



## **Tumori del rene e della vescica**

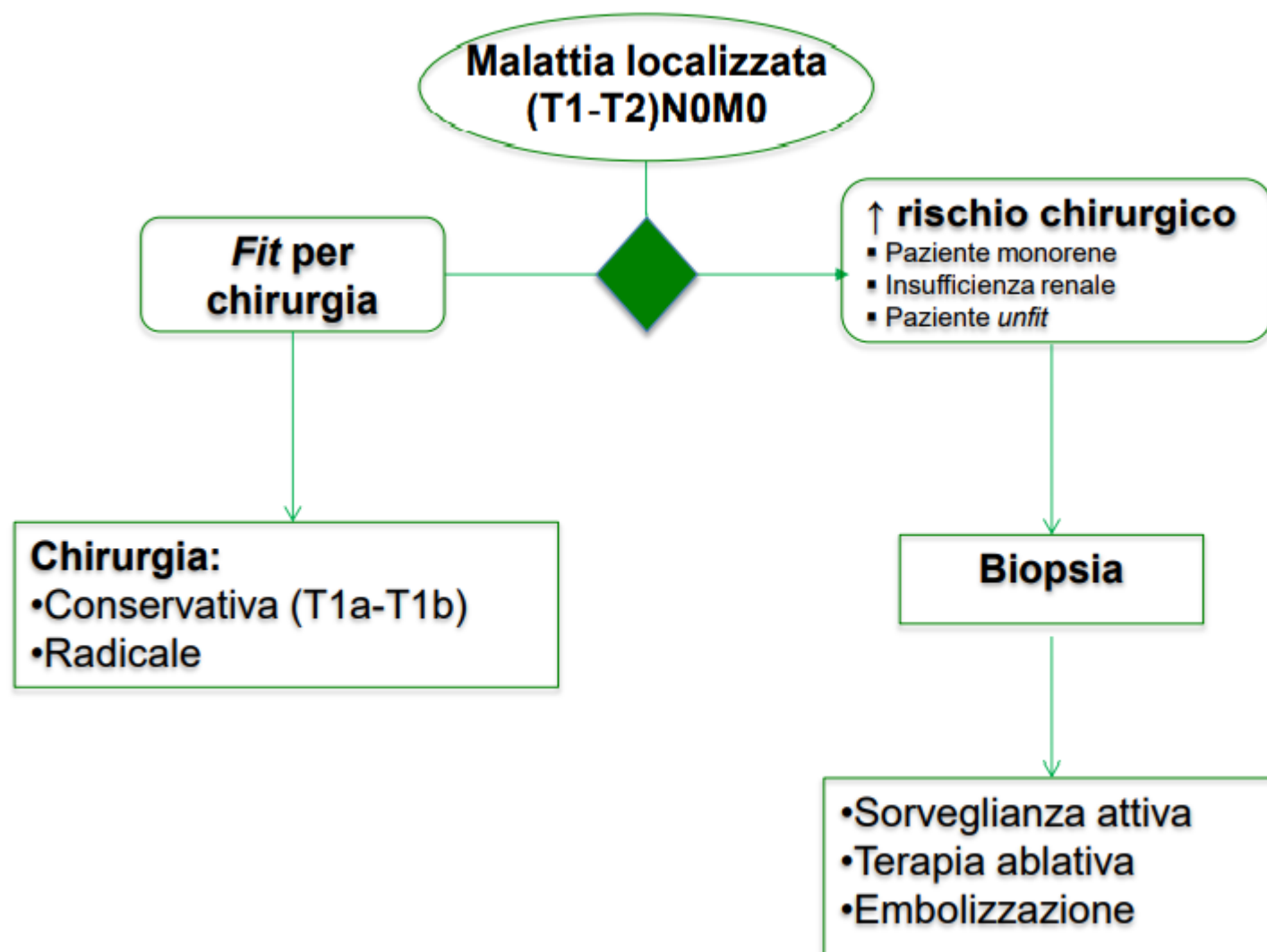
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[Emilia.cocorocchio@gavazzeni.it](mailto:Emilia.cocorocchio@gavazzeni.it)

Sede	Maschi	Femmine	Totale
	N. (%)	N. (%)	N. (%)
Vie Aero Digestive Superiori -VADS*	7.276 (3,7)	2.580 (1,4)	9.856 (2,6)
Esofago	1.710 (0,9)	684 (0,4)	2.394 (0,6)
Stomaco	8.458 (4,3)	6.098 (3,4)	14.556 (3,9)
Colon-Retto	23.420 (12,0)	20.282 (11,2)	43.702 (11,6)
Fegato	8.978 (4,6)	4.034 (2,2)	13.012 (3,5)
Pancreas	6.847 (3,5)	7.416 (4,1)	14.263 (3,8)
Colecisti e vie biliari	2400 (1,2)	3.000 (1,7)	5.400 (1,4)
Polmone	27.554 (14,1)	13.328 (7,3)	40.882 (10,9)
Melanomi	8.147 (4,2)	6.716 (3,7)	14.863 (4,0)
Mesotelioma	1.523 (0,8)	463 (0,3)	1.986 (0,5)
Mammella		54.976 (30,3)	54.976 (14,6)
Ovaio		5.179 (2,8)	5.179 (1,4)
Utero (cervice)		2.365 (1,3)	2.365 (0,6)
Utero (corpo)		8.335 (4,6)	8.335 (2,2)
Prostata	36.074 (18,5)		36.074 (9,6)
Testicolo	2.289 (1,2)		2.289 (0,6)
Rene, vie urinarie**	9.049 (4,6)	4.472 (2,5)	13.521 (3,6)
Vescica***	20.477 (10,5)	5.015 (2,8)	25.492 (6,8)
Sistema Nervoso Centrale	3.533 (1,8)	2.589 (1,4)	6.122 (1,6)
Tiroide	3.333 (1,7)	9.850 (5,4)	13.183 (3,5)
Linfomi di Hodgkin	1.222 (0,6)	929 (0,5)	2.151 (0,6)
Linfomi non Hodgkin	7.011 (3,6)	6.171 (3,4)	13.182 (3,5)
Mieloma multiplo	3.019 (1,6)	2.740 (1,5)	5.759 (1,5)
Leucemie, tutte	4.738 (2,4)	3.229 (1,8)	7.967 (2,1)
<b>Totale</b>	<b>194.754 ****</b>	<b>181.857</b>	<b>376.611</b>

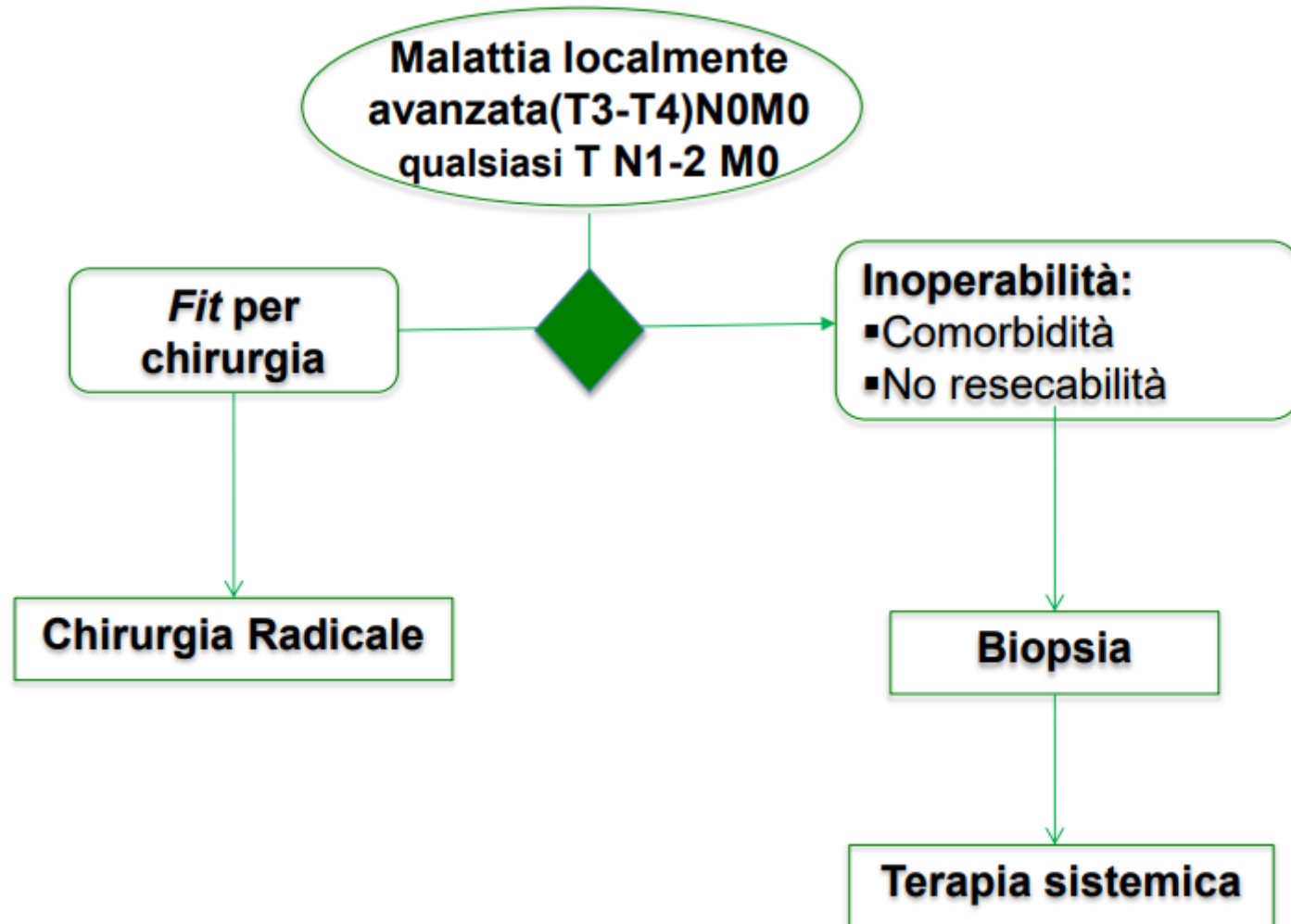
**TABELLA 6.** Numero di nuovi casi di tumore (e percentuali sul totale) stimati per il 2020 in base al sesso e per le sedi più frequenti. Sono esclusi i carcinomi della cute non melanomi

