

« En Vrac » RETROSPECTIVE ASCO 1991-2021 : Exemple du cancer du rein



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AVEC 30 ans d'écart ...



1 génération



ASCO 1991

▪ **Sein :**

- Tout début Histoire HER2 : facteur pronostique +++
- On parlait à peine des Taxanes et des anti-aromatases

▪ **Colon :**

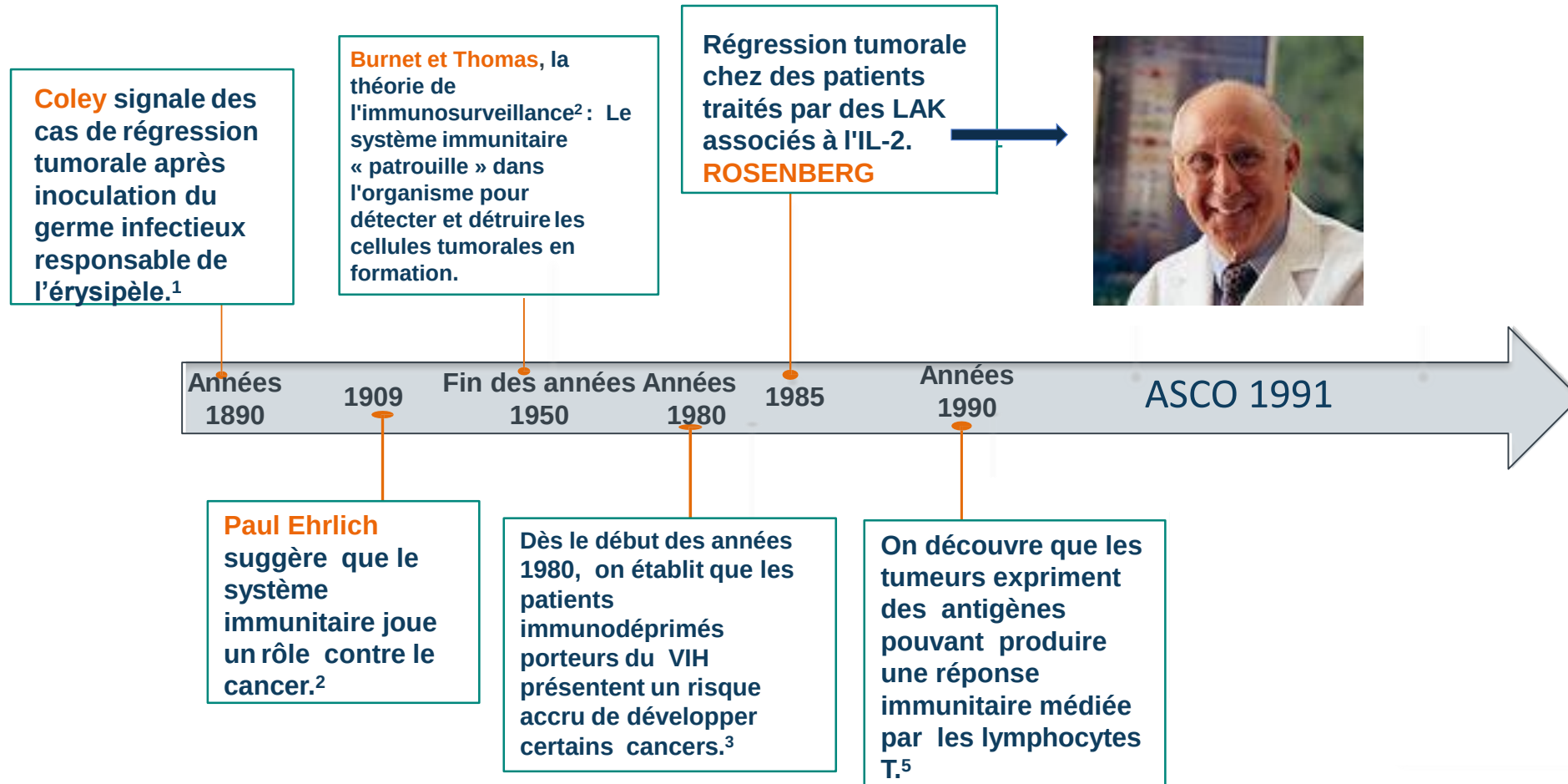
- Supériorité schéma 5FU continu : pas d'OXALIPLATINE, CAMPTO, anti-EGFR, Béva et encore moins d'immuno ...

▪ **Poumon :** ère des chimio « modernes » :

- Doublets Platine + Navelbine, Gemcitabine ...

▪ **Prostate :** agonistes seul ...

