Fig.1 Change in levels of blood biomarkers after E7080 treatment

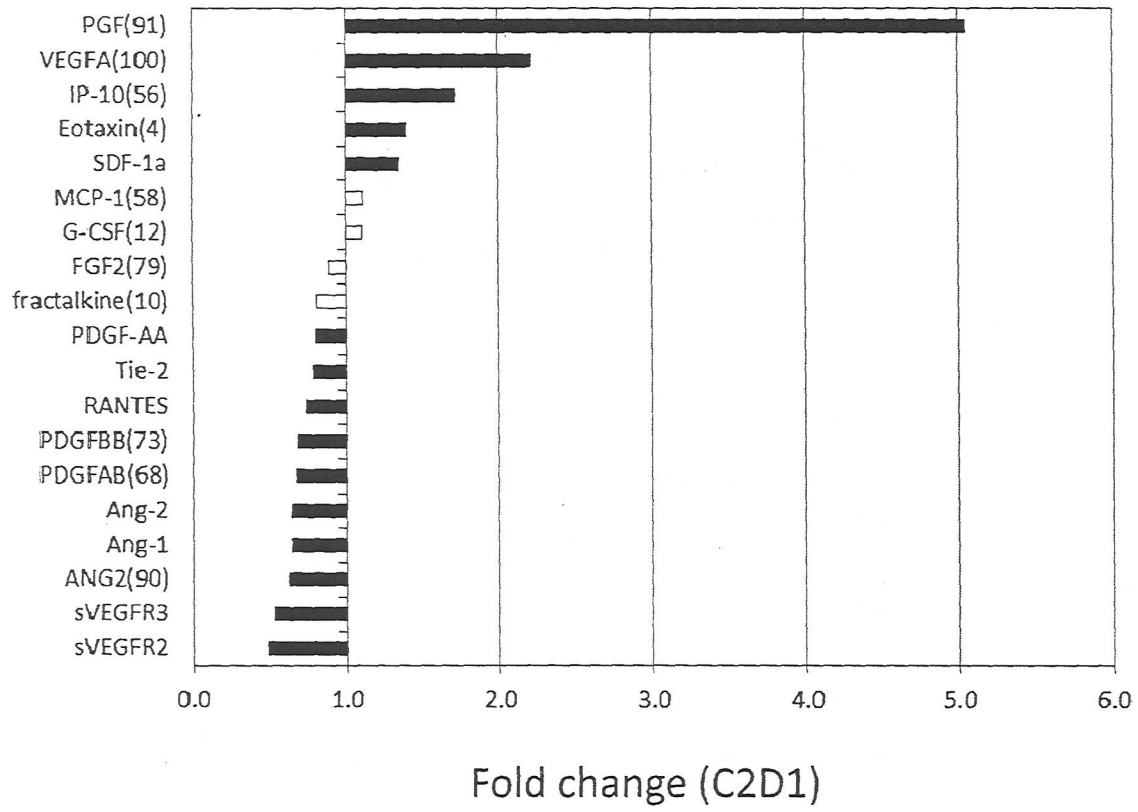
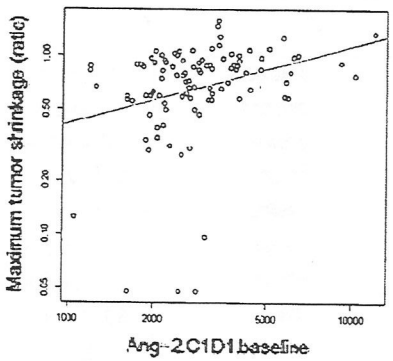


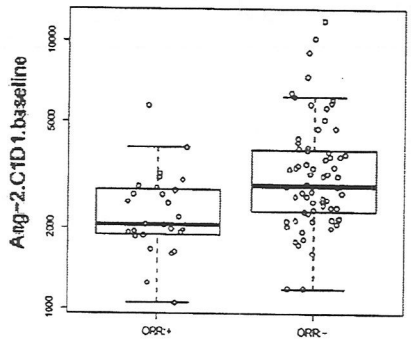
Fig. 2 Correlation of baseline CAFs with tumor response

