)	17 (30.9)		91.6 (165.7)		52.0 (35.0, 60.0)	
National Ca	ncer Database, 20	021						
Stage II-III	Overall	3972 (52.0)	3662 (48.0)		67.7 (69.4)		42.0 (29.0, 74.0)	
(n=7655)	Race/Ethnicity							
	Asian or			.11	72.5 (69.6)	<.001	45.0 (29.0-86.5)	<.001
	Pacific Islander	175 (46.5)	201 (53.5)					
	Black	927 (54.1)	788 (45.9)		77.4 (79.1)]	48.0 (33.0-84.0)]
	Hispanic	380 (51.5)	358 (48.5)		76.3 (71.9)		49.5 (33.0-92.0)	
	White	2397 (51.8)	2228 (48.2)		62.8 (65.4)]	40.0 (28.0-65.0)]
	Other	93 (51.7)	87 (48.3)		58.1 (50.2)		43.0 (25.0-76.0)	

Abbreviations: No., number; SD, standard deviation; IQR, interquartile range; API.

^a P values were calculated using ANOVA, Kruskal-Wallis, or Pearson's X² tests, as appropriate.

eTable 4. Association between Immunotherapy Use and Pathologic Complete Response among Patients With Stage II-III, Triple-Negative Breast Cancer in the 2021 National Cancer Database

	Pathologic complete response			Logistic regression	
	Did not achieve	Achieved			
Characteristic	(n=4749 [62.0%])	(n=2906 [38.0%])			
	No. (row %)	No. (row %)	P value a	AOR (95% CI) b	P value
Receipt of immunotherapy					
No (neoadjuvant chemotherapy alone)	2641 (66.5)	1331 (33.5)	<.001	1 [Reference]	
Yes (neoadjuvant chemotherapy and immunotherapy)	2095 (57.2)	1567 (42.8)		1.45 (1.31-1.61)	<.001
Age at diagnosis					
Mean (SD)	56.0 (13.5)	52.9 (13.3)	<.001	0.83 (0.80-0.86) ^c	<.001
Median (IQR)	57.0 (46.0, 66.0)	53.0 (43.0, 63.0)	<.001	NA	
Race/Ethnicity					
Asian or Pacific Islander	232 (61.7)	144 (38.3)	.04	0.88 (0.70-1.11)	.29
Black	1104 (64.2)	617 (35.8)		0.90 (0.80-1.02)	.11
Hispanic	425 (57.4)	315 (42.6)		1.06 (0.89-1.25)	.54
White	2878 (62.1)	1759 (37.9)		1 [Reference]	
Other	110 (60.8)	71 (39.2)		1.05 (0.76-1.45)	.78
Histologic type					
Ductal	4276 (60.7)	2765 (39.3)	<.001	1 [Reference]	
Ductal and lobular	46 (75.4)	15 (24.6)		0.55 (0.28-1.06)	.07
Inflammatory breast cancer	141 (71.2)	57 (28.8)		0.84 (0.56-1.27)	.41
Lobular	61 (81.3)	14 (18.7)		0.72 (0.38-1.34)	.30
Metaplastic breast cancer	147 (83.1)	30 (16.9)		0.38 (0.25-0.59)	<.001
Other	78 (75.7)	25 (24.3)		0.65 (0.39-1.09)	.10
Tumor grade					
1	40 (85.1)	7 (14.9)	<.001	1 [Reference]	
2	857 (76.0)	270 (24.0)		1.42 (0.61-3.29)	.42
3	3763 (59.3)	2586 (40.7)		2.69 (1.17-6.18)	.02
Charlson-Deyo Comorbidity Index					
0	3843 (60.8)	2477 (39.2)	<.001	1 [Reference]	

1	635 (65.9)	329 (34.1)		0.90 (0.77-1.06)	.21
≥2	271 (73.1)	100 (27.0)		0.73 (0.56-0.94)	.02
AJCC clinical T stage					
cT2	3232 (60.5)	2109 (39.5)	<.001	1 [Reference]	
cT3	783 (68.8)	355 (31.2)		0.67 (0.58-0.78)	<.001
cT4	386 (70.4)	162 (29.6)		0.73 (0.56-0.93)	.01
AJCC clinical nodal status					
Negative (cN0)	2478 (60.6)	1613 (39.4)	<.001	1 [Reference]	
Positive (cN1+)	1917 (65.5)	1011 (34.5)		0.84 (0.75-0.93)	.001

Abbreviations: SD, standard deviation; IQR, interquartile range; No., number; AOR, adjusted odds ratio; CI, confidence interval; AJCC, American Joint Committee on Cancer.

^a P values were generated based on Student's t, Wilcoxon rank-sum, or Pearson's X^2 tests, as appropriate.

^b This multivariable logistic regression model included all variables in the table, including receipt of immunotherapy, age at diagnosis, race/ethnicity, histologic type, tumor grade, and the Charlson-Deyo Comorbidity index.