# Tumori del rene

## Trattamento chirurgico della malattia localizzata



## Trattamento chirurgico della malattia localizzata

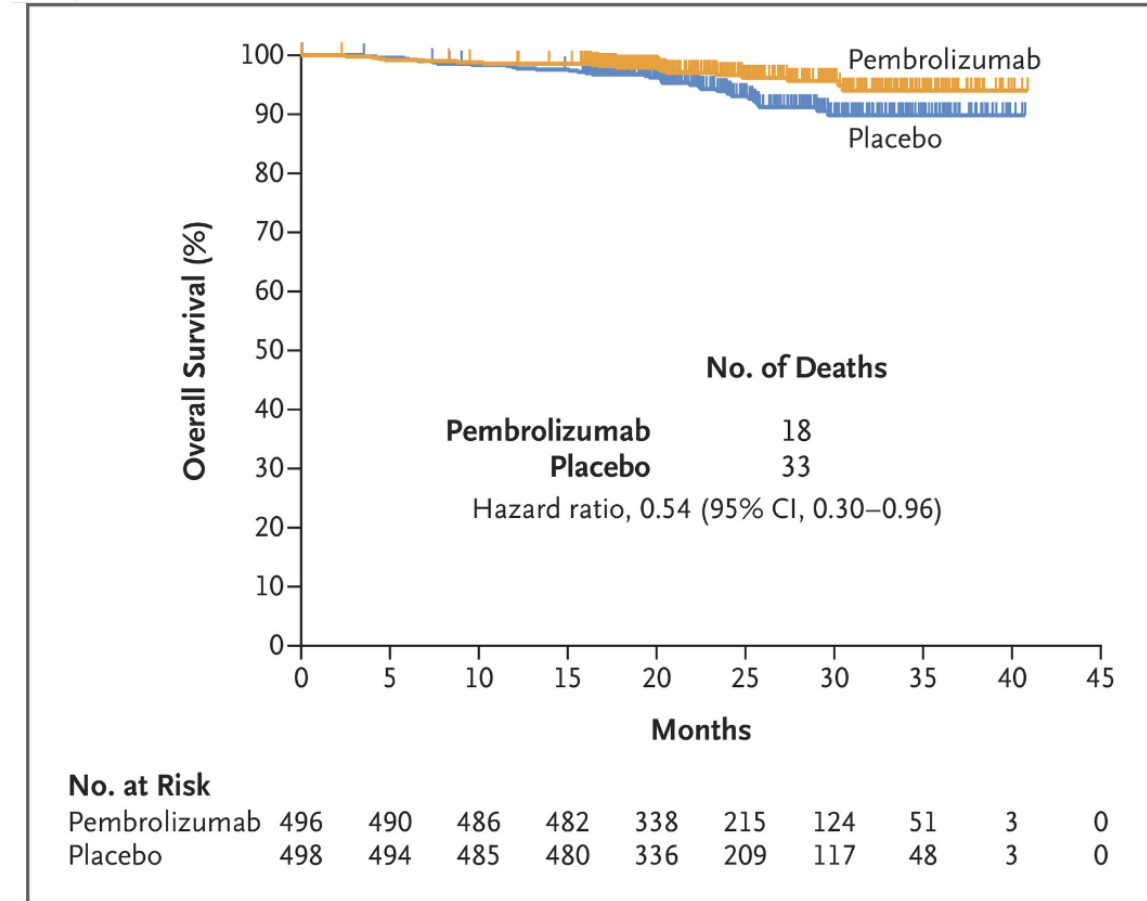
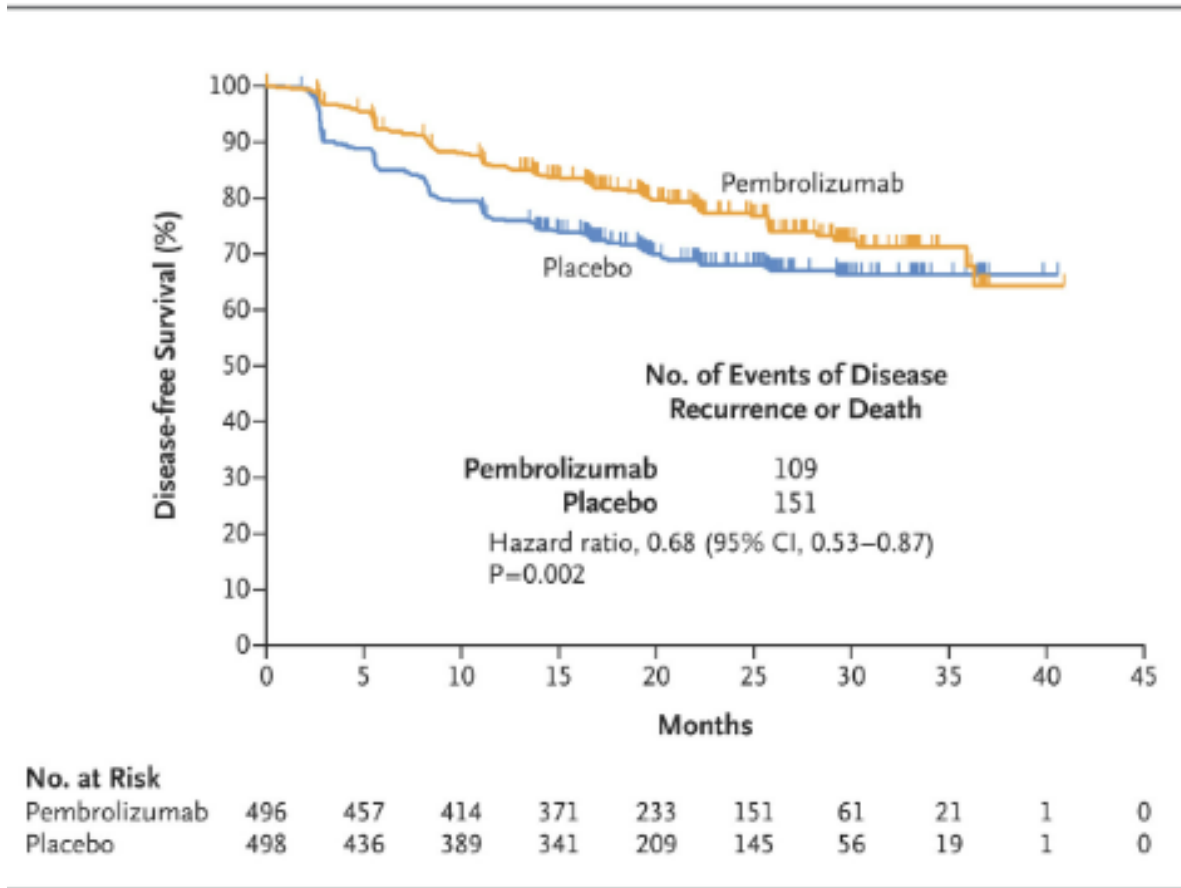


# Adjuvant Pembrolizumab (Keynote 564) in high risk pts

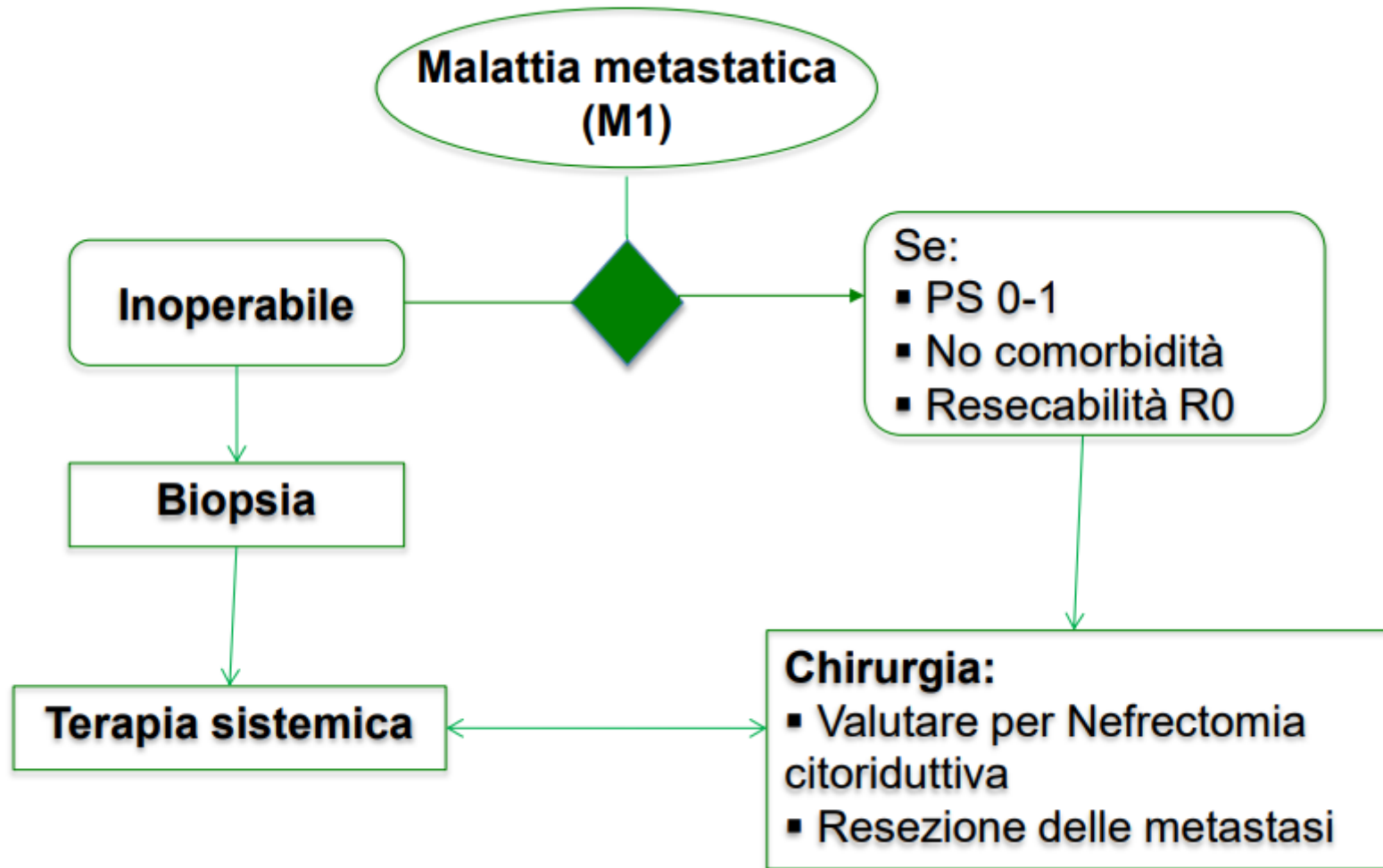
stage II G4 or sarcomatoid differentiation, stage III or higher, N+, or stage M1 with NED

**DFS**

**OS**



# Trattamento chirurgico della malattia metastatica



# Systemic treatment options for advanced/metastatic clear cell RCC

## Immunotherapies: cytokines<sup>a</sup>

IFN- $\alpha$  (Roche)<sup>1</sup>  
IL-2 (Prometheus)<sup>1</sup>

## Immunotherapies: checkpoint inhibitors

Nivolumab (Bristol Myers Squibb)<sup>2,3</sup>  
Nivolumab plus ipilimumab (Bristol Myers Squibb)<sup>4,5</sup>  
Pembrolizumab (Merck)<sup>6,7b</sup>

## Antiangiogenic therapies: mTOR inhibitors

Temsirolimus (Pfizer)<sup>1</sup>  
Everolimus (Novartis)<sup>1</sup>

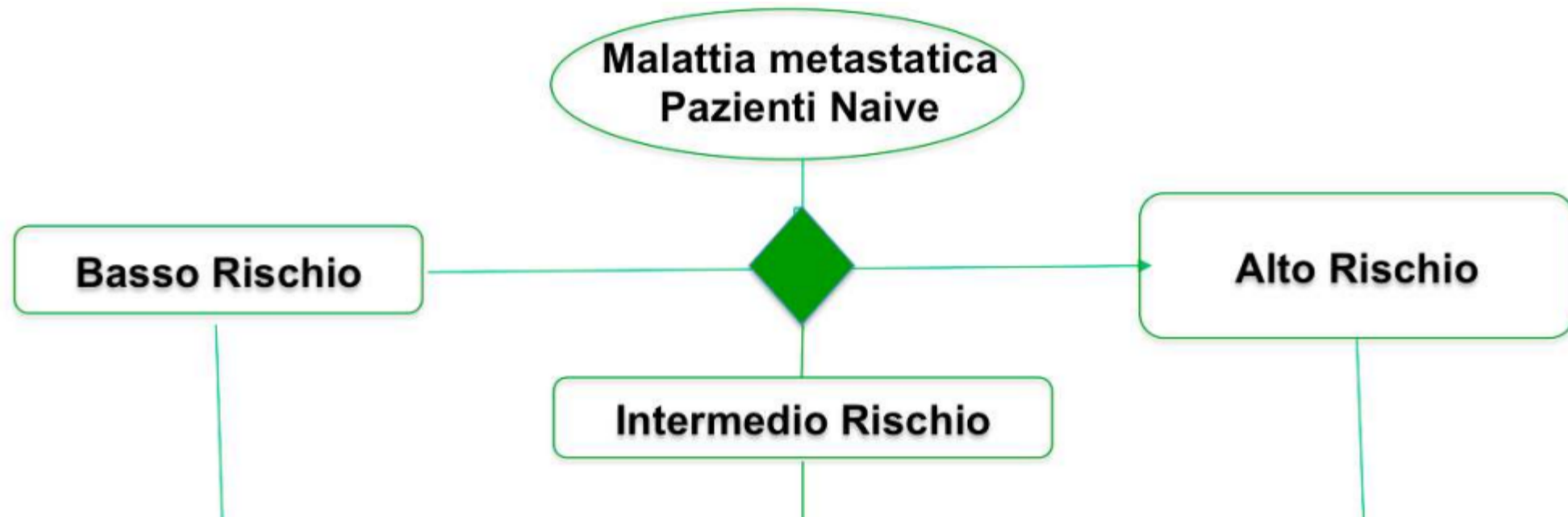
## Antiangiogenic therapies: Anti-VEGF antibodies/VEGFR inhibitors

Bevacizumab <sup>c</sup> (Roche) <sup>1</sup>	Axitinib (Pfizer) <sup>10,11</sup>
Sorafenib (Bayer Pharma AG) <sup>1</sup>	Cabozantinib (Exelixis/Ipsen) <sup>12-15</sup>
Sunitinib (Pfizer) <sup>1</sup>	Lenvatinib (Eisai) <sup>16,17,d</sup>
Pazopanib (Novartis) <sup>8,9</sup>	Tivozanib (Aveo) <sup>18,19</sup>

