

# Postprogression survival after lenvatinib treatment for advanced hepatocellular carcinoma



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## INTRODUCTION

Lenvatinib treatment has been established as first line systemic chemotherapy for advanced hepatocellular carcinoma (HCC), however, sequential treatment after lenvatinib remains obscure. As for multi-targeting agent (MTA) treatment for advanced HCC, it is well-known that not progression-free survival (PFS) but postprogression survival (PPS) is well correlated with overall survival (OS). PPS in the patients treated with lenvatinib remains obscure.

## AIM

To investigate PPS for advanced HCC treated with lenvatinib.

## METHOD

Patients: Sixty advanced HCC patients treated with lenvatinib at our institution (3 patients participated REFLECT trial and 57 patients were treated in a real-world practice between July 2018 and December 2019) were reviewed.

- Assessment
- ✓ Clinical characteristics
  - ✓ radiological progression pattern
  - ✓ following treatment after progression disease
  - ✓ Overall survival
  - ✓ Postprogression survival

## RESULTS

Table 1. Characteristics at starting lenvatinib

		n = 44
age	median	74 (68-78)
gender	male/female	36/8
ECOG PS	0/1	38/6
etiology	B/C/NBNC	7/12/25
BCLC	B/C	16/28
Extrahepatic spread		23 (52.2%)
Macrovascular invasion		11 (25.0%)
Child-Pugh	5/6/7/8/9	24/12/3/4/1
	A/B	36/8
ALBI	score	-2.42 (-2.75--1.95)
	grade 1/2a/2b/3	17/10/14/3
AFP		64.2 (8.6-164.2)
L3		21.4 (6.7-56.1)
DCP		
treatment line	1/2/3	29/8/7