Immunothérapie et cancer avant 1990



ANNEES 1990... REIN

- **Début traitement par IL 2 IV (schéma West 5 j): TOXICITE + + +**
 - **Velbé/IFN puis IFN alfa seul ss/cut**
 - **Association IFN et IL2 ss/cut**
 - **1ers essais de traitement anti-angiogénique...**
 - **Région Nouvelle-Aquitaine :**
Pr Alain RAVAUD (Bordeaux) et Pr Jean-Marc TOURANI (Poitiers)
- **Résultats : 5% de patients en RC à long terme (métas pulmonaires)**

ASCO 2008

mRCC 1ère ligne : actualisation de l'étude de Motzer

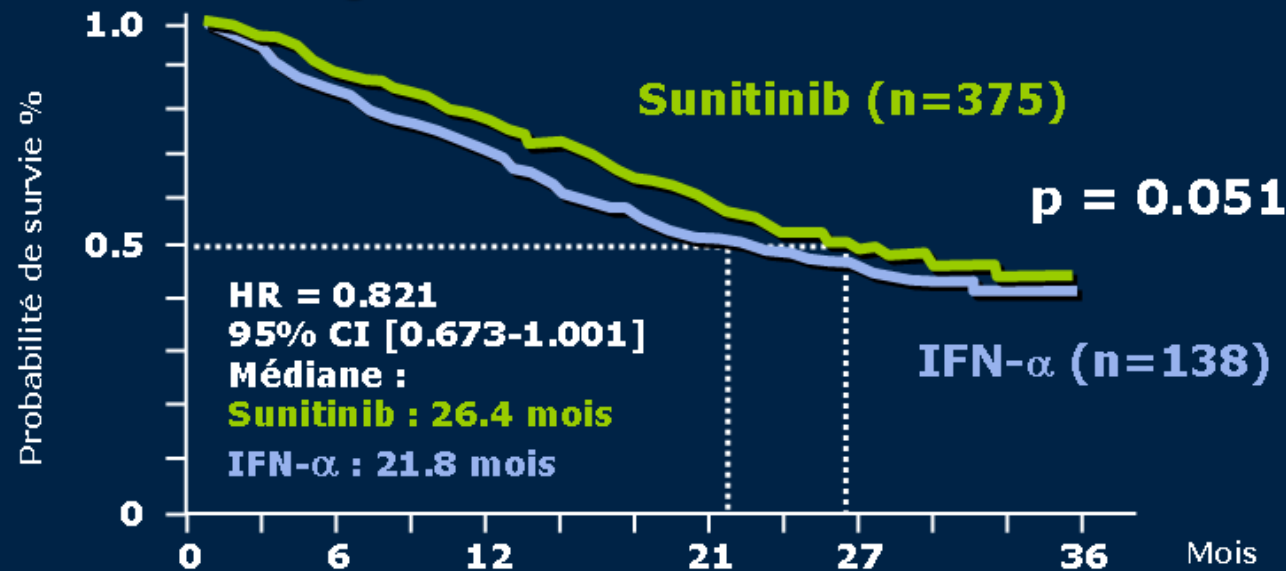
R

sunitinib (50 mg/j 4 semaines sur 6)