## Combination immunotherapies plus antiangiogenic therapy

Avelumab plus axitinib (EMD Serono, Inc; Pfizer)<sup>20,21</sup>  
Pembrolizumab plus axitinib (Merck; Pfizer)<sup>22,23</sup>  
Nivolumab plus cabozantinib (Bristol Myers Squibb; Exelixis)<sup>24,25</sup>  
Pembrolizumab plus lenvatinib (Merck; Eisai)<sup>26,27</sup>

# Trattamento medico mRCC istotipo a cellule chiare

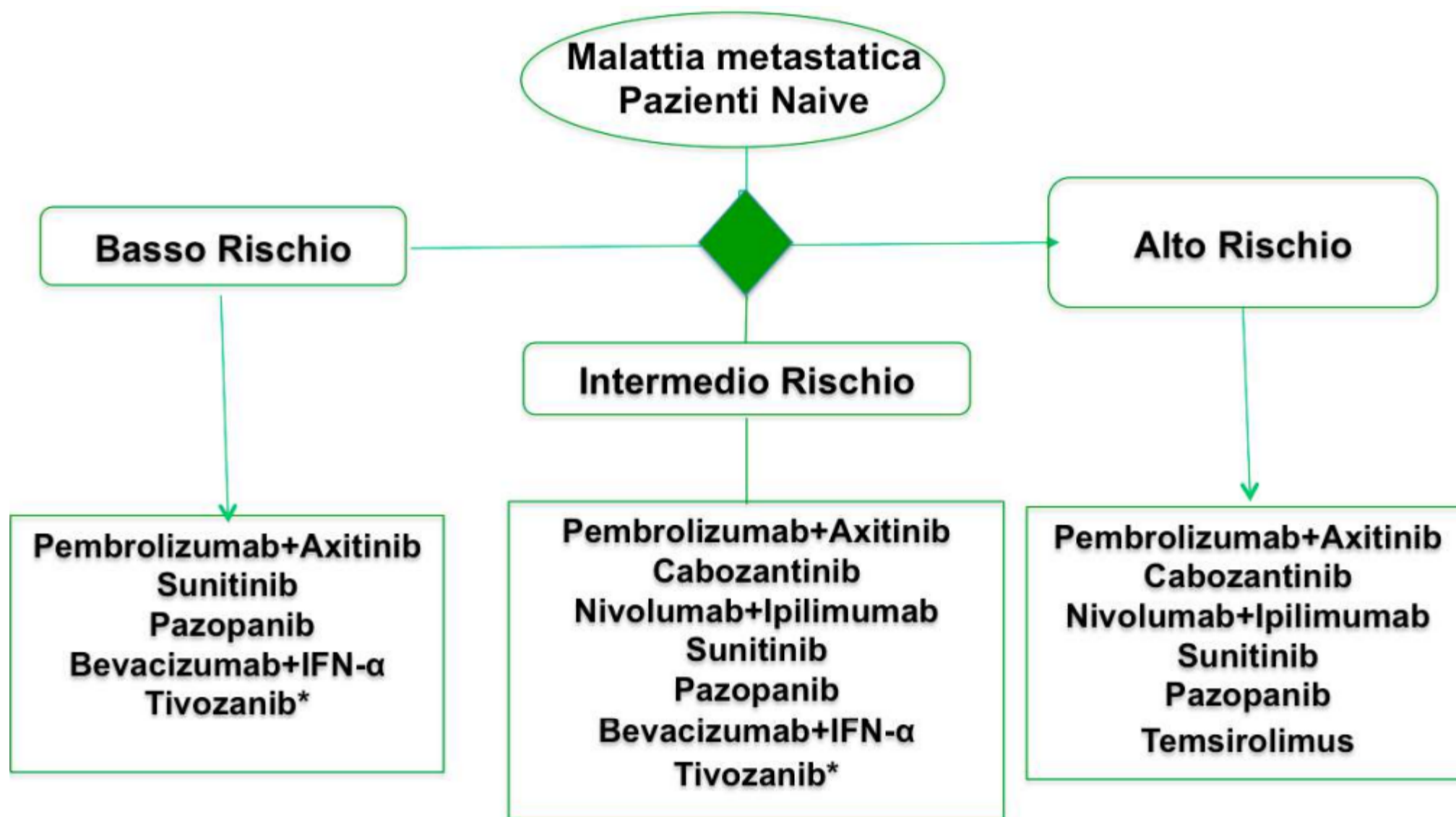


# Criteri di Heng

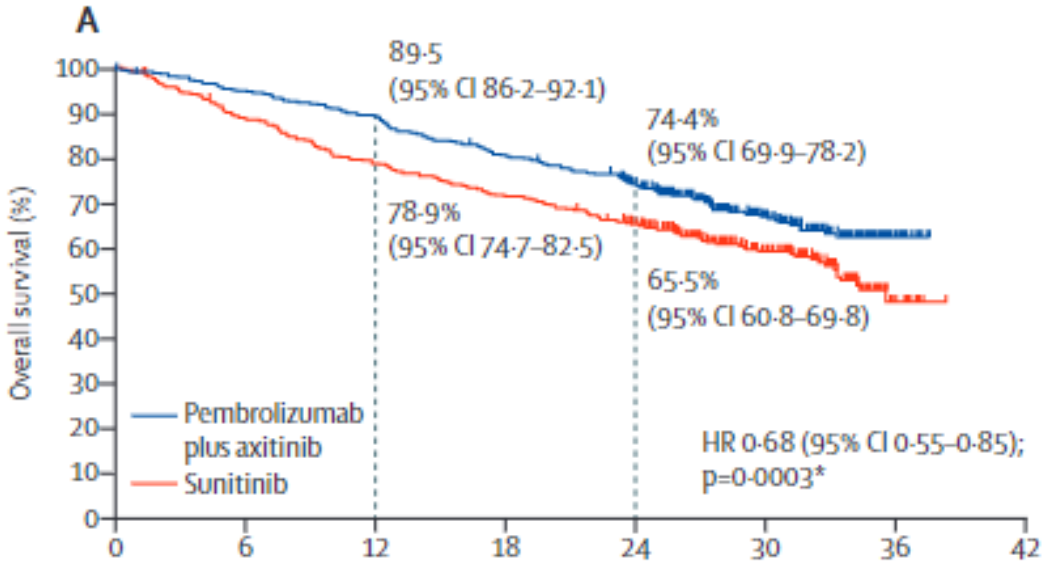
<b>Karnofsky PS</b>	< 80%
<b>Tasso di emoglobinemia</b>	< limite inferiore del <i>range</i> di normalità
<b>Calcio corretto</b>	> 10 mg/dl
<b>Periodo dalla diagnosi al trattamento</b>	< 1 anno
<b>Conta assoluta dei neutrofilii</b>	> limite superiore del <i>range</i> di normalità
<b>Conta piastrinica</b>	> limite superiore del <i>range</i> di normalità

<b>Prognosi</b>	<b>Numero di fattori</b>	<b>Sopravvivenza mediana</b>	<b>Sopravvivenza a 2 anni</b>
Favorevole	0	NR	75%
Intermedia	1-2	27 mesi	53%
Sfavorevole	3-6	8.8 mesi	7%

# Trattamento medico mRCC istotipo a cellule chiare

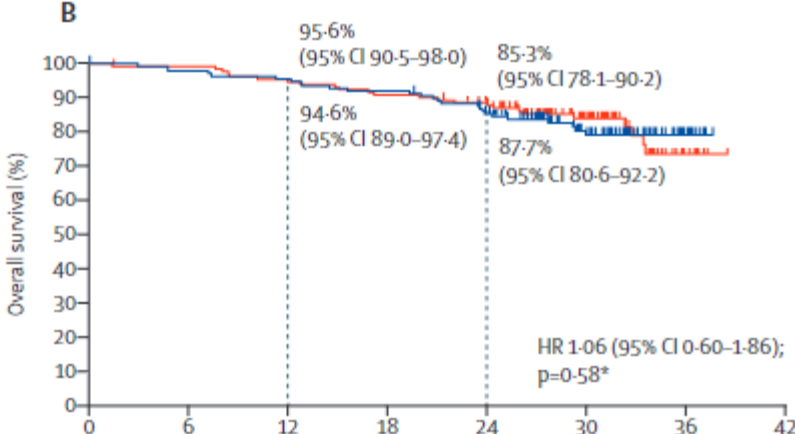


# Keynote 426: Pembrolizumab + Axitinib vs Sunitinib



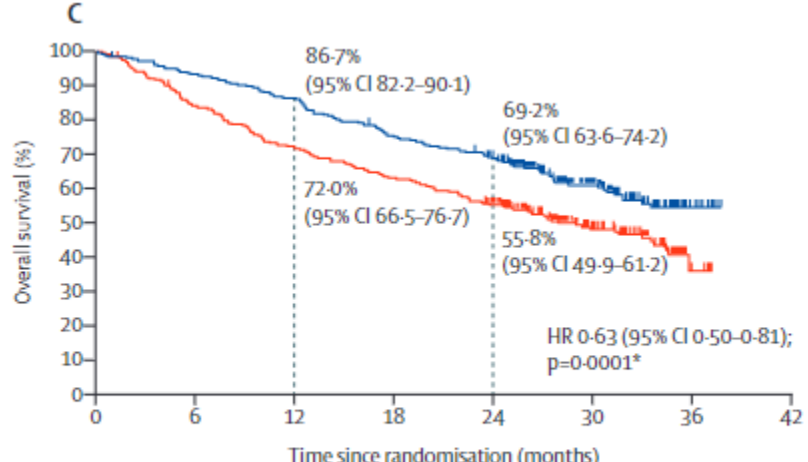
**Number at risk (number censored)**

Pembrolizumab plus axitinib	432 (0)	408 (2)	385 (2)	346 (3)	305 (17)	163 (135)	23 (267)	0 (290)
Sunitinib	429 (0)	379 (3)	336 (3)	306 (3)	268 (14)	134 (129)	16 (235)	0 (251)



**Number at risk (number censored)**

Pembrolizumab plus axitinib	138 (0)	134 (1)	131 (1)	126 (1)	110 (8)	63 (49)	12 (100)	0 (112)
Sunitinib	131 (0)	129 (1)	123 (1)	118 (1)	108 (7)	60 (51)	9 (98)	0 (107)



**Number at risk (number censored)**

Pembrolizumab plus axitinib	294 (0)	274 (1)	254 (1)	220 (2)	195 (9)	100 (86)	11 (167)	0 (178)
Sunitinib	298 (0)	250 (2)	213 (2)	188 (2)	160 (7)	74 (78)	7 (137)	0 (144)

# CheckMate 9ER

Phase 3, randomized, open-label trial of nivolumab combined with cabozantinib versus sunitinib in patients with previously untreated advanced or metastatic RCC<sup>1-5</sup>