Ang-2 baseline

%MTS

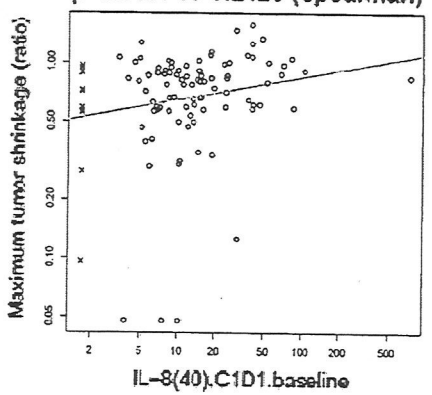


ORR



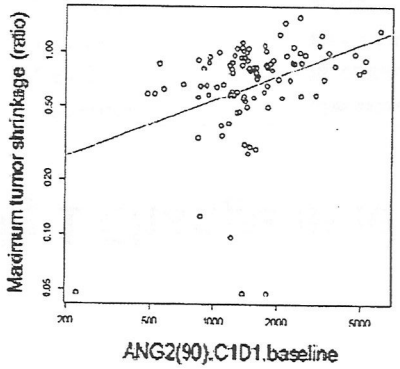
IL-8 baseline

%MTS



ANG(90) baseline

%MTS



ORR

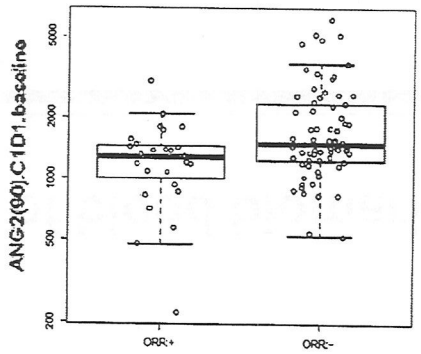


Fig. 3 No significant correlation of weight, age and histology with PFS and OS

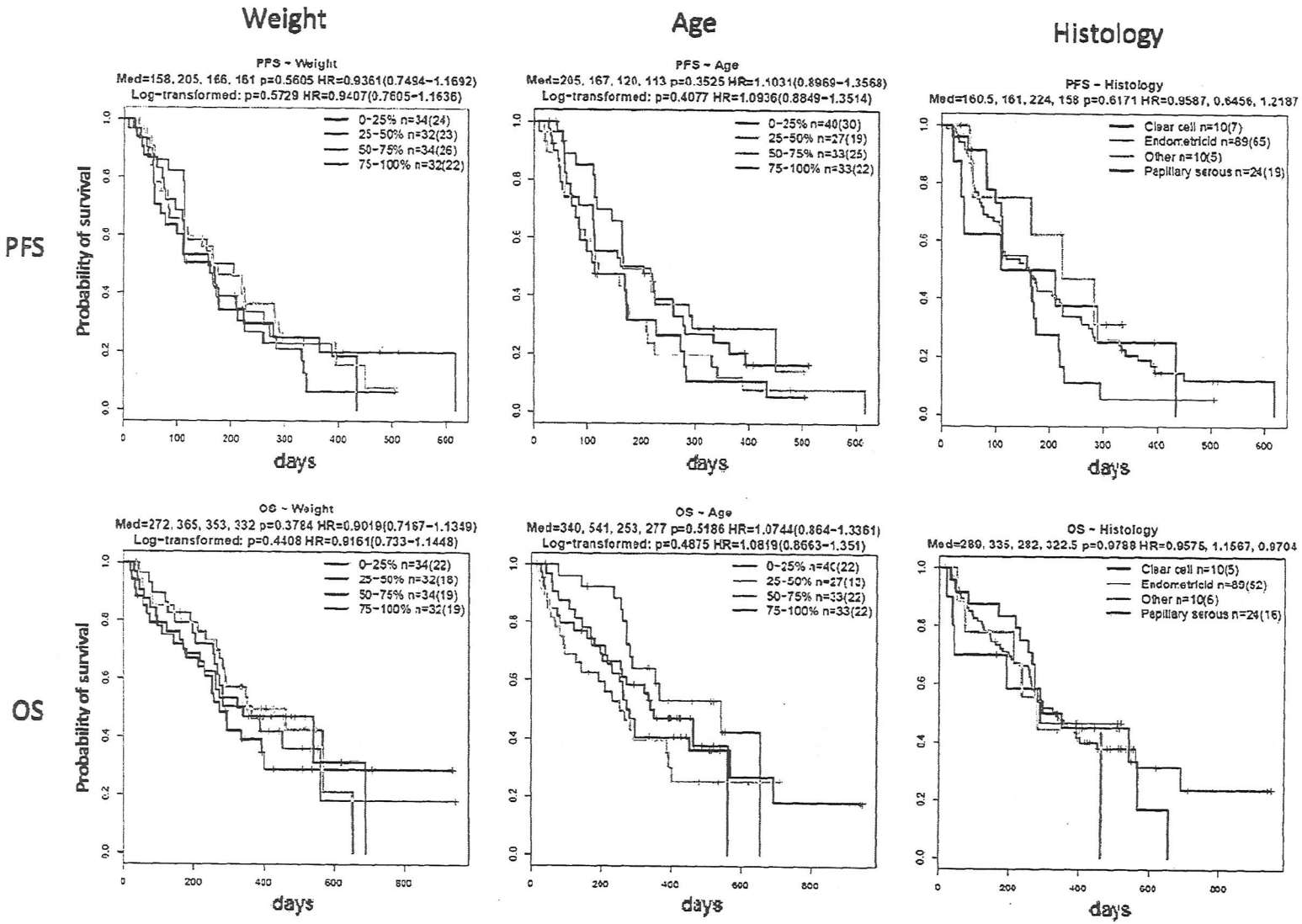


Fig. 4 Multivariate analysis does not identify potential combination factors with Ang-2

MTS

	Estimate	Std. Error	t value	Pr(>|t|)	
(Intercept)	-23.269	2.850	-8.163	1.14e-12	***
`Ang-2.C1D1.baseline`	12.517	3.592	3.485	0.000738	***

ORR