Figure 1. Kaplan-Meier survival curve

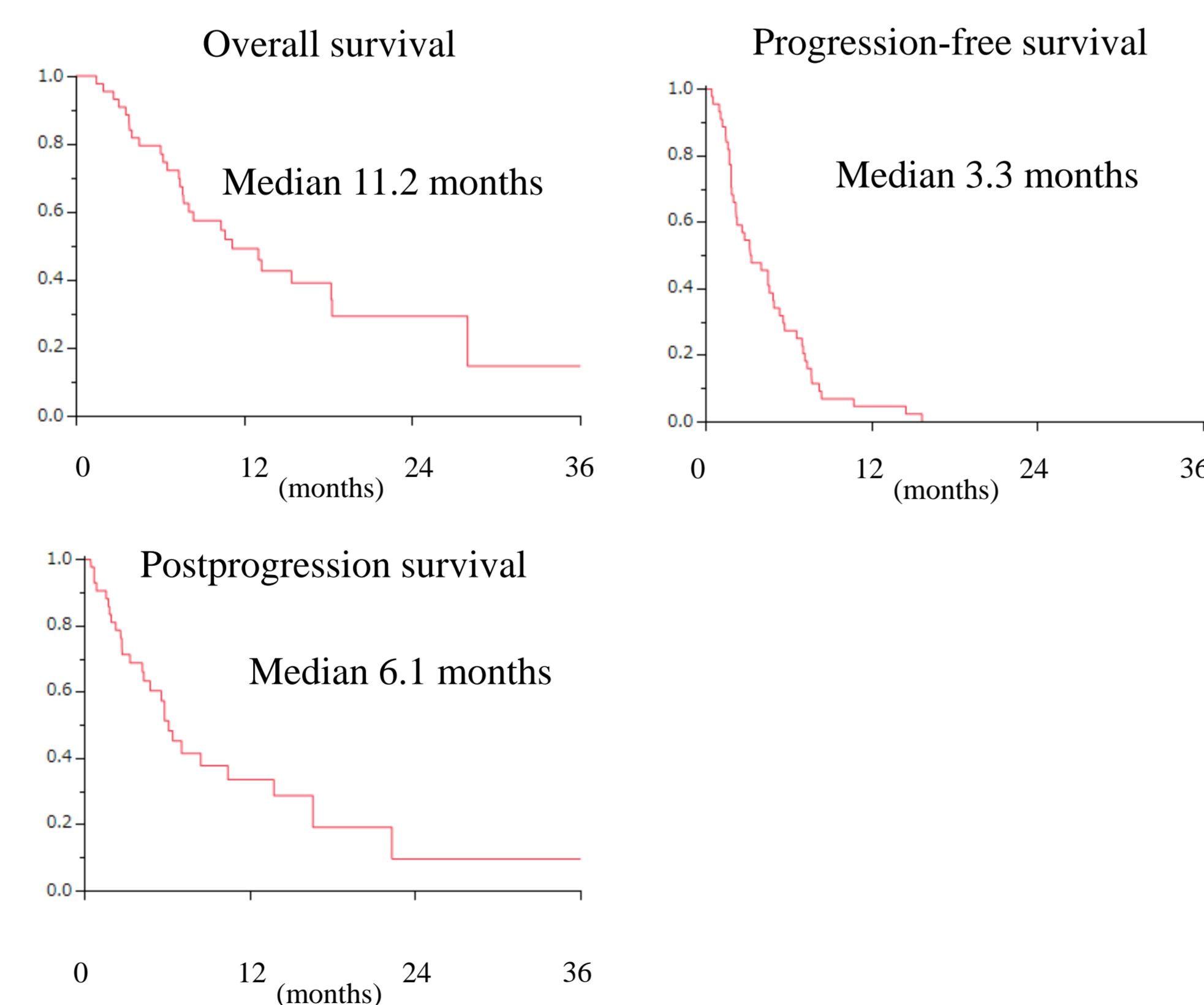


Table 2. Changes at radiological disease progression

		n (%)
impairment of PS	+1	24 (54.5%)
impairment of Child-Pugh	≥+1	19 (43.2%)
	≥+2	9 (20.5%)
radiologic progression pattern	NIH	19 (43.2%)
	IHG	30 (68.2%)
	NEH	13 (29.5%)
	EHG	15 (34.1%)

NIH, new intrahepatic lesion; IHG, intrahepatic growth; NEH, new extrahepatic lesion; EHG, extrahepatic growth

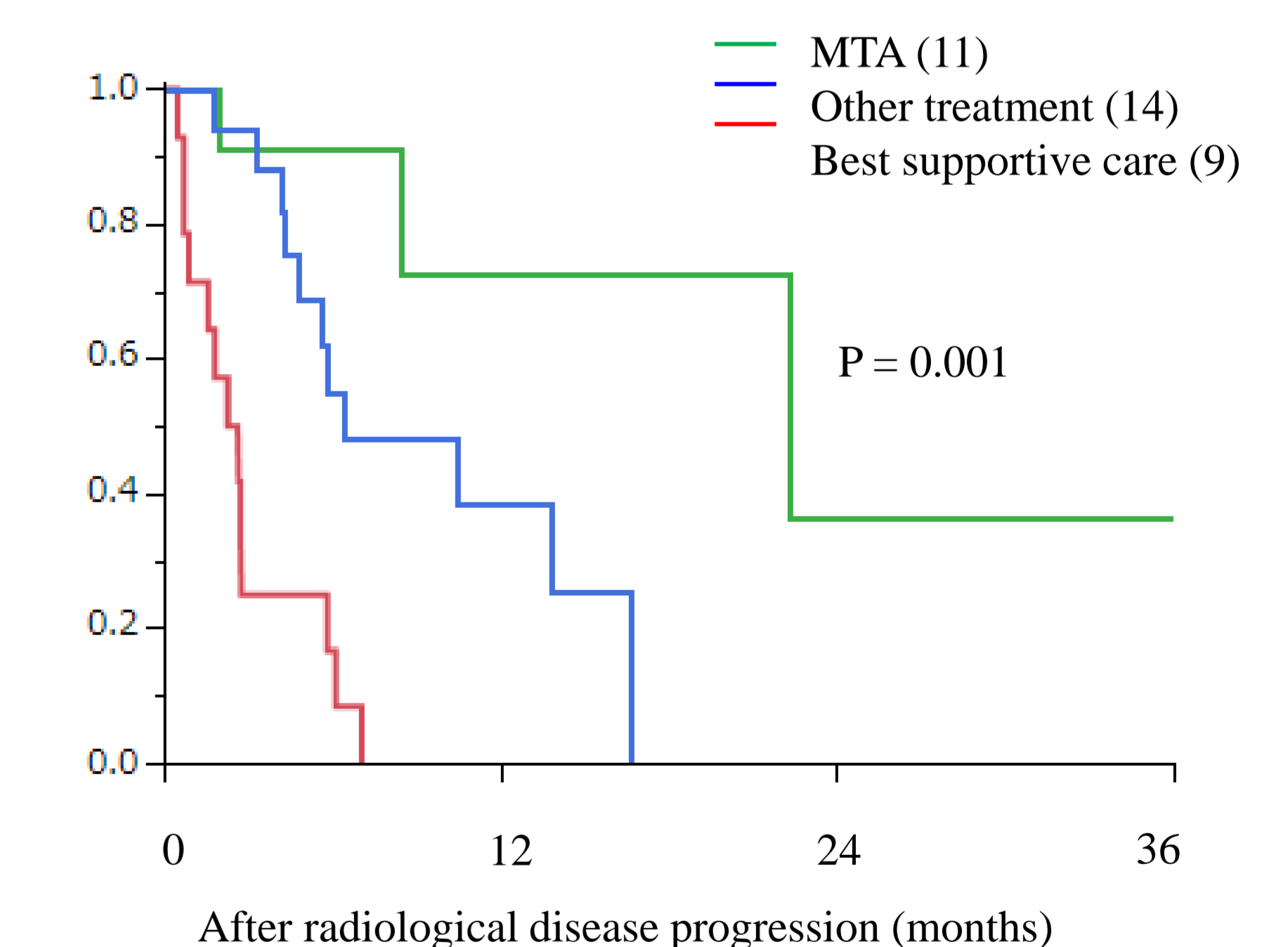
Table 3. predictive factors for overall survival