## OS, PFS, and DOR in the ITT population

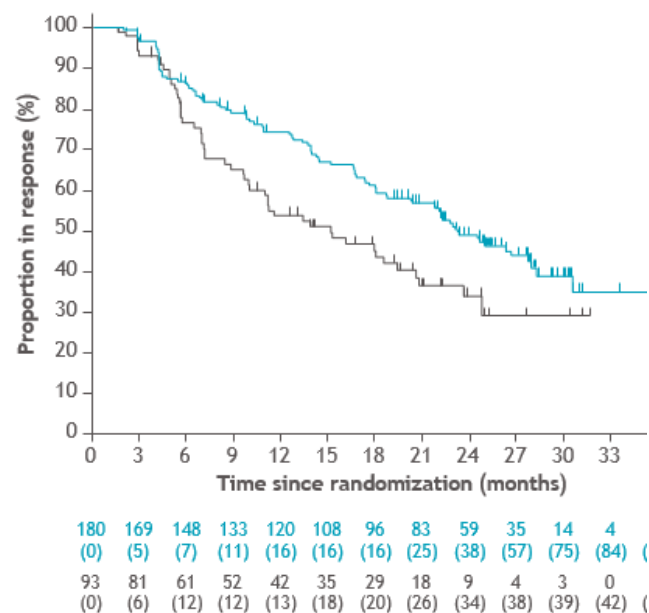
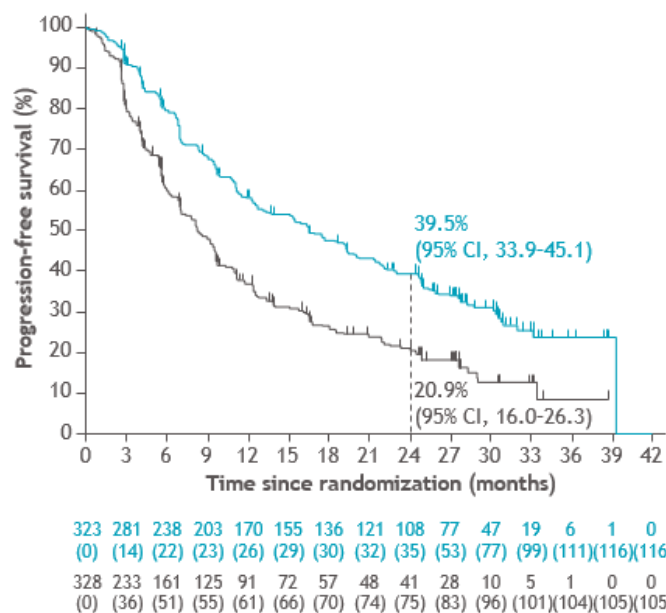
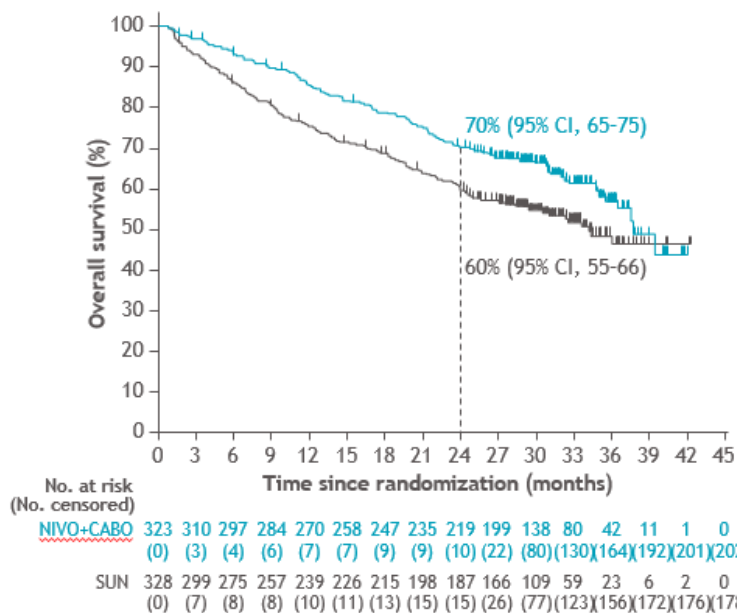
Median OS (95% CI), months <sup>a,b</sup>	
NIVO+CABO	SUN
37.7 (35.5-NE)	34.3 (29.0-NE)

HR (95% CI), 0.70 (0.55-0.90);  $P = 0.0043^d$

Median PFS (95% CI), months <sup>c</sup>	
NIVO+CABO	SUN
16.6 (12.8-19.8)	8.3 (7.0-9.7)

HR (95% CI), 0.56 (0.46-0.68);  $P < 0.0001^d$

Median DOR (95% CI), months	
NIVO+CABO	SUN
23.1 (20.2-27.9)	15.1 (9.9-20.5)



- Median follow-up for survival (IOR) was 32.9 (30.4-35.9) months; minimum 25.4 months

Vertical lines denote censored patients.

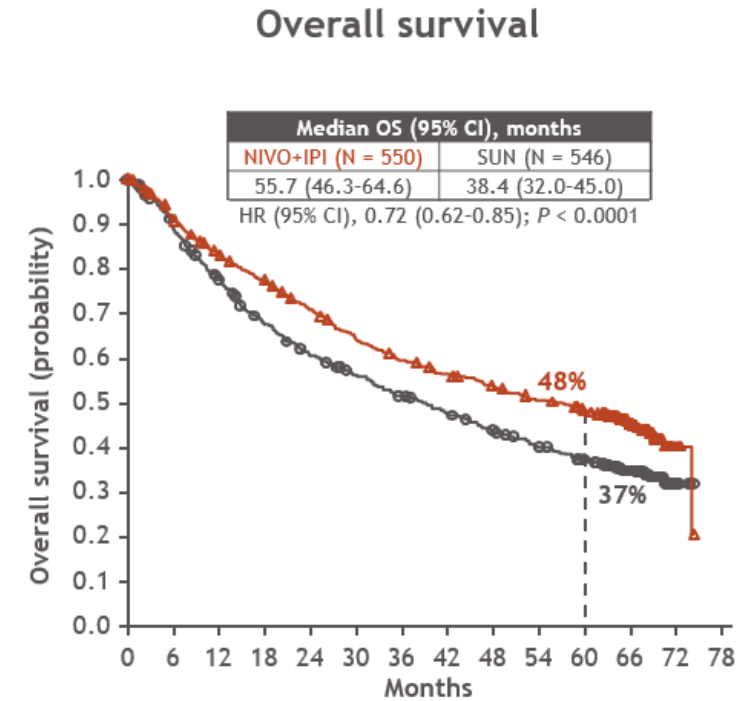
<sup>a</sup>Preplanned final analysis of OS according to the protocol set to occur after 254 events. <sup>b</sup>271 events occurred (121 events with NIVO+CABO and 150 events with SUN). <sup>c</sup>207 (64%) of 323 patients had an event with NIVO+CABO; 223 (68%) of 328 patients had an event with SUN. <sup>d</sup> $P$  values are descriptive.

Motzer RJ et al. *Lancet Oncol* 2022;23:888-898.

# CheckMate 214 study design

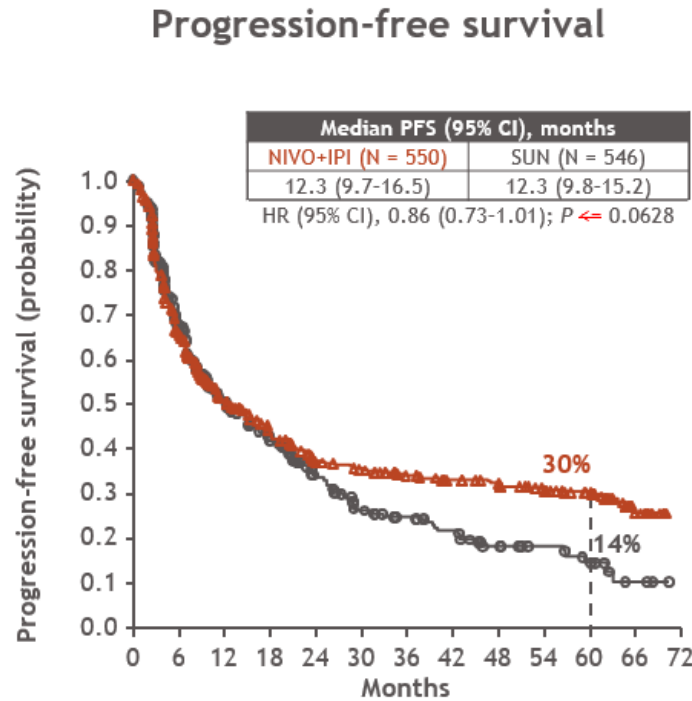
Phase 3, randomized, open-label trial of nivolumab combined with ipilimumab vs sunitinib monotherapy in treatment-naïve patients with advanced or metastatic clear cell RCC<sup>1</sup>

## OS, PFS, and DOR in ITT patients



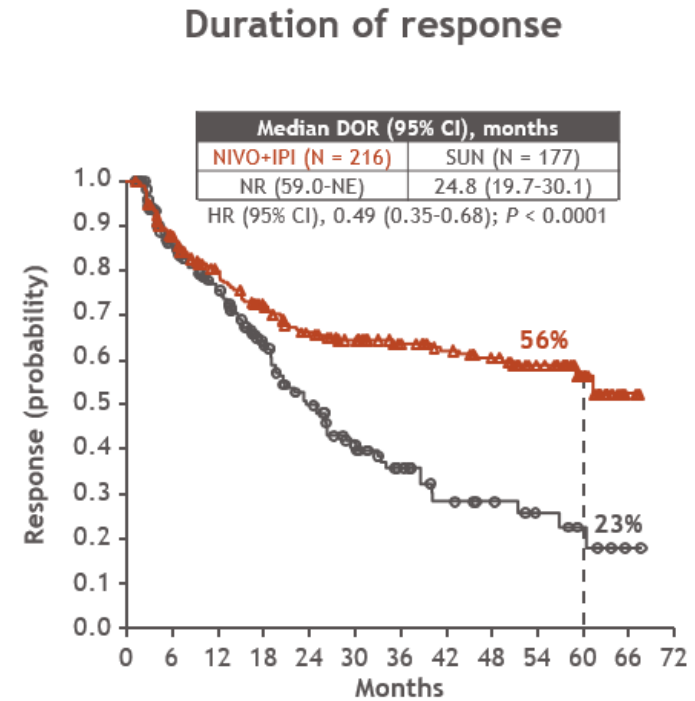
No. at risk

	0	6	12	18	24	30	36	42	48	54	60	66	72	78
NIVO+IPI	550	493	444	411	372	337	309	291	274	256	236	138	5	0
SUN	546	472	405	347	310	281	257	234	213	192	171	108	6	0



No. at risk

	0	6	12	18	24	30	36	42	48	54	60	66	72
NIVO+IPI	550	315	217	171	132	121	103	92	86	75	62	14	0
SUN	546	285	178	130	87	59	42	33	21	15	10	3	0



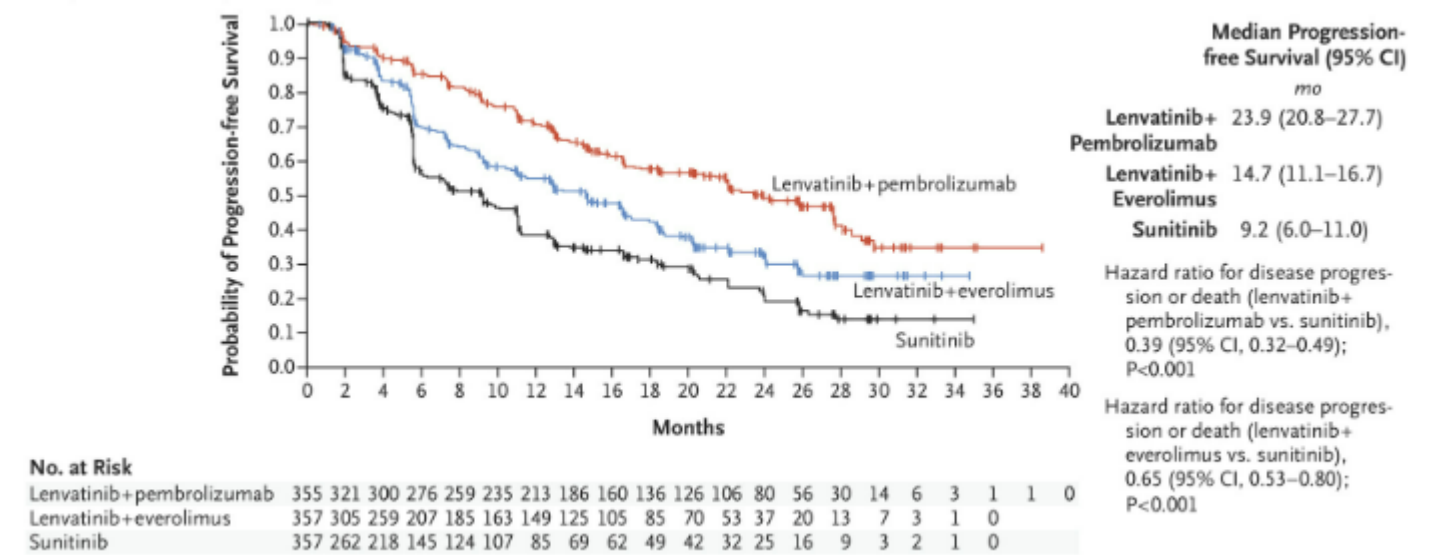
No. at risk

	0	6	12	18	24	30	36	42	48	54	60	66	72
NIVO+IPI	216	177	151	130	110	99	88	80	74	61	33	4	0
SUN	177	128	104	76	54	39	24	15	12	8	5	1	0

