

Supplemental Tables for:
Sorafenib Effectiveness in Advanced Hepatocellular Carcinoma
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Supplemental Table 1: Covariate Balance Following Propensity Score Matching

Characteristic	Before Matching		p-value	After Matching		P-value
	No Treatment n=565 (70%)	Sorafenib N=242 (30%)		No Treatment N=223	Sorafenib N=223	
Year of Diagnosis						
2008	141 (25%)	44 (18%)	0.003	43 (19%)	43 (19%)	0.855
2009	164 (29%)	53 (22%)		56 (25%)	52 (23%)	
2010	131 (23%)	66 (27%)		64 (29%)	60 (27%)	
2011	129 (23%)	79 (33%)		60 (27%)	68 (30%)	
Age						
Median (Q1, Q3)	74 (67, 82)	70 (64, 77)		71 (64, 79)	70 (64, 78)	
less than 64	119 (21%)	63 (26%)	<.001	56 (25%)	58 (26%)	0.890
65-74 yr	181 (32%)	99 (41%)		93 (42%)	88 (39%)	
75+ yr	265 (47%)	80 (33%)		74 (33%)	77 (35%)	
Sex						
Male	380 (67%)	186 (77%)	0.006	162 (73%)	168 (75%)	0.517
Female	185 (33%)	56 (23%)		61 (27%)	55 (25%)	
Race						
White	346 (61%)	139 (57%)	0.467	138 (62%)	128 (57%)	0.743
African American	67 (12%)	38 (16%)		30 (13%)	36 (16%)	
Asian	73 (13%)	29 (12%)		23 (10%)	27 (12%)	
Other	79 (14%)	36 (15%)		32 (14%)	32 (14%)	
Marital Status						
Married	208 (37%)	115 (48%)	0.011	98 (44%)	102 (46%)	0.858
Separated/Divorced/Widowed	231 (41%)	71 (29%)		72 (32%)	69 (31%)	
Single	100 (18%)	43 (18%)		53 (24%)	40 (18%)	
Other/Unknown	26 (5%)	13 (5%)		†	12 (5%)	
Census Tract Region						
Metro	482 (85%)	221 (91%)	0.053	201 (90%)	202 (91%)	0.966
Urban	35 (6%)	21 (9%)		22 (10%)	21 (9%)	
Rural	48 (8%)	†		†	†	
Census Tract % Below Poverty Line						
1 st Quartile (lowest)	119 (21%)	67 (28%)	0.118	66 (30%)	59 (26%)	0.409

2 nd Quartile	137 (25%)	51 (21%)		48 (22%)	48 (22%)	
3 rd Quartile	158 (28%)	56 (23%)		39 (17%)	53 (24%)	
4 th Quartile (highest)	141 (25%)	67 (28%)		70 (31%)	63 (28%)	
Modified Charlson Score						
0	52 (9%)	28 (12%)	0.129	20 (9%)	24 (11%)	0.817
1	76 (13%)	43 (18%)		39 (17%)	38 (17%)	
2+	437 (77%)	171 (71%)		164 (74%)	161 (72%)	
Number of Liver Comorbidities						
Mean (STD)	0.9 (1.2)	0.6 (1.1)	0.001	0.6 (0.9)	0.7 (1.1)	0.816
Prior AFP Screening						
0	360 (64%)	133 (55%)	0.054	137 (61%)	126 (57%)	0.537
1	102 (18%)	58 (24%)		43 (19%)	51 (23%)	
2+	103 (18%)	51 (21%)		43 (19%)	46 (21%)	
Hepatitis B Virus	42 (7%)	26 (11%)	0.121	21 (9%)	19 (9%)	0.740
Hepatitis C Virus	174 (31%)	76 (31%)	0.864	64 (29%)	71 (32%)	0.471
Alcoholic Liver Disease	86 (15%)	31 (13%)	0.373	24 (11%)	29 (13%)	0.464
Other Cause of Liver Disease	76 (13%)	27 (11%)	0.371	26 (12%)	26 (12%)	1.000
Tumor Extent						
Multiple lesions, no vascular invasion	231 (41%)	117 (48%)	<.001	107 (48%)	104 (47%)	0.852
Multiple lesions with vascular invasion	93 (16%)	57 (24%)		48 (22%)	53 (24%)	
Extension beyond liver, NOS	241 (43%)	68 (28%)		68 (30%)	66 (30%)	
Maximum Tumor Size						
<= 3cm	61 (11%)	35 (14%)	<.001	30 (14%)	34 (15%)	0.939
3 to 5cm	74 (13%)	†		†	†	
> 5cm	195 (35%)	123 (51%)		107 (48%)	108 (48%)	
Unknown	235 (42%)	84 (35%)		86 (39%)	81 (36%)	

† n<11. Cell merged with above to preserve confidentiality.