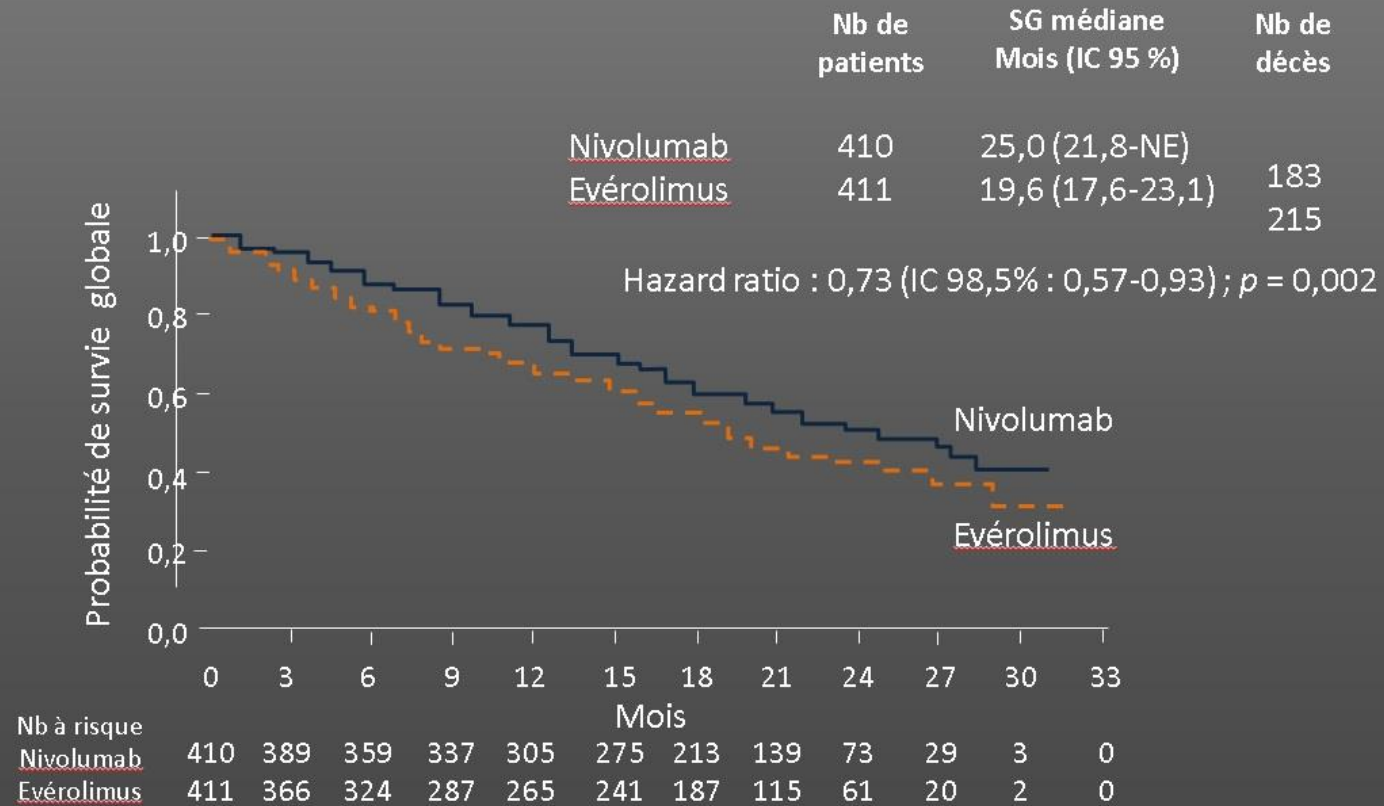
interféron- α par voie SC 3 fois par semaine

(doses progressives de 3 MU à 9 MUI)

► Survie globale par traitement

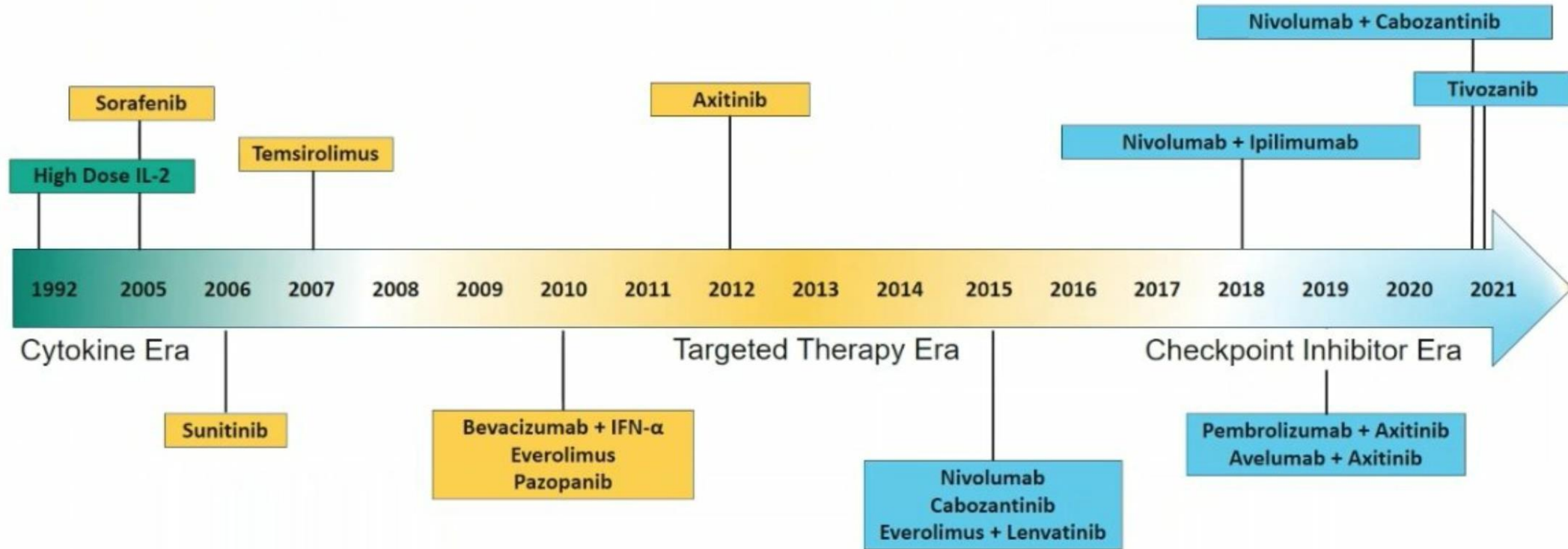


ESMO 2015 : Bénéfice en survie globale de l'immunothérapie Nivolumab versus Everolimus



D'après la communication orale de Sharma P, et al. *Abst LBA3, ECC 2015*.