	bbbEstimate	Std. Error	z value	Pr(>|z|)	
(Intercept)	-1.4782	0.3643	-4.057	4.96e-05	***
`Ang-2.C1D1.baseline`	-1.6398	0.6193	-2.648	0.00811	**

PFS

	coef	exp(coef)	se(coef)	z	Pr(>|z|)	
`Ang-2.C1D1.baseline`	0.6422	1.9007	0.1099	5.846	5.04e-09	***

OS

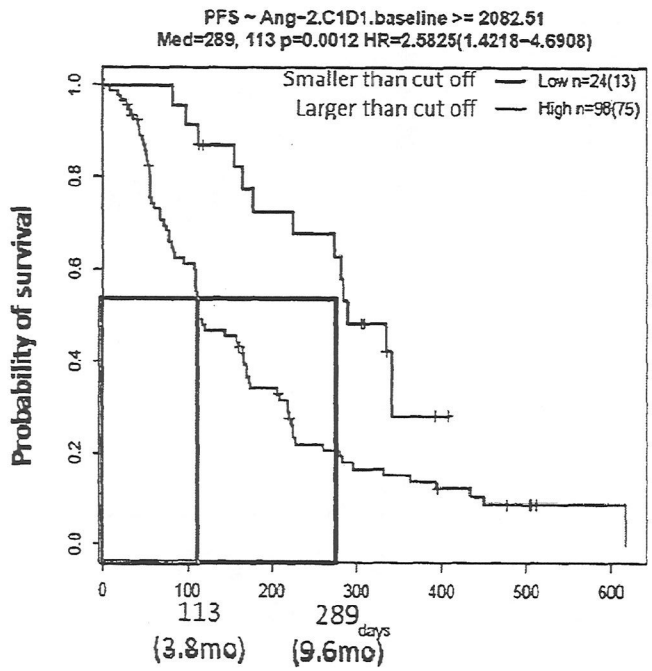
	coef	exp(coef)	se(coef)	z	Pr(>|z|)	
`Ang-2.C1D1.baseline`	0.53042	1.69964	0.11519	4.605	4.13e-06	***
`FGF-2(6).C1D1.baseline`	0.33595	1.39927	0.11400	2.947	0.00321	**
`PGF(91).C1D1.baseline`	0.19499	1.21530	0.08356	2.334	0.01962	*
`HGF(86).C1D1.baseline`	0.18837	1.20728	0.09339	2.017	0.04369	*