Table S2: Characteristics of Initial versus Subsequent Sorafenib Users

A	Initial Sorafenib N=422	Subsequent Sorafenib N=232	p-value
Year of Diagnosis			
2008	90 (21%)	54 (23%)	0.680*
2009	106 (25%)	65 (28%)	
2010	100 (24%)	52 (22%)	
2011	126 (30%)	61 (26%)	
First Line Therapy			
TACE	NA	149 (64%)	NA
Y90 Radioembolization		15 (6%)	
Curative surgery		29 (13%)	
Ablative surgery		16 (7%)	
Radiation (any)		23 (9%)	
Other drug therapy		+	
Age			
Median (Q1, Q3)	70 (63, 77)	70 (62, 76)	
less than 64	112 (27%)	65 (28%)	0.837*
65-74 yr	168 (40%)	94 (41%)	
75+ yr	142 (34%)	73 (31%)	
Sex			
Male	311 (74%)	154 (66%)	0.048*
Female	111 (26%)	78 (34%)	
Race			
White	254 (60%)	127 (55%)	0.282*
African American	58 (14%)	31 (13%)	
Asian	52 (12%)	41 (18%)	
Other	58 (14%)	33 (14%)	
Marital Status			
Married	192 (45%)	107 (46%)	0.635*
Separated/Divorced/Widowed	128 (30%)	78 (34%)	
Single	77 (18%)	47 (20%)	
Other/Unknown	25 (6%)	+	
Residence Area			
Metro	378 (90%)	209 (90%)	0.977*
Urban	15 (4%)	23 (9%)	

Rural	29 (7%)	†	
Census Tract % <Poverty			
1 st Quartile (lowest)	104 (25%)	63 (27%)	0.290*
2 nd Quartile	99 (24%)	66 (29%)	
3 rd Quartile	109 (26%)	54 (23%)	
4 th Quartile (highest)	108 (26%)	48 (21%)	
Modified Charlson Score			
0	45 (11%)	24 (10%)	0.881*
1	77 (18%)	39 (17%)	
2+	300 (71%)	169 (73%)	
Mean (STD) # of Liver Comorbidites	0.7 (1.0)	0.7 (1.1)	0.959**
Prior AFP Screening			
0	242 (57%)	95 (41%)	<.001*
1	99 (23%)	55 (24%)	
2+	81 (19%)	82 (35%)	
Hepatitis B Virus	41 (10%)	32 (14%)	0.113*
Hepatitis C Virus	133 (32%)	98 (42%)	0.01
Alcoholic Cirrhosis	58 (14%)	34 (15%)	0.75
Other Cause of Liver Disease	48 (11%)	30 (13%)	0.56
Tumor Extent			
Multiple lesions, no vascular invasion	210 (50%)	146 (63%)	0.005*
Multiple lesions with vascular invasion	96 (23%)	38 (16%)	
Extension beyond liver, NOS	116 (27%)	48 (21%)	

† n<11. Cell merged with above to preserve confidentiality.

Table S3. Effect of Initial Sorafenib Compared to No Treatment on the Risk of Death From the Time of Diagnosis of HCC. Rates of death by treatment group and the mortality Risk Ratio (RR) are shown for multiple time points from the date of diagnosis.

Time Point	MODEL	No Treatment		Sorafenib		Risk Difference	RR, 95% CI	p-value
		Events/n	% Deceased	Events/n	% Deceased			
3 Months	Unmatched	124/565	22%	52/242	21%	1%	0.98, 0.74-1.30	0.89
	PS Matched	53/223	24%	51/223	23%	1%	0.96, 0.69-1.35	0.82
6 Months	Unmatched	307/565	54%	124/242	51%	3%	0.94, 0.82-1.09	0.43
	PS Matched	127/223	57%	118/223	53%	4%	0.93, 0.79-1.10	0.39
12 Months	Unmatched	434/565	77%	184/242	76%	1%	0.99, 0.91-1.08	0.81
	PS Matched	177/223	79%	174/223	78%	1%	0.98, 0.89-1.08	0.73

Table S4A, Sensitivity Analysis of the Effect of Varying Landmarks (30, 60, 90 days) on Mortality Risk Ratios Anchored at Diagnosis in the Propensity Score Matched Cohort

Model	Treatment group	3 months from Diagnosis		6 months from Diagnosis		12 months from Diagnosis	
		Events/n	RR, 95% CI	Events/n	RR, 95% CI	Events/n	RR, 95% CI
30 day	None	74/150	0.86, 0.68- 1.11	97/150	1.00, 0.85- 1.18	119/150	1.05, 0.94, 1.17
	Sorafenib	64/150		97/150		125/150	
60 day	None	53/223	0.96, 0.69-1.35	127/223	0.93, 0.79-1.10	177/223	0.98, 0.89-1.08
	Sorafenib	51/223		118/223		174/223	
90 day	None	0/181	na	81/181	0.86, 0.68- 1.10	125/181	1.07, 0.94-1.22
	Sorafenib	0/181		70/181		134/181	

Table S4B, Sensitivity Analysis of the Effect of Varying Landmarks (30, 60, 90 days) on Mortality Risk Ratios Anchored at 60 days Post-Diagnosis in the Propensity Score Matched Cohort

Model	Treatment group	3 months from 60d Anchor		6 months from 60d Anchor		12 months from 60d Anchor	
		Events/n	RR, 95% CI	Events/n	RR, 95% CI	Events/n	RR, 95% CI
30 day	None	49/150	1.00, 0.72-1.38	64/150	1.16, 0.90-1.48	85/150	1.05, 0.86-1.27
	Sorafenib	49/150		74/150		89/150	
60 day	None	113/223	0.88, 0.72-1.07	153/223	0.97, 0.86-1.11	188/223	0.95, 0.87-1.04
	Sorafenib	99/223		149/223		179/223	
90 day	None	65/181	0.77, 0.57-1.04	98/181	1.04, 0.86-1.25	135/181	1.04, 0.92-1.16
	Sorafenib	50/181		102/181		140/181	