Apparition d'options thérapeutiques en situation métastatique



Toxicités décroissantes, efficacité croissante
Depuis 2015 : Associations immunothérapies

Question actuelle : Usage en traitement adjuvant?

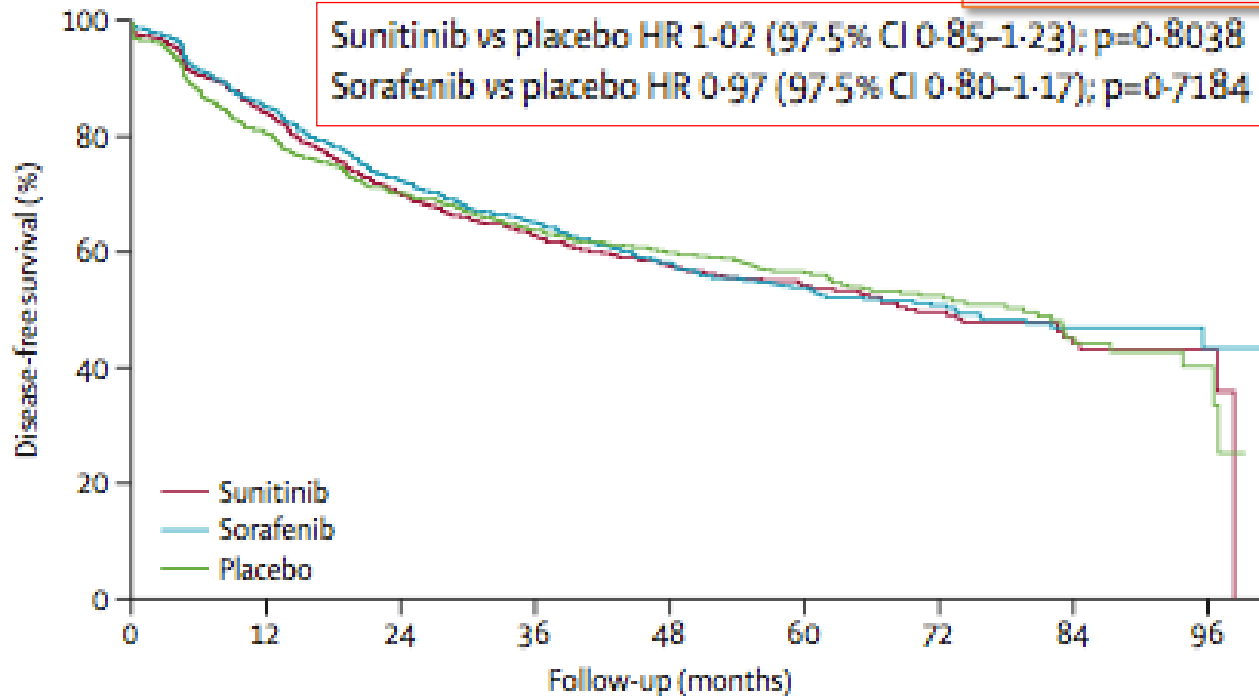
Balance pour décider d'un traitement adjuvant



- Disease-free survival
- Overall survival
- Risk of over treatment
- Side effect of therapy
- Quality of life
- Financial cost