➤ Avelumab + intermittent axitinib

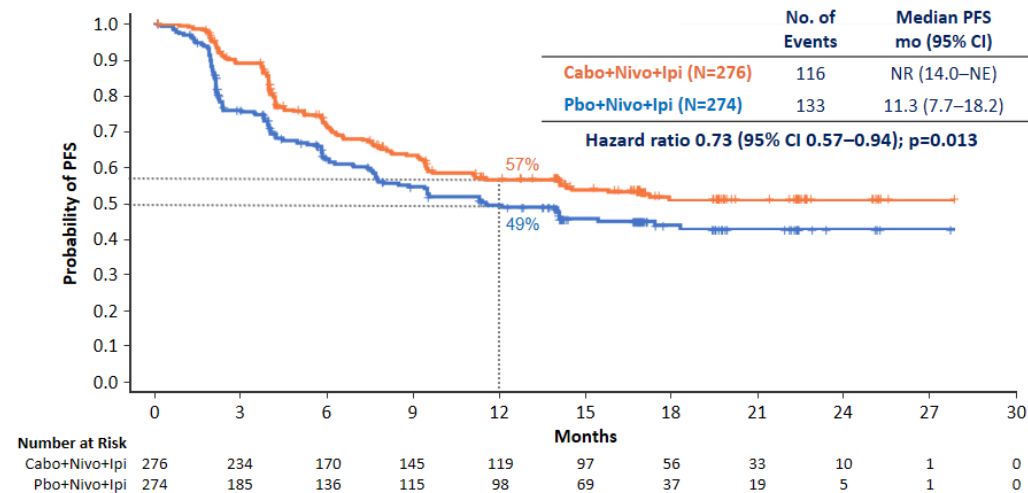
➤ Lenvatinib + pembrolizumab

A Kaplan–Meier Analysis of Progression-free Survival

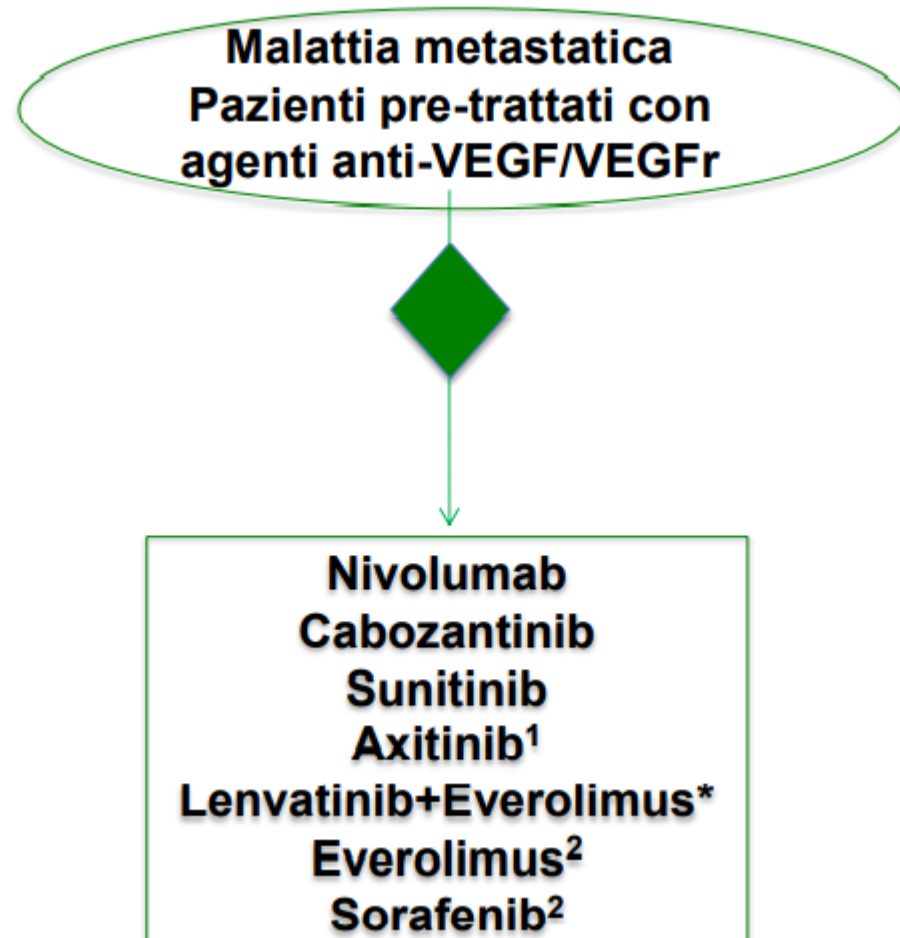


Ipilimumab + nivolumab + cabozantinib

Progression-Free Survival: Final Analysis (PITT Population)

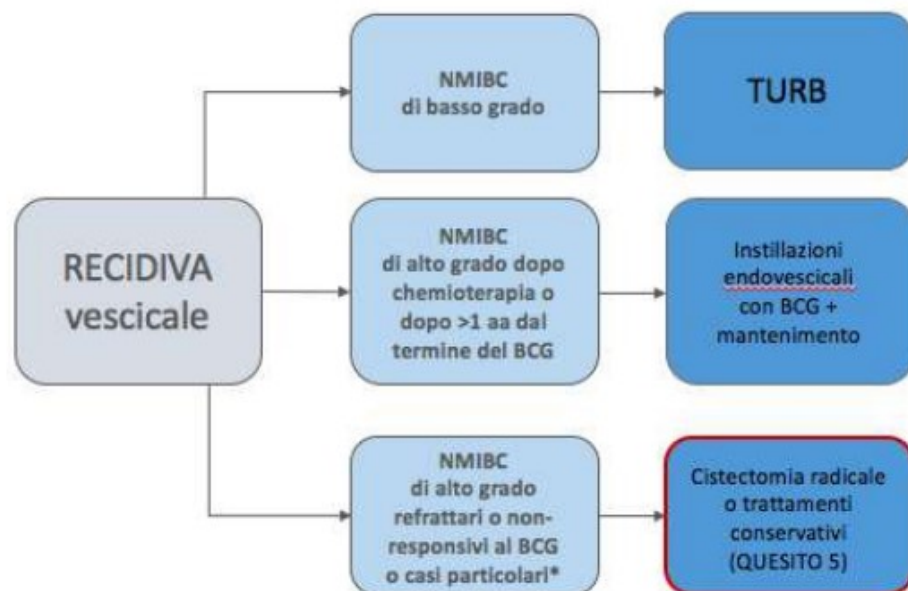
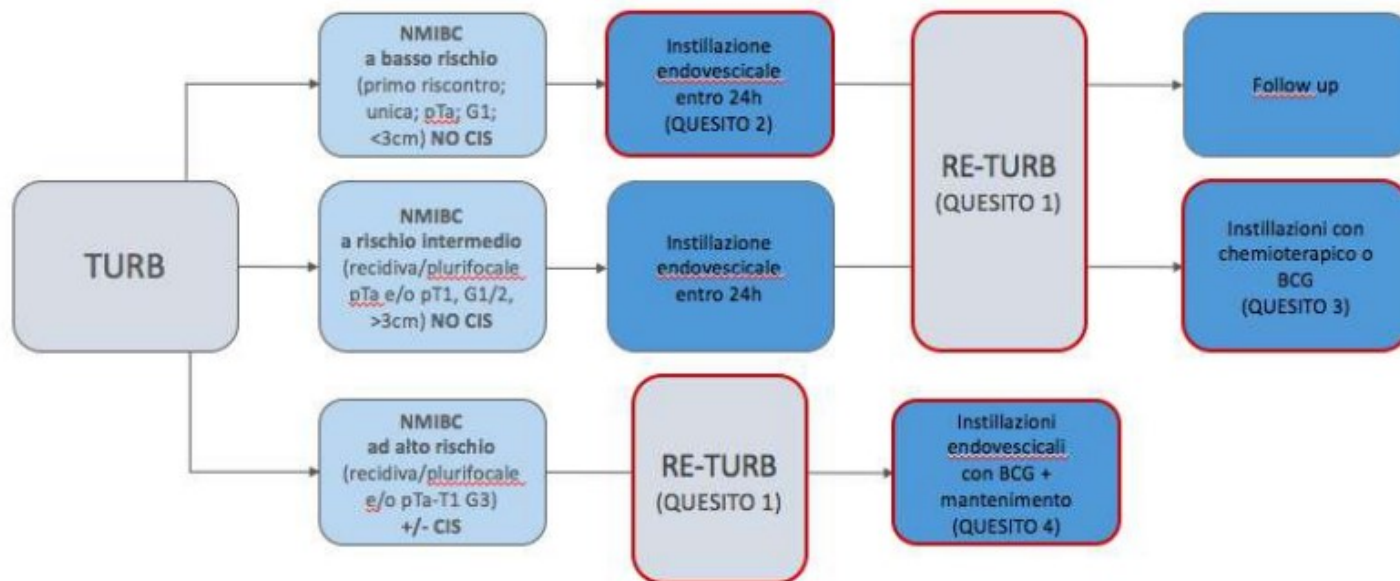