^c Age at diagnosis was considered a continuous variable and therefore, the corresponding AOR and 95% CI are listed per 10-year increase in age.

eTable 5. Assessment of Racial and Ethnic Differences in Pathologic Complete Response among Patients With Stage II-III, Triple-Negative Breast Cancer Who Received Immunotherapy and Neoadjuvant Chemotherapy in the 2021 National Cancer Database

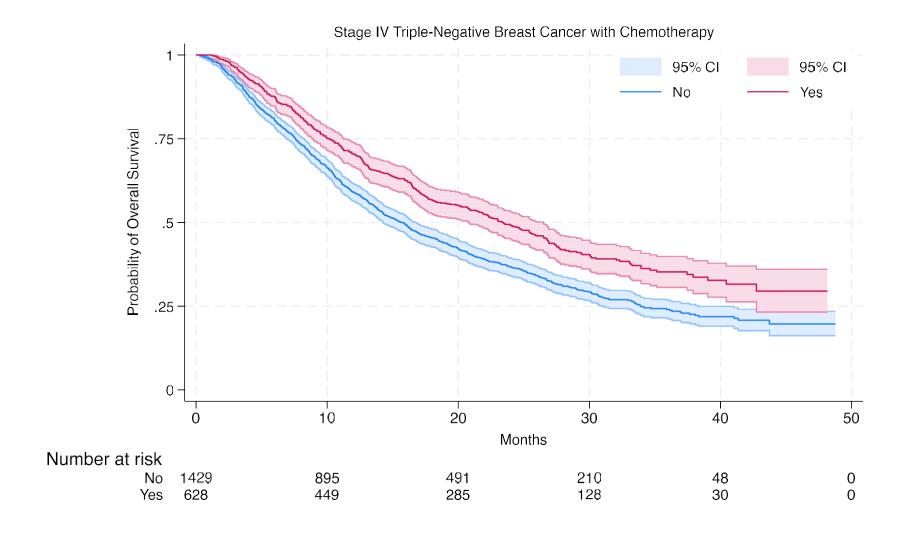
	Pathologic comp	lete response	Logistic regressi	ogistic regression		
	Did not achieve	Achieved	Model 1		Model 2	
Characteristic	(n=2095 [57.2%])	(n=1567 [42.8%])				
	No. (row %)	No. (row %)	AOR (95% CI)	P value	AOR (95% CI) ^a	P value
Race/Ethnicity						
Asian or Pacific Islander	121 (60.2)	80 (39.8)	0.76 (0.55-1.04)	.08	0.82 (0.53-1.26)	.37
Black	469 (59.5)	319 (40.5)	0.88 (0.74-1.06)	.17	0.92 (0.73-1.17)	.49
Hispanic	189 (52.8)	169 (47.2)	1 [Reference]		1 [Reference]	
White	1272 (57.1)	956 (42.9)	1.31 (0.83-2.05)	.25	1.09 (0.60-1.96)	.78
Other	44 (50.6)	43 (49.4)	0.86 (0.81-0.91)	<.001	0.84 (0.74-0.94)	.004
Age at diagnosis, per 10-year increase			0.76 (0.55-1.04)	.08	0.82 (0.53-1.26)	.37
Histologic type						
Ductal	1,874 (55.8)	1485 (44.2)	1 [Reference]		1 [Reference]	
Ductal and lobular	24 (75.0)	8 (25.0)	0.41 (0.16-1.03)	.06	0.38 (0.12-1.19)	.10
Inflammatory breast cancer	70 (63.6)	40 (36.4)	1.08 (0.64-1.83)	.77	0.99 (0.52-1.89)	.97
Lobular	27 (79.4)	7 (20.6)	0.65 (0.27-1.56)	.34	0.42 (0.14-1.26)	.12
Metaplastic breast cancer	69 (85.2)	12 (14.8)	0.28 (0.15-0.53)	<.001	0.37 (0.18-0.75)	.006
Other	31 (67.4)	15 (32.6)	0.72 (0.36-1.45)	.36	0.54 (0.21-1.39)	.20
Tumor grade						
1	15 (83.3)	3 (16.7)	1 [Reference]		1 [Reference]	
2	353 (69.8)	153 (30.2)	1.67 (0.45-6.19)	.45	1.01 (0.20-5.24)	.99
3	1691 (54.8)	1395 (45.2)	2.86 (0.78-10.5)	.11	1.75 (0.34-8.96)	.50
Charlson-Deyo Comorbidity Index					·	
0	1742 (56.4)	1347 (43.6)	1 [Reference]		1 [Reference]	
1	245 (58.8)	172 (41.3)	0.99 (0.79-1.24)	.92	0.99 (0.74-1.32)	.94
≥2	108 (69.2)	48 (30.8)	0.74 (0.51-1.09)	.12	0.85 (0.54-1.33)	.47
AJCC clinical T stage		, , ,			,	
cT2	1346 (55.0)	1102 (45.0)	1 [Reference]		1 [Reference]	

cT3	400 (65.1)	214 (34.9)	0.66 (0.55-0.81)	<.001	0.72 (0.56-0.92)	.01
cT4	184 (65.9)	95 (34.1)	0.64 (0.45-0.91)	.01	0.83 (0.54-1.33)	.37
AJCC clinical nodal status						
Negative (cN0)	991 (54.8)	816 (45.2)	1 [Reference]		1 [Reference]	
Positive (cN1+)	935 (61.2)	593 (38.8)	0.83 (0.72-0.97)	.02	0.86 (0.71-1.25)	.11

Abbreviations: No., number; AOR, adjusted odds ratio; CI, confidence interval; AJCC, American Joint Committee on Cancer.