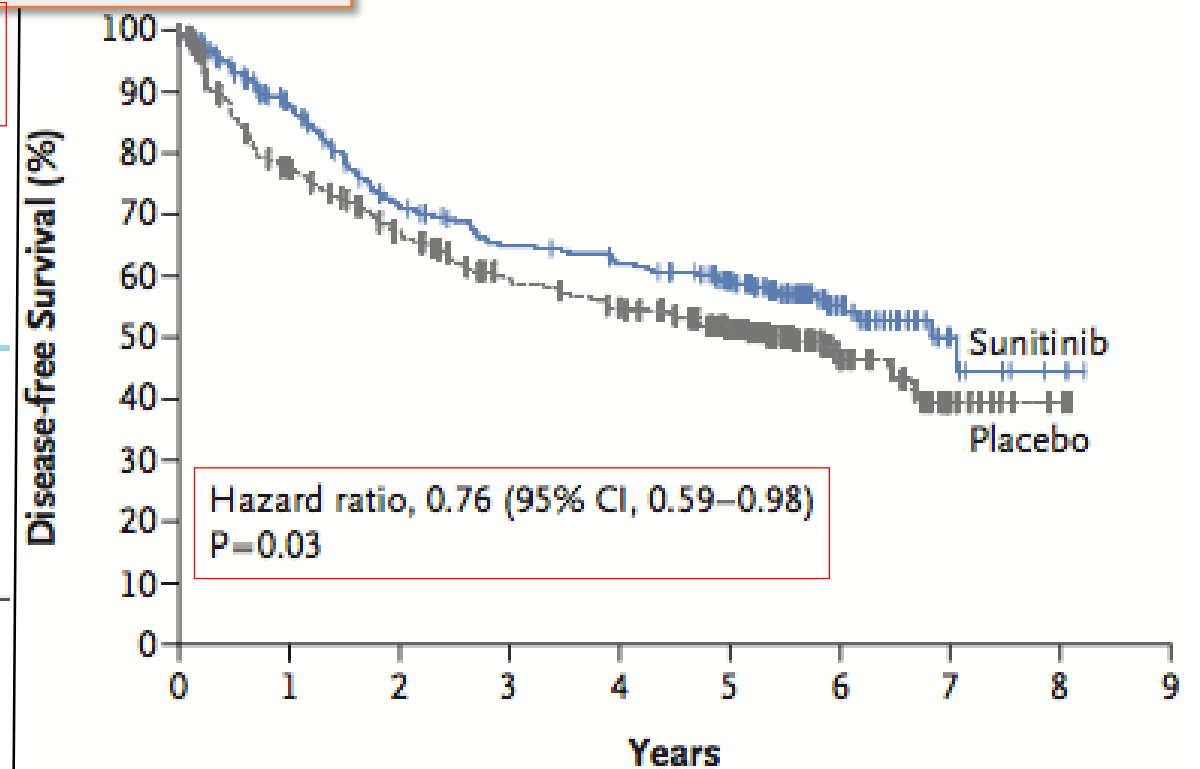
Essai ASSURE

Disease-Free Survival



- **Médiane DFS** de 5.8, 6.1 et 6.6 années pour Sunitinib, Sorafenib et Placebo

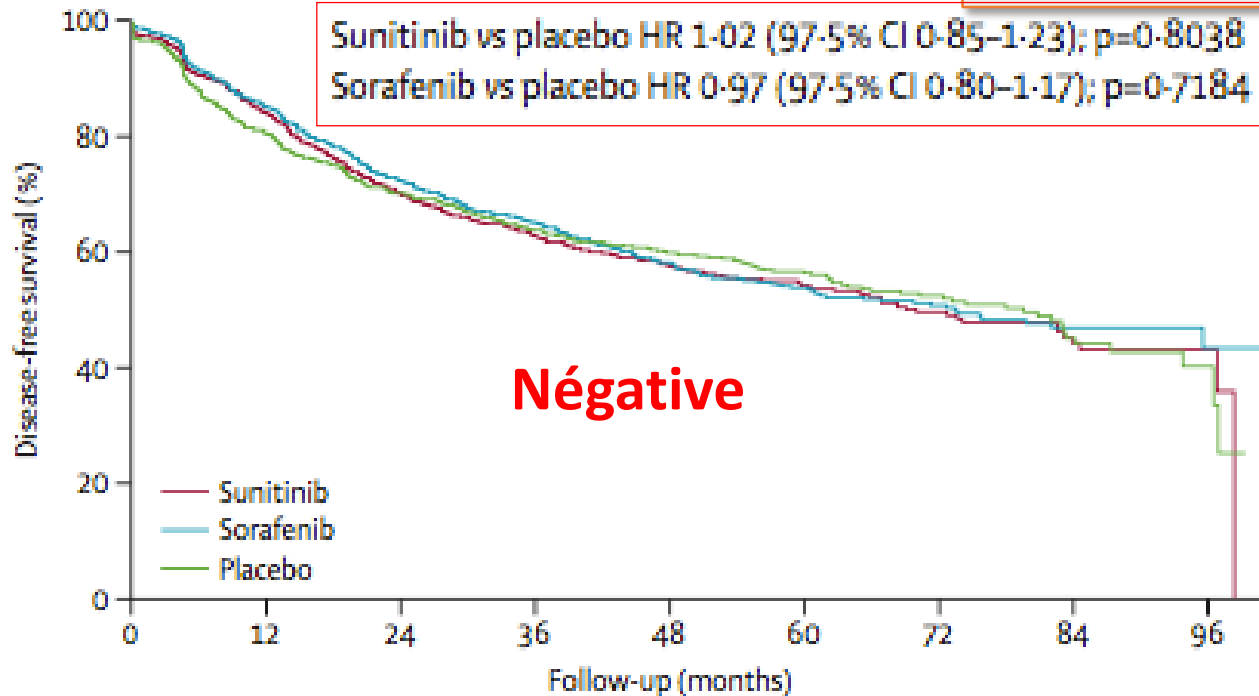
Essai S-TRAC



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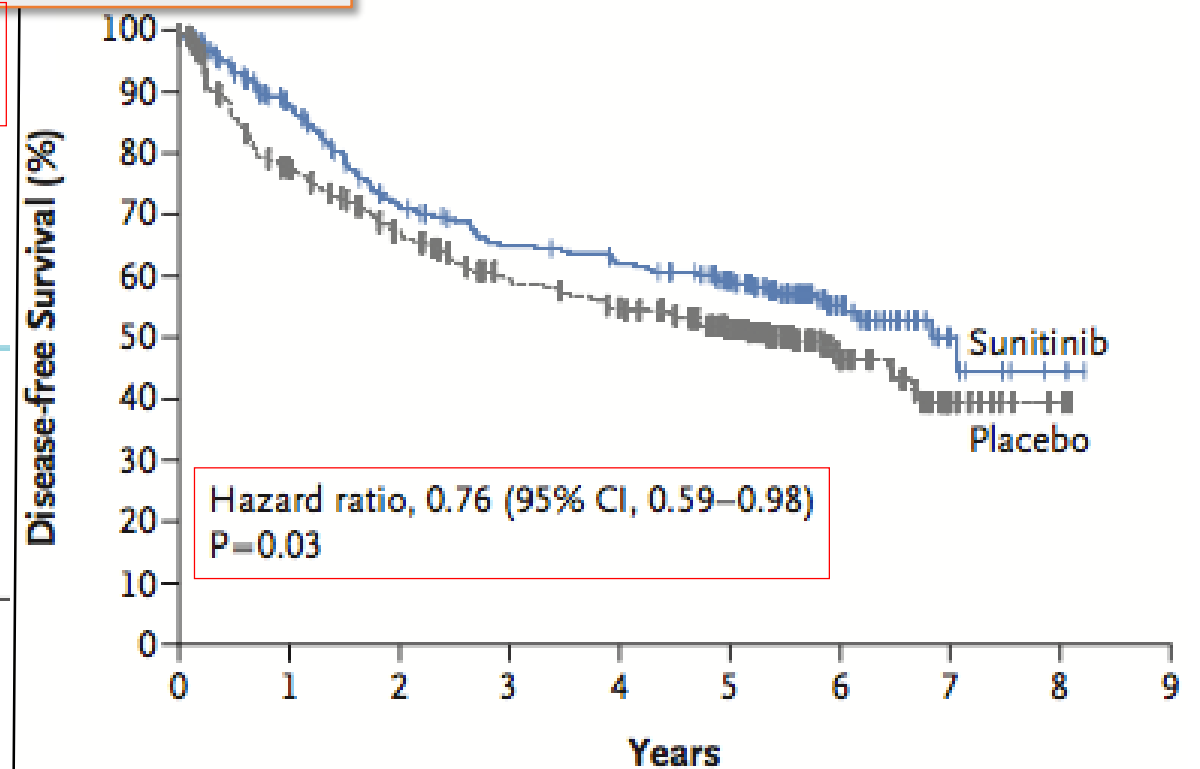