# Trattamento medico mRCC istotipo a cellule chiare

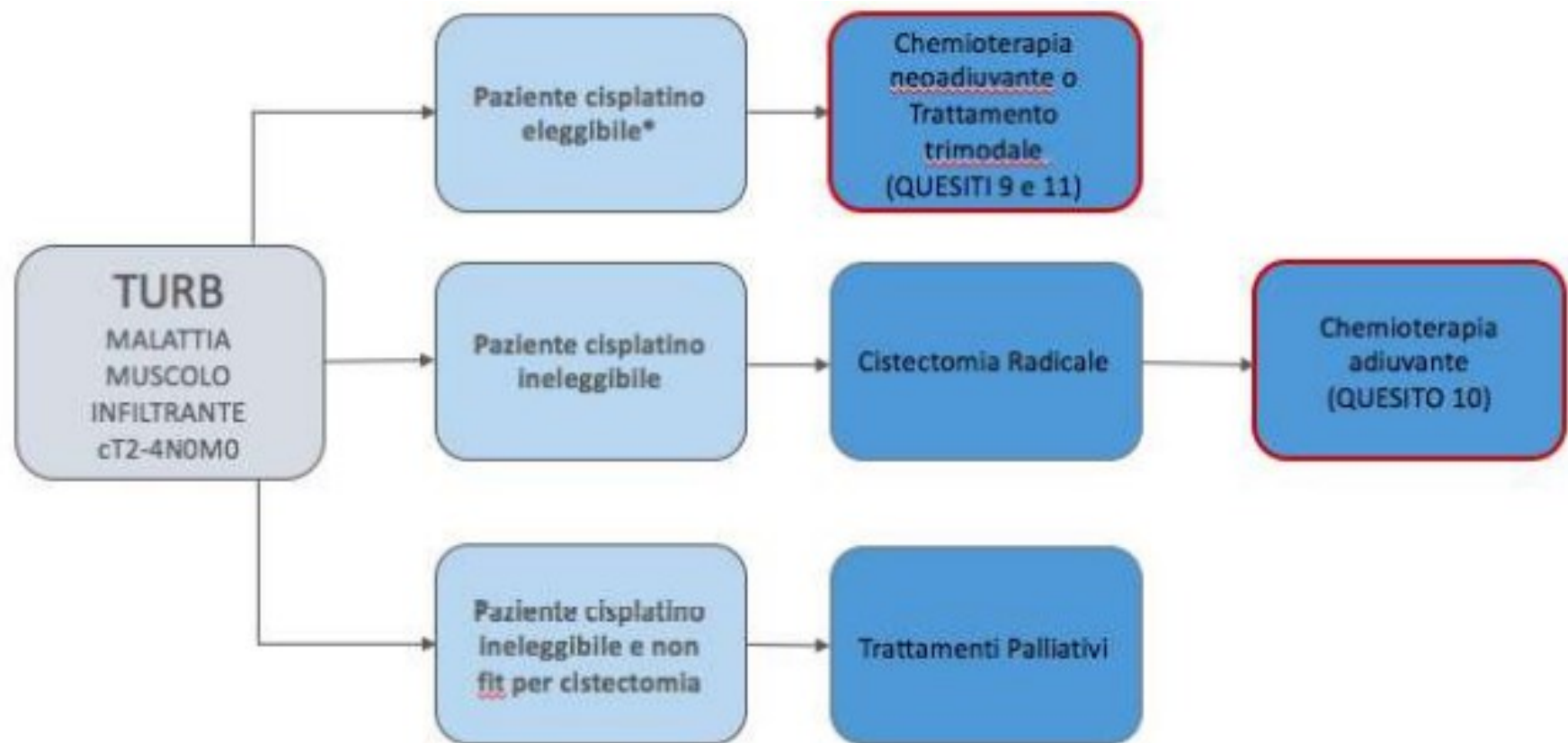


# Tumori della vescica

## Trattamento della Malattia Vescicale Non-Muscolo Infiltrante (NMIBC)

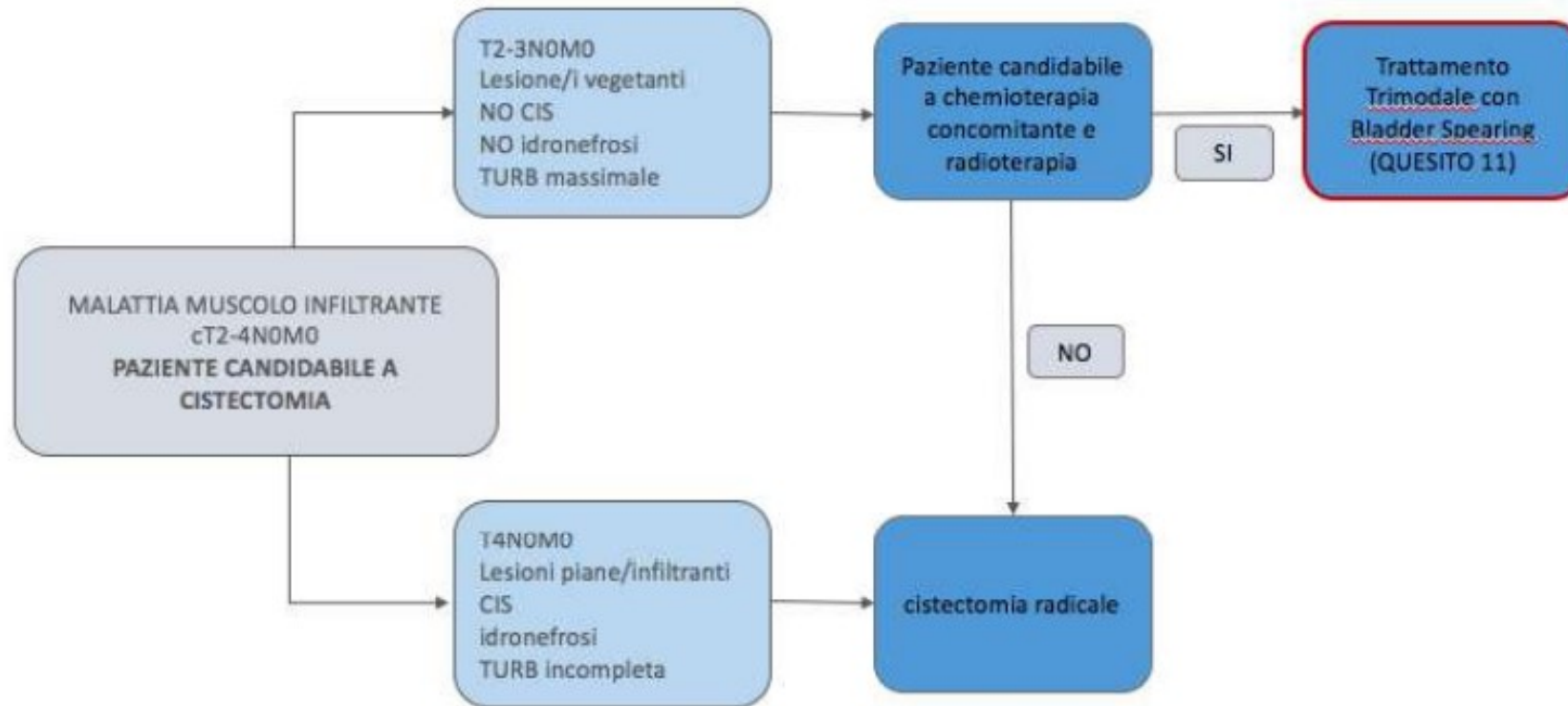


## Trattamento della malattia vescicale muscolo infiltrante (MIBC, T2-4)

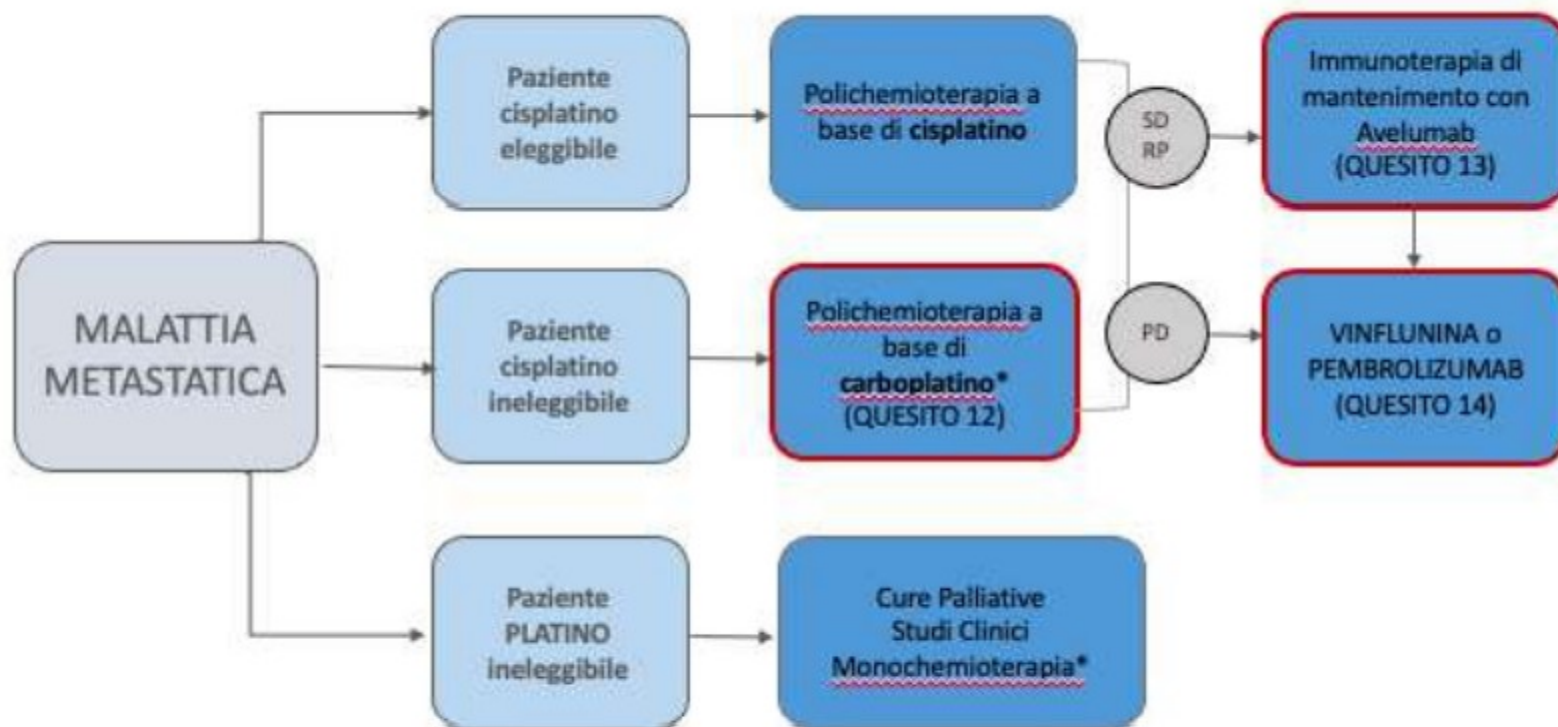


\* Cisplatino eleggibilità: ECOG PS 0-1; clearance creatinina  $>60$  ml/min; non alterazioni dell'udito; non neuropatia periferica

## Trattamento integrato di Bladder-Sparing (“Trimodale”)



## Terapia della Malattia Uroteliale Avanzata



\* L'immunoterapia con PEMBROLIZUMAB è indicata per pazienti con malattia avanzata, cisplatino ineleggibili, PD-L1 positivi, ma non rimborsato da AIFA

# Treatment of mUC: NCCN Guidelines®

## 1L systemic therapy

Cisplatin eligible	Cisplatin ineligible
<ul style="list-style-type: none"> <li>Gemcitabine and cisplatin (category 1) followed by avelumab maintenance therapy (category 1)<sup>a</sup></li> <li>ddMVAC with growth factor support (category 1) followed by avelumab maintenance therapy (category 1)<sup>a</sup></li> </ul>	<p><b>Platinum eligible:</b></p> <ul style="list-style-type: none"> <li>Gemcitabine and carboplatin followed by avelumab maintenance therapy (category 1)<sup>a</sup></li> <li>Atezolizumab<sup>a,b</sup></li> </ul> <p><b>Platinum ineligible:</b></p> <ul style="list-style-type: none"> <li>Atezolizumab<sup>a,b</sup></li> <li>Pembrolizumab<sup>a</sup></li> <li>Gemcitabine</li> <li>Gemcitabine and paclitaxel</li> <li>Ifosfamide, doxorubicin, and gemcitabine<sup>c</sup></li> </ul>

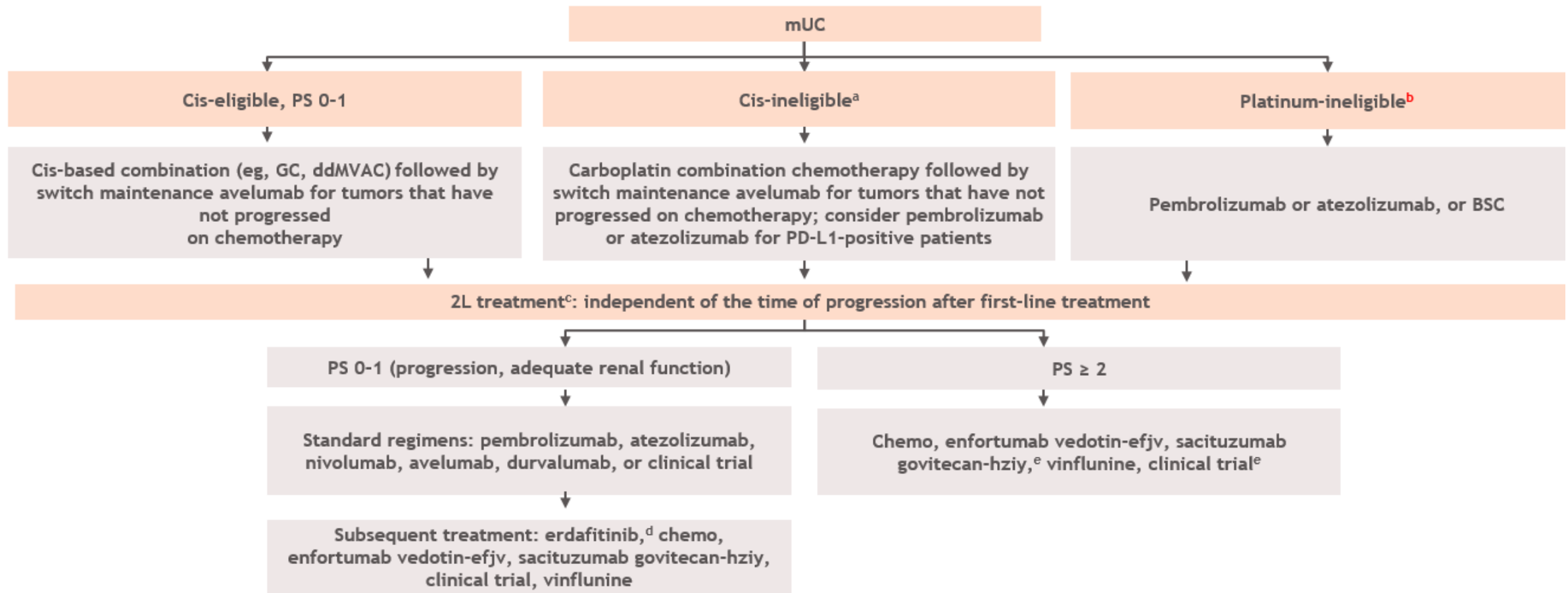
## 2L systemic therapy (post-platinum or other chemotherapy)