Variables use in analyses

WEIGHT
AGE
Ang-1
Ang-2
ANG2(90)
FGF-2(6)
FGF4(75)
HGF(86)
IL-8(40)
PDGF-AA
PDGFAB(68)
PDGFBB(73)
PGF(91)
Tie-2
VEGF(86)
VEGFA(100)
VEGFD(78)

Fig. 5 mPFS and mOS of sub-groups stratified by baseline Ang-2 level

PFS and OS: based on cut off (2082.5 pg/ml)



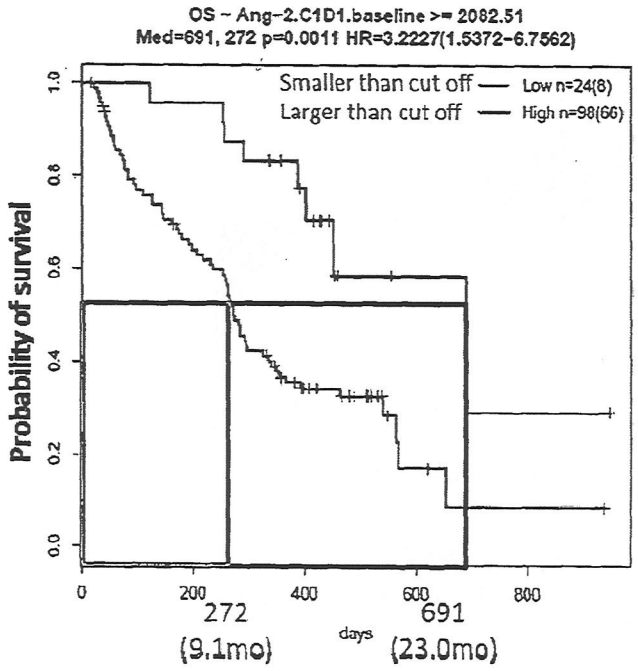
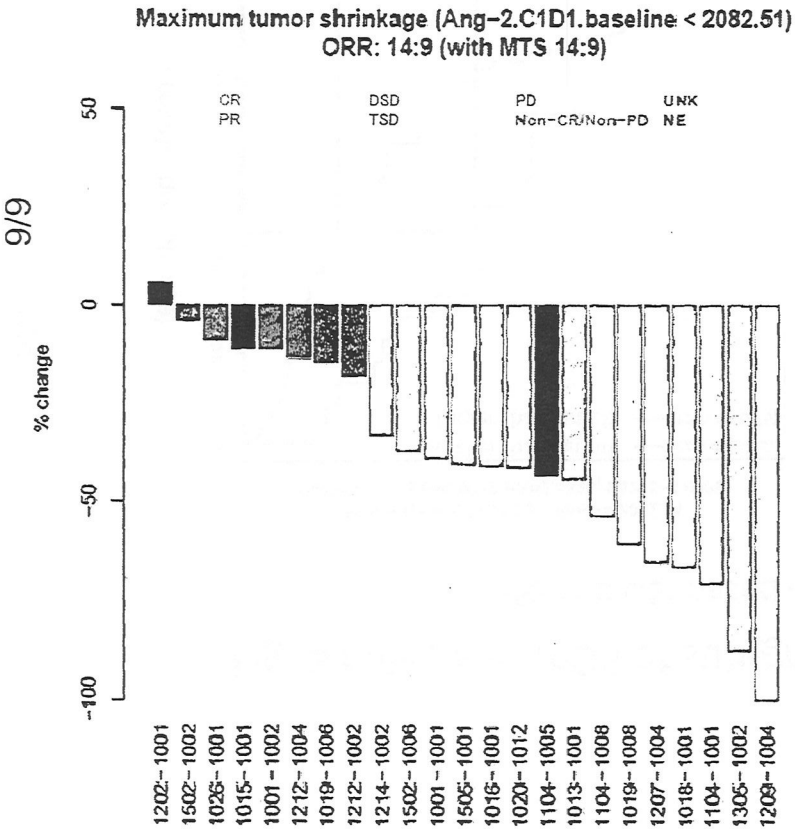


Fig. 6 Enrichment of patients population with better ORR based on baseline Ang-2 level

Ang-2 < 2082.5 pg/ml (23/96 = 24%)
ORR = 60.9%



Ang-2 > 2082.5 pg/ml (73/96 = 76%)
ORR = 17.8%