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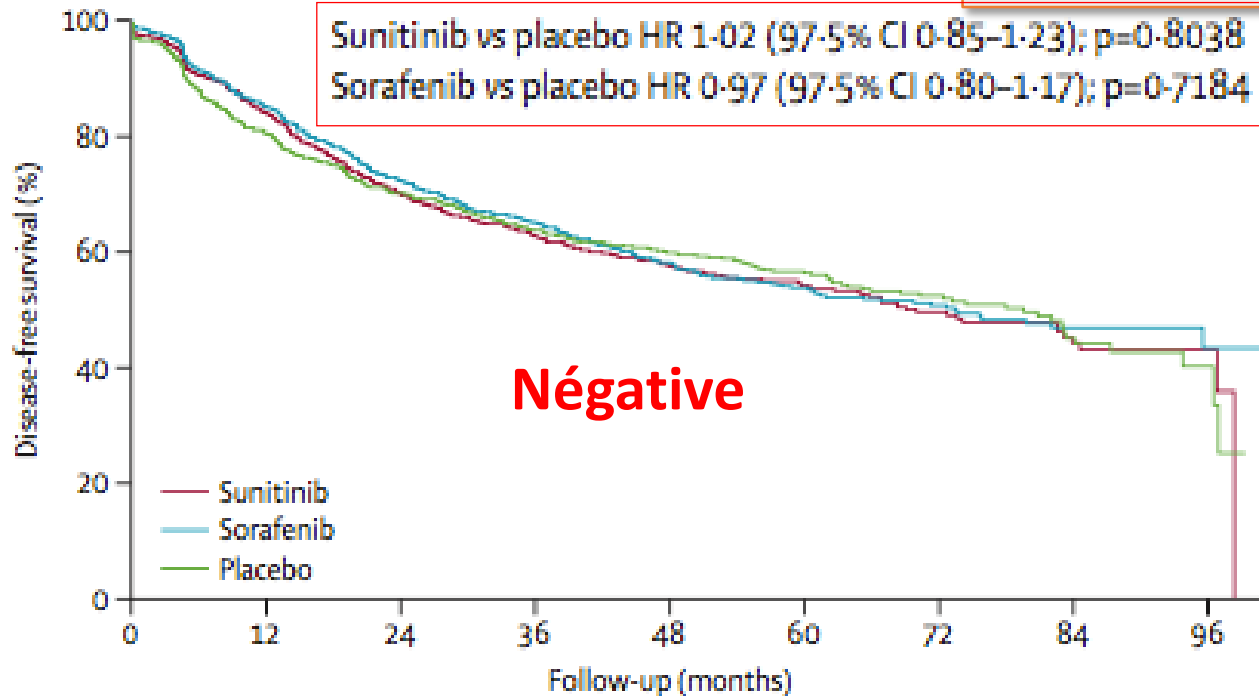
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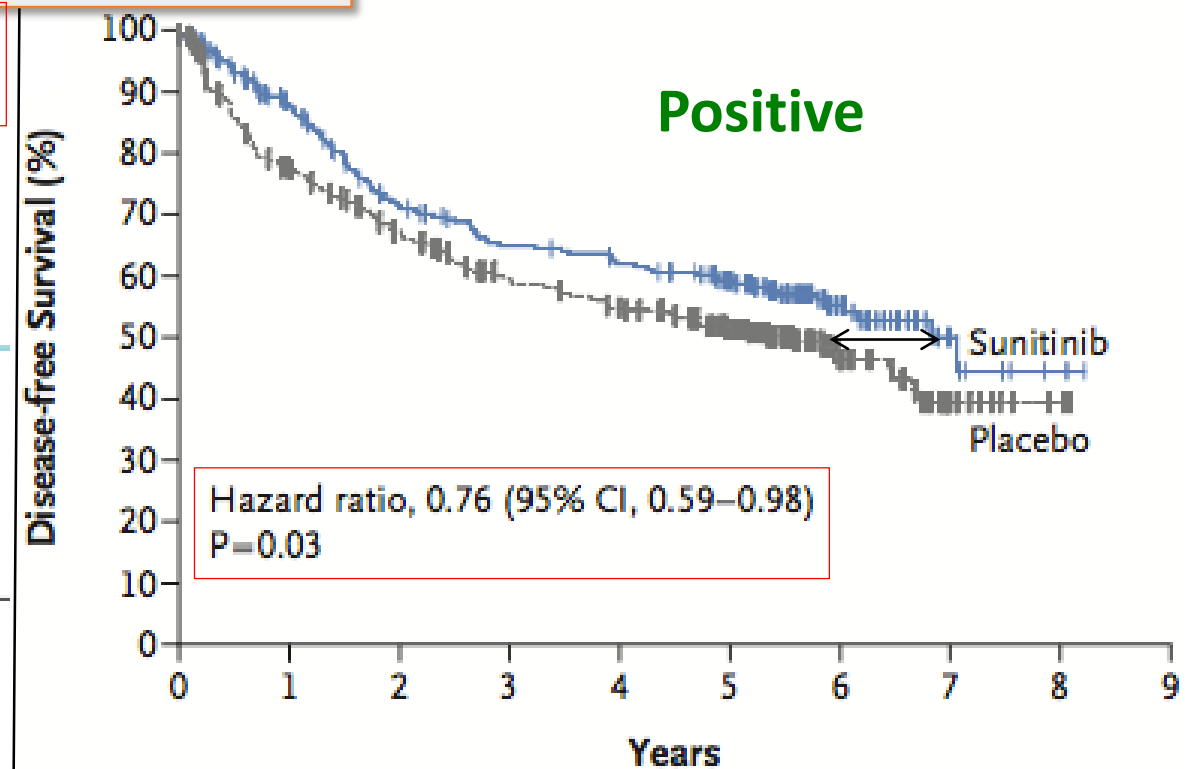
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Disease-Free Survival



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Un bref historique des essais d'immunothérapie adjuvants

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We came a long way... ASCO 1992: 1st RCT of adjuvant immunotherapy in RCC

How it started...