variables	univariate			multivariate		
	HR	95% CI	p-value	HR	95% CI	p-value
age	≥70	1.23(0.54-2.84)	0.61			
gender	male	1.57(0.47-5.27)	0.47			
ECOG PS	1	3.88(1.42-10.5)	0.008	<b>9.63(2.41-38.5)</b>		<b>0.001</b>
impairment of PS		2.17(0.92-5.09)	0.08	0.77(0.21-2.80)		0.7
etiology	HBV	0.78(0.29-2.09)	0.62			
	HCV	1.29(0.54-3.09)	0.56			
BCLC	C	2.18(0.94-5.00)	0.07	1.45(0.44-4.81)		0.54
EHS		1.17(0.55-2.49)	0.69			
MVI		3.69(1.64-8.31)	0.002			
Child-Pugh	B	3.71(1.59-8.65)	0.002	1.32(0.36-4.81)		0.67
impairment of Child-Pugh	≥1	1.57(0.72-3.41)	0.26	0.74(0.20-2.77)		0.66
ALBI	1or2a	3.59(1.63-7.91)	0.002	0.43(0.08-2.26)		0.32
treatment line	1st line	0.73(0.34-1.59)	0.43			
following treatment	yes	0.18(0.08-0.42)	<0.001	<b>1.17(0.04-0.66)</b>		<b>0.01</b>
progression pattern	NIH	2.05(0.93-4.50)	0.07	<b>3.77(1.38-10.3)</b>		<b>0.01</b>
	IHG	2.01(0.81-5.00)	0.13			
	NEH	1.19(0.51-2.73)	0.69			
	EHG	1.3(0.59-2.86)	0.51			

Table 4. predictive factors for postprogression survival

variables	univariate		multivariate		
	HR95% CI	p-value	HR95% CI	p-value	
age	≥70	1.25(0.54-2.89)	0.61		
gender	male	1.21(0.36-4.07)	0.76		
ECOG PS	1	2.39(0.87-6.54)	0.09	<b>4.02(1.06-15.17)</b>	<b>0.04</b>
impairment of PS		2.46(1.03-5.85)	0.042	0.47(0.08-2.74)	0.4
etiology	HBV	0.7(0.26-1.91)	0.48		
	HCV	0.75(0.31-1.78)	0.51		
BCLC	C	2.16(0.93-5.00)	0.07	2.17(0.70-6.72)	0.18
Child-Pugh	B	3.86(1.64-9.08)	0.002	0.97(0.27-3.54)	0.97
impairment of Child-Pugh	≥1	1.92(0.86-4.26)	0.11		
ALBI	1or2a	0.32(0.14-0.70)	0.0048	2.99(0.57-15.80)	0.2
treatment line	1st line	0.65(0.30-1.42)	0.28		
following treatment	yes	0.14(0.06-0.35)	<0.0001	<b>0.09(0.02-0.58)</b>	<b>0.01</b>
progression pattern	NIH	2.31(1.03-5.16)	0.04	<b>4.91(1.75-13.77)</b>	<b>0.0025</b>
	IHG	1.56(0.63-3.88)	0.34		
	NEH	1.18(0.50-2.77)	0.71		
	EHG	1.09(0.49-2.40)	0.83		

Figure 2. Comparison of postprogression survival among following treatments



MTA treatment was administered to 11 patients (regorafenib, ramucirumab, and sorafenib 1, 1, and 9 patients, respectively). The PPS of 22.3 months of patients treated with sequential MTA was higher than that of patients with other treatments (PPS, 10.4 months) or BSC (PPS, 2.5 months) (p<0.0001).

## CONCLUSIONS

OS and PPS are influenced by the progression pattern, which are important in prognostic predictions. Moreover, sequential MTA treatment after lenvatinib may improve OS and PPS.

## CONTACT INFORMATION

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