^a Additionally adjusted for percent no high school degree quartile, median household income quartile, type of health insurance, rural-urban area, and facility type.

eFigure 1. Kaplan-Meier Curves of Overall Survival among Patients With Stage IV, Triple-Negative Breast Cancer with Chemotherapy in the 2019-2021 National Cancer Database, by Immunotherapy Use



eTable 6. Kaplan-Meier Estimate of Median Overall Survival Time among Patients With Stage IV, Triple-Negative Breast Cancer with Chemotherapy in the 2019-2021 National Cancer Database

Characteristic	No. of patients	No. of events (%)	Median survival time in months ^a (95% CI)	P value ^a
Overall	2057	1297	17.3 (16.4-18.7)	
Receipt of immunotherapy				
No (chemotherapy alone)	1429	948 (66.3)	15.6 (14.3-16.8)	<.001
Yes (chemotherapy and immunotherapy)	628	349 (55.6)	23.4 (20.8-26.5)	
Among patients who received immunotherapy				
Histologic type				
Ductal	465	246 (52.9)	25.4 (22.3-27.8)	.05
Ductal and lobular	7	5 (71.4)	16.5 (8.5)	
Inflammatory breast cancer	81	53 (65.4)	16.5 (11.9-23.3)	
Lobular	16	11 (68.8)	18.4 (7.4-23.5)	
Metaplastic breast cancer	18	9 (50.0)	33.1 (13.2)	
Other	41	25 (61.0)	20.7 (8.3)	

Abbreviations: IQR, interquartile range, No., number; CI, confidence interval.

^a *P* value was generated using the log-rank test.

eTable 7. Immunotherapy Use and Overall Survival among Patients With Stage IV, Triple-Negative Breast Cancer with Chemotherapy in the 2019-2021 National Cancer Database

	Cox proportional hazards regression					
Characteristic	Model 1		Model 2			
	aHR (95% CI)	P value	aHR (95% CI) a	P value		
Receipt of immunotherapy						
Yes (chemotherapy and immunotherapy)	0.76 (0.66-0.87)	<.001	0.80 (0.68-0.94)	.006		
No (chemotherapy alone)	1 [Reference]		1 [Reference]			
Age at diagnosis, per 10-year increase	1.02 (0.98-1.07)	.34	1.12 (1.03-1.22)	.007		
Race and ethnicity						
Asian or Pacific Islander	0.90 (0.61-1.32)	.58	0.85 (0.53-1.37)	.50		
Black	1.07 (0.93-1.22)	.34	1.03 (0.87-1.22)	.70		
Hispanic	0.88 (0.68-1.13)	.30	0.74 (0.52-1.04)	.08		
White	1 [Reference]		1 [Reference]			
Other	0.95 (0.60-1.50)	.81	1.39 (0.84-2.30)	.20		
Histologic type						
Ductal	1 [Reference]		1 [Reference]			
Ductal and lobular	1.29 (0.78-2.13)	.32	1.31 (0.69-2.48)	.40		
Inflammatory breast cancer	1.27 (1.07-1.51)	.006	1.31 (1.07-1.60)	.008		
Lobular	1.08 (0.72-1.64)	.70	0.90 (0.55-1.45)	.66		
Metaplastic breast cancer	0.65 (0.40-1.08)	.10	0.89 (0.53-1.50)	.67		
Other	1.62 (1.19-2.21)	.002	1.52 (1.05-2.19)	.03		
Tumor grade						
1	1 [Reference]		1 [Reference]			
2	2.14 (0.68-6.71)	.19	2.56 (0.63-10.43)	.19		
3	2.14 (0.69-6.68)	.19	2.57 (0.64-10.38)	.19		
Charlson-Deyo Comorbidity Index						
0	1 [Reference]		1 [Reference]			
1	1.12 (0.94-1.32)	.21	1.14 (0.94-1.38)	.18		
≥2	1.52 (1.20-1.93)	.001	1.46 (1.12-1.92)	.01		
Type of health insurance						

Medicaid	NA	NA	1.36 (1.10-1.69)	.004
Medicare	NA	NA	0.95 (0.77-1.18)	.65
Other government/unknown	NA	NA	1.17 (0.74-1.87)	.50
Private/managed care	NA	NA	1 [Reference]	
Uninsured	NA	NA	1.41 (1.01-1.96)	.04
Type of cancer program				
Academic/research	NA	NA	1 [Reference]	
Community	NA	NA	1.24 90.92-1.67)	.15
Comprehensive community	NA	NA	1.30 (1.10-1.53)	.002
Integrated network	NA	NA	1.08 (0.88-1.32)	.45

Abbreviations: No., number; aHR, adjusted hazard ratio; CI, confidence interval; NA, not applicable.

^a Additionally adjusted for percent no high school degree quartile, median household income quartile, and rural-urban area.