<ul style="list-style-type: none"> <li>Pembrolizumab (category 1 post-platinum)<sup>a</sup></li> <li>Nivolumab<sup>a</sup></li> <li>Avelumab<sup>a</sup></li> <li>Erdafitinib<sup>a,d</sup></li> <li>Enfortumab vedotin-ejfv<sup>a,e</sup></li> </ul>	<ul style="list-style-type: none"> <li>Paclitaxel or docetaxel</li> <li>Gemcitabine</li> <li>Ifosfamide, doxorubicin, and gemcitabine</li> <li>Gemcitabine and paclitaxel</li> <li>Gemcitabine and cisplatin</li> <li>ddMVAC with growth factor support</li> </ul>
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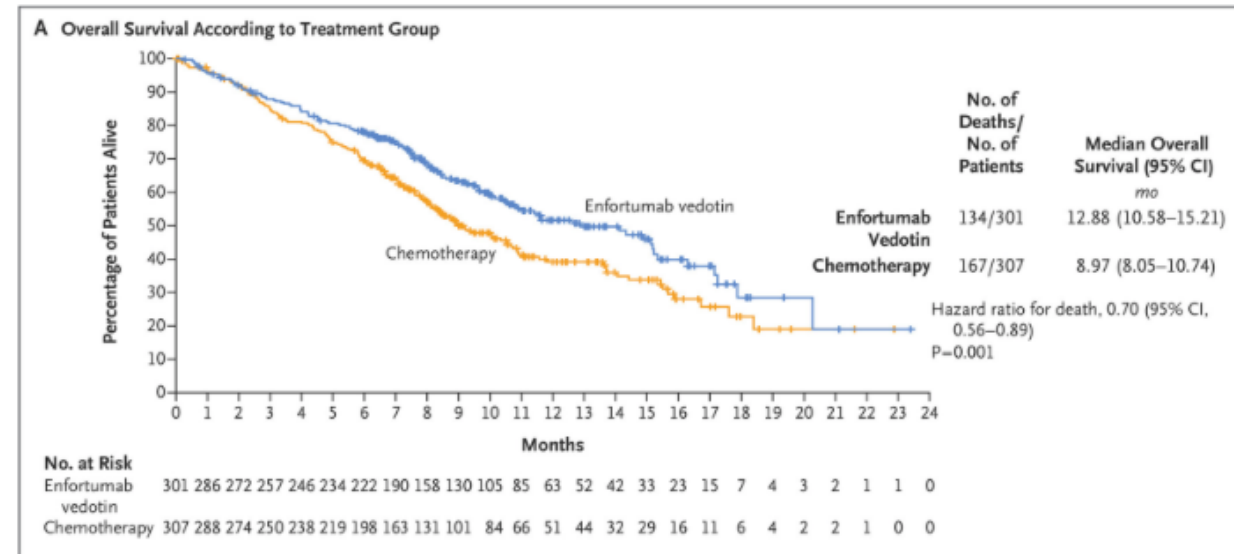
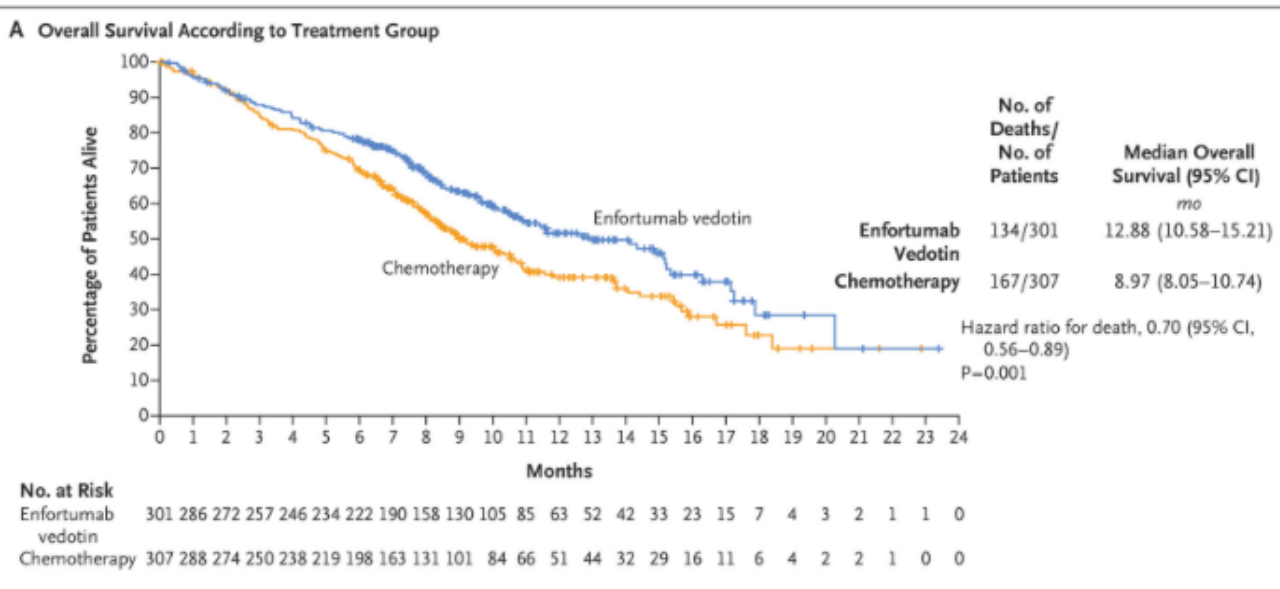
## 2L systemic therapy (post-checkpoint inhibitor)

Cisplatin eligible	Cisplatin ineligible <sup>f</sup>
<ul style="list-style-type: none"> <li>Gemcitabine and cisplatin<sup>a</sup></li> <li>ddMVAC with growth factor support<sup>a</sup></li> </ul>	<ul style="list-style-type: none"> <li>Enfortumab vedotin-ejfv<sup>a</sup></li> <li>Gemcitabine/carboplatin<sup>a</sup></li> </ul>
<ul style="list-style-type: none"> <li>Erdafitinib<sup>d,g</sup></li> <li>Paclitaxel or docetaxel<sup>g</sup></li> <li>Gemcitabine<sup>g</sup></li> </ul>	<ul style="list-style-type: none"> <li>Ifosfamide, doxorubicin, and gemcitabine<sup>h</sup></li> <li>Gemcitabine and paclitaxel<sup>h</sup></li> </ul>

# Treatment of metastatic urothelial carcinoma: EAU, ESMO guidelines<sup>1,2</sup>

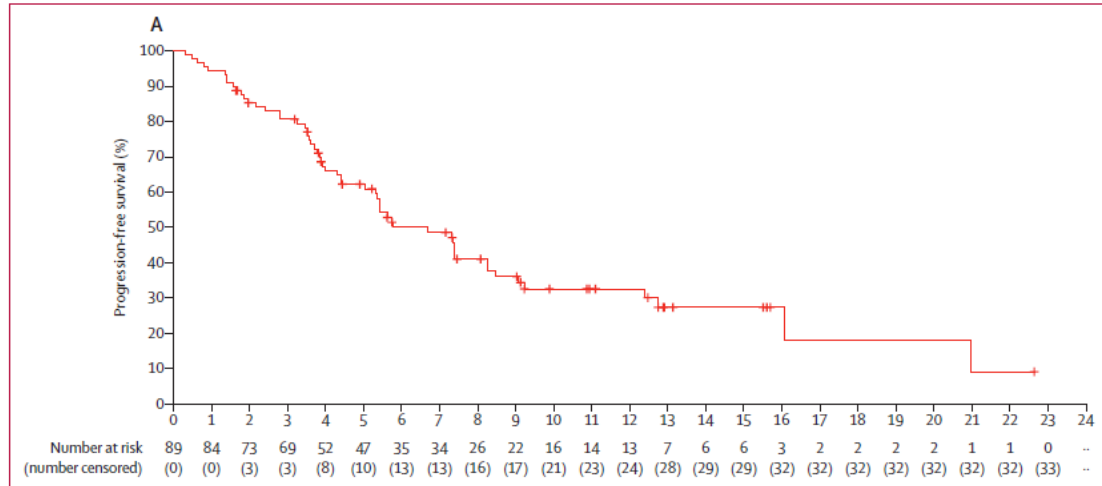


# Enfortumab Vedotin in Previously Treated Advanced Urothelial Carcinoma

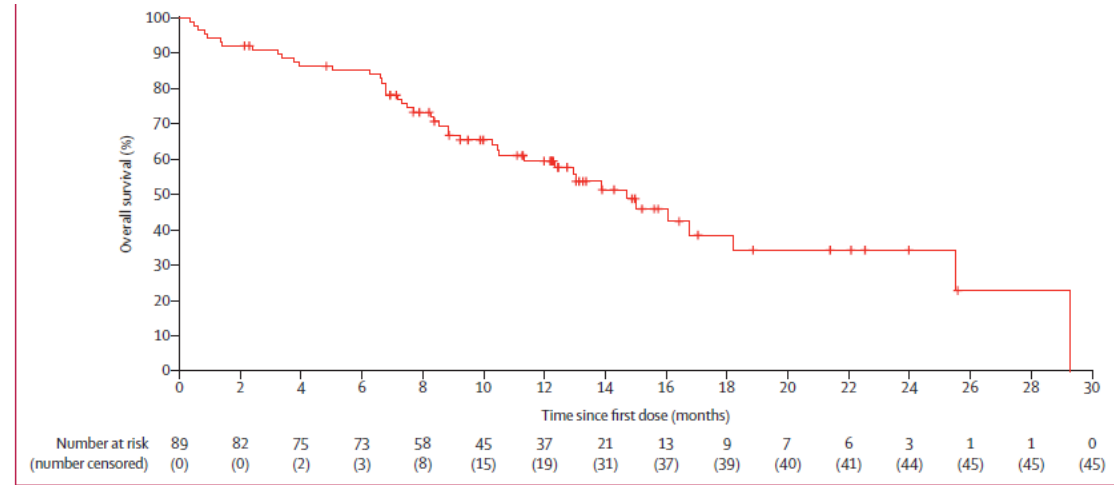


# Enfortumab vedotin after PD-1 or PD-L1 inhibitors in cisplatin-ineligible patients with advanced urothelial carcinoma (EV-201): a multicentre, single-arm, phase 2 trial

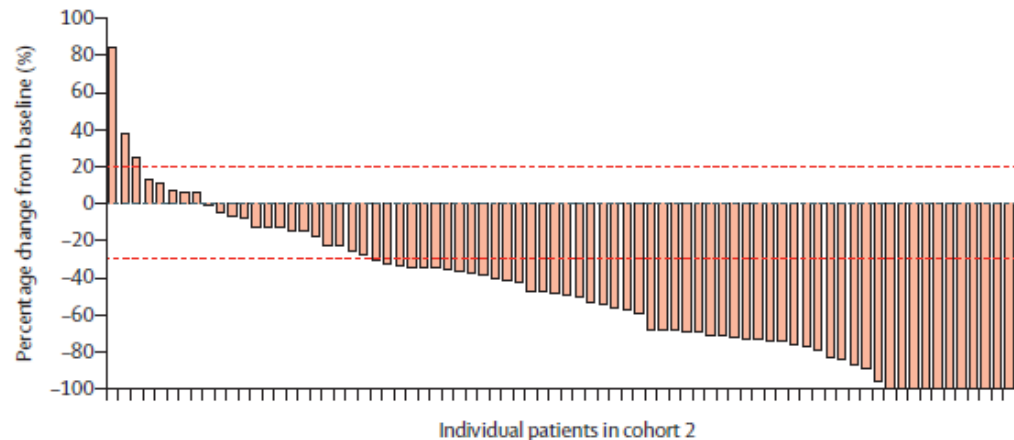
(median FUP: 13.4 months)



The median progression-free survival was 5.8 months (95% CI 5.03–8.28)

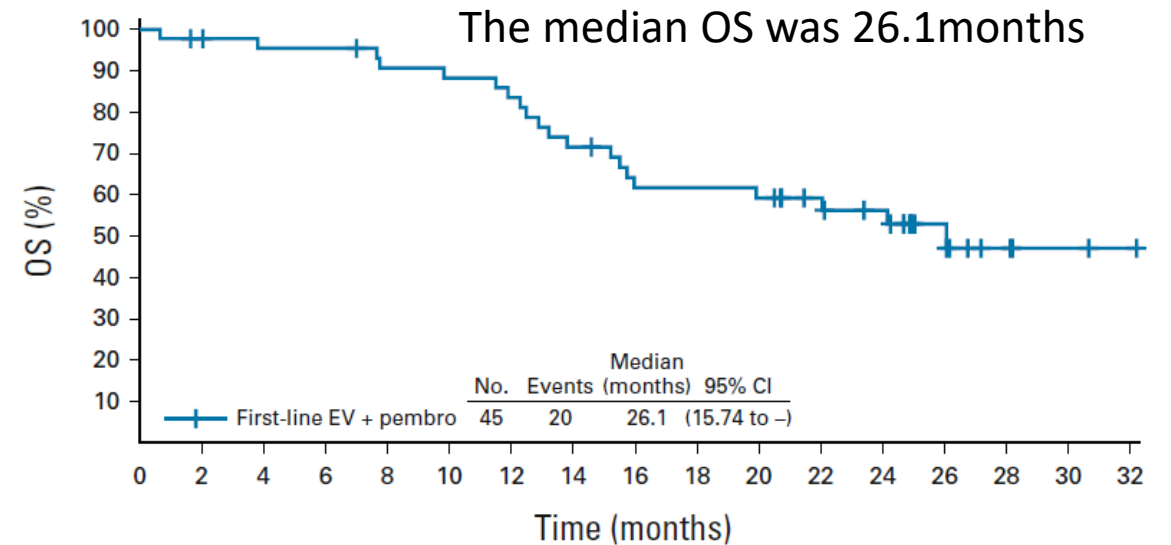
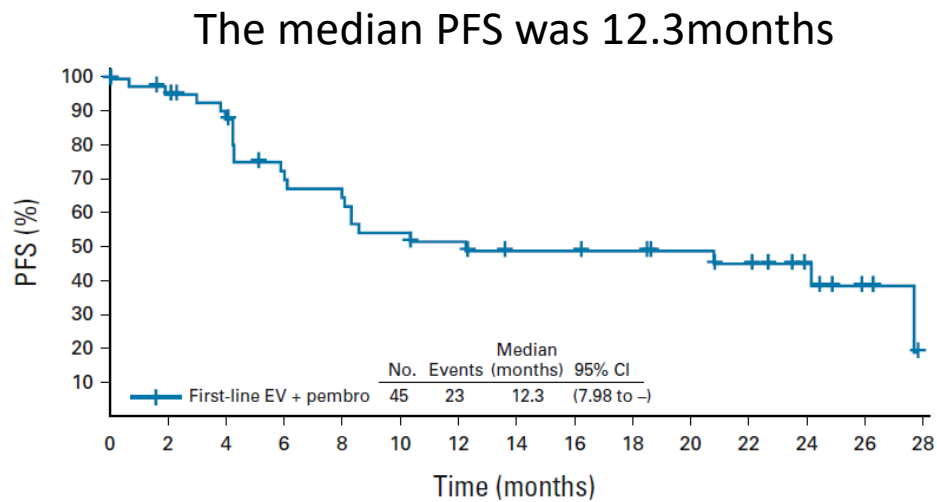
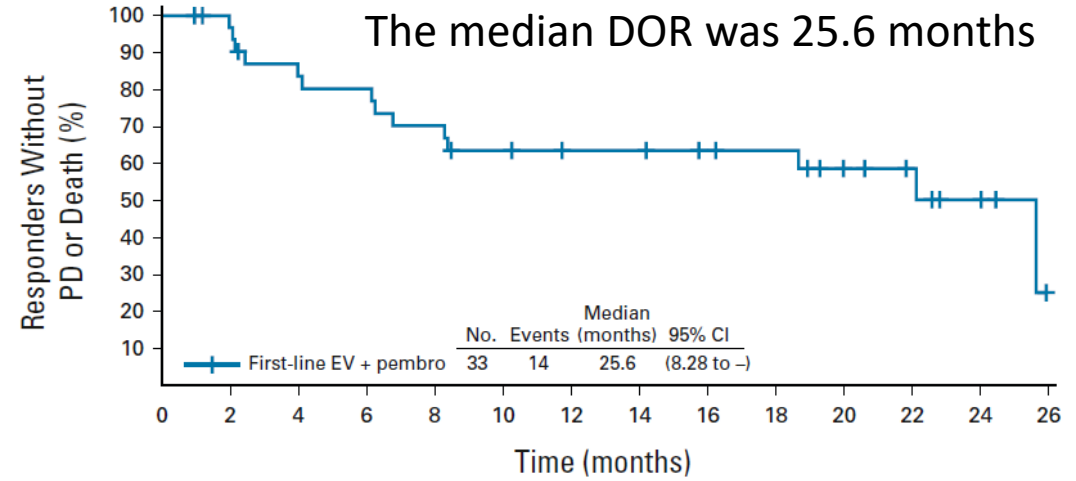
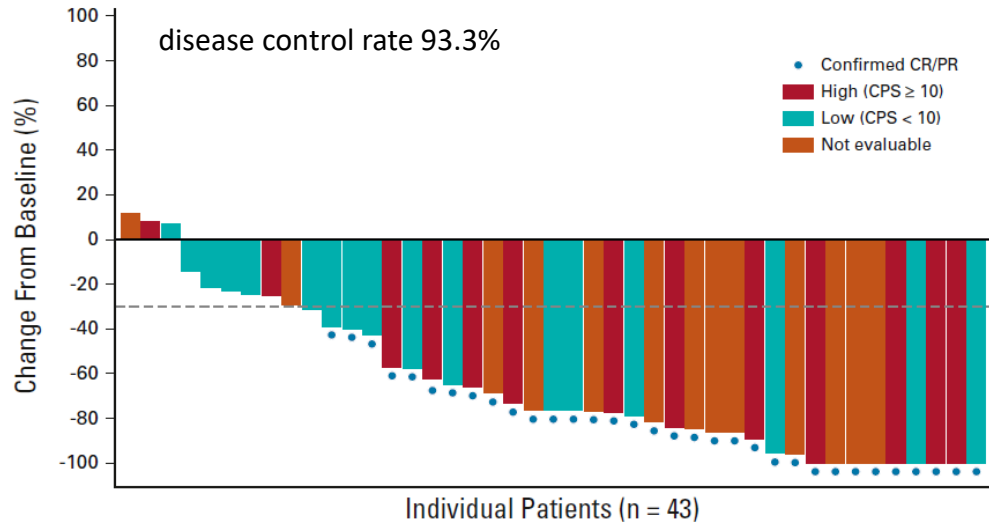


The median overall survival was 14.7 months (95% CI 10.51–18.20; figure 3B).



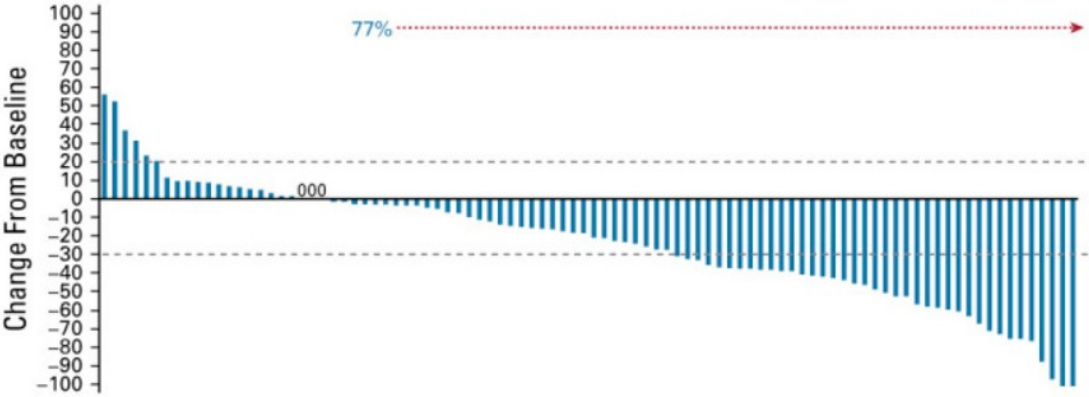
objective response rate: 52%

# Enfortumab Vedotin Plus Pembrolizumab in Previously Untreated Advanced Urothelial Cancer

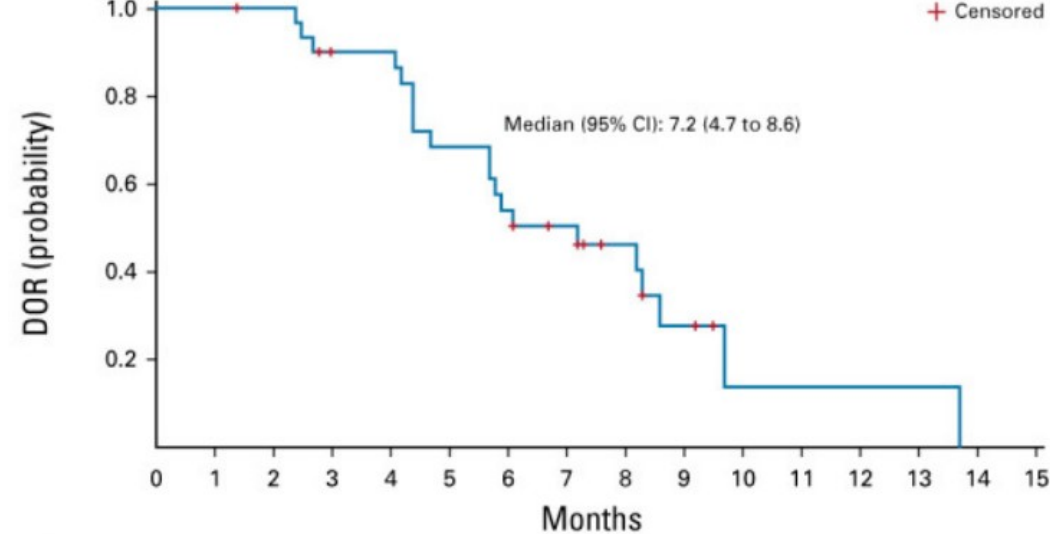
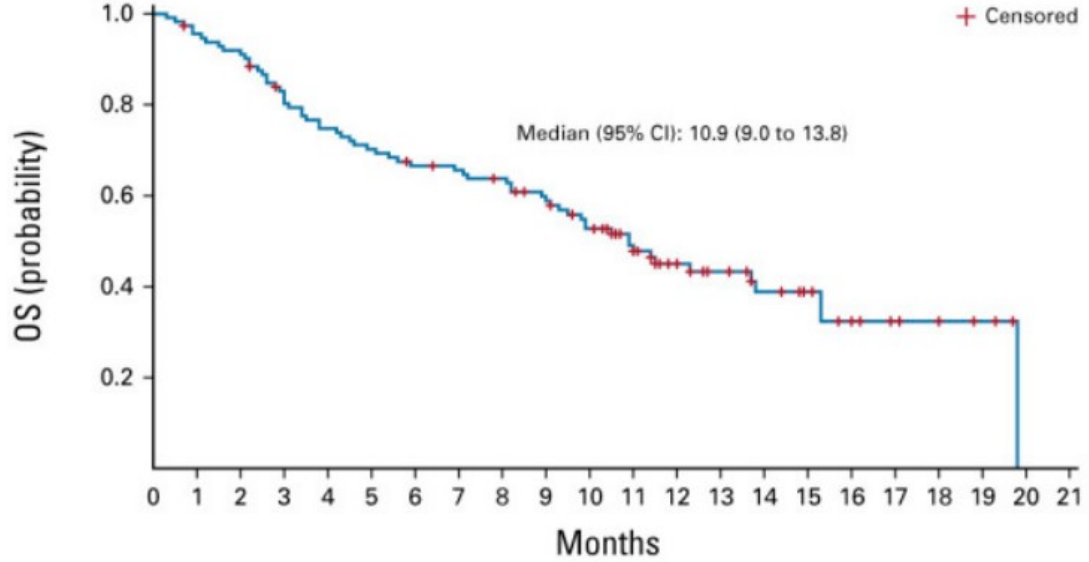
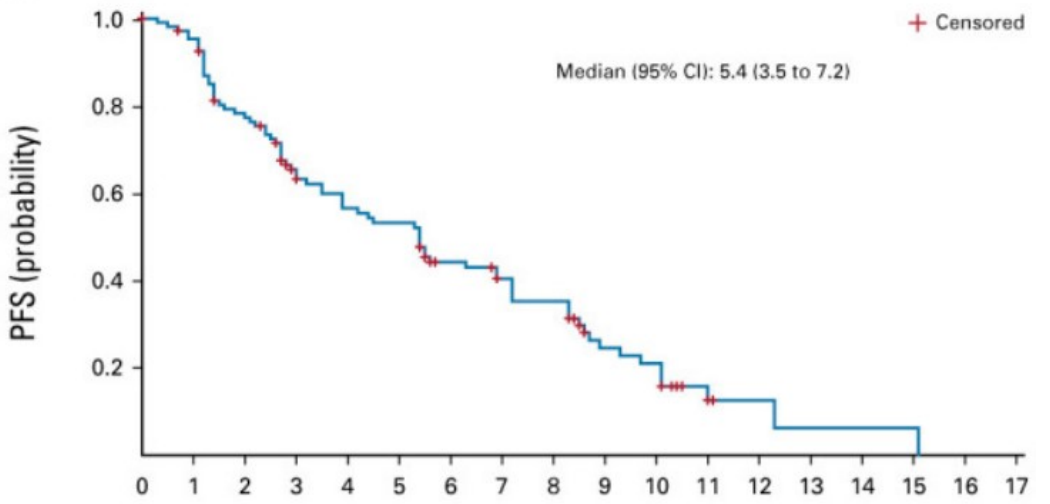


# TROPHY-U-01: A Phase II Open-Label Study of Sacituzumab Govitecan in Patients With Metastatic Urothelial Carcinoma Progressing After Platinum-Based Chemotherapy and Checkpoint Inhibitors

**A**



median DOR:7.2 months



# Gruppo multidisciplinare (incontro settimanale)

- Urologo
  - Discussione dei nuovi casi di neoplasia
- Oncologo
  - Discussione dei casi che richiedano un cambio di terapia
- Radioterapista
  - Revisione della letteratura
- Anatomopatologo
- Radiologo