ASCO 1996
Cytokine therapy
Lymphoblastoid
Interferon

1996
Randomized, controlled trial of adjuvant therapy with lymphoblastoid interferon (LBI) in resectable renal cell carcinoma (RCC). The purpose of this study was to evaluate the efficacy of LBI in the adjuvant setting. Patients were randomized to receive either LBI or placebo. The primary endpoint was overall survival. Secondary endpoints included time to recurrence, time to progression, and quality of life. The study was conducted in a multicenter setting. The results of the study showed that LBI was not superior to placebo in terms of overall survival or any of the other endpoints. However, there was a trend towards improved time to recurrence in the LBI group. The study was limited by a small number of patients and a relatively short follow-up period.

How it is going...

ASCO 2021
KEYNOTE-564

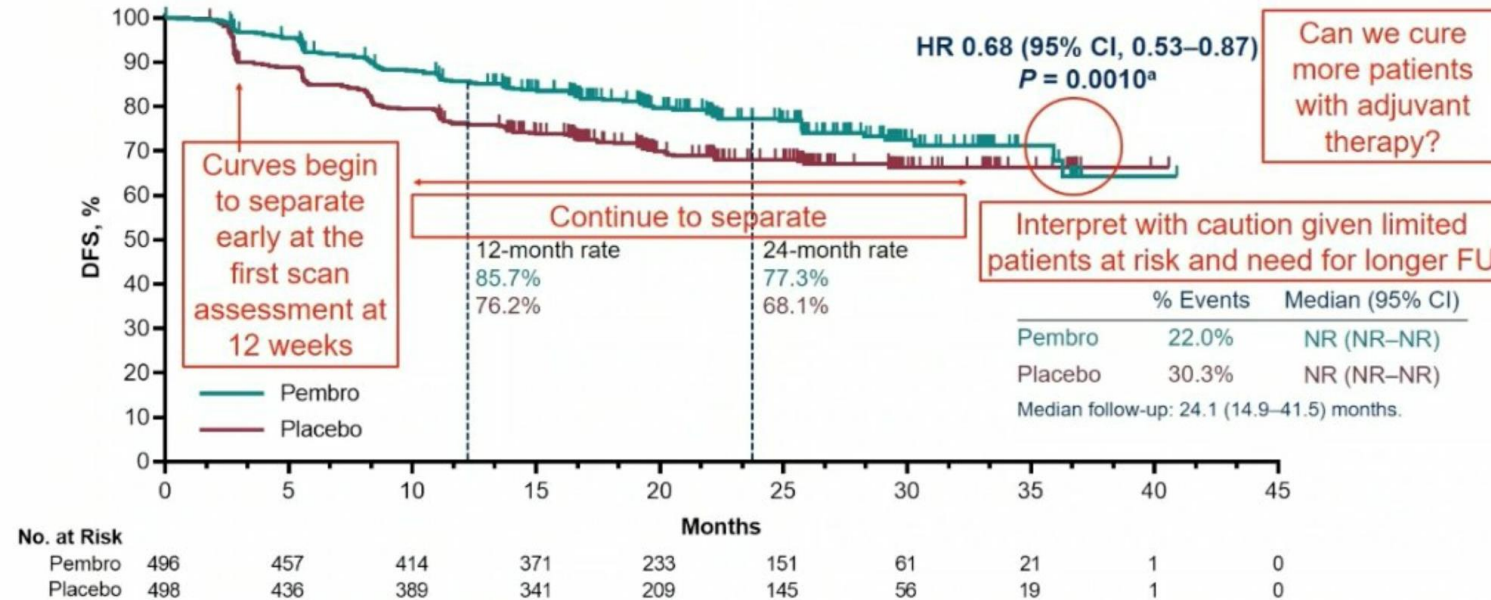
ASCO 1992
Cytokine therapy
Interferon alpha-2a

1992
Randomized, controlled trial of adjuvant therapy with interferon alpha-2a (IFN-α2a) in resectable renal cell carcinoma (RCC). The purpose of this study was to evaluate the efficacy of IFN-α2a in the adjuvant setting. Patients were randomized to receive either IFN-α2a or placebo. The primary endpoint was overall survival. Secondary endpoints included time to recurrence, time to progression, and quality of life. The study was conducted in a multicenter setting. The results of the study showed that IFN-α2a was not superior to placebo in terms of overall survival or any of the other endpoints. However, there was a trend towards improved time to recurrence in the IFN-α2a group. The study was limited by a small number of patients and a relatively short follow-up period.

2000s

- Autologous tumor cell vaccines
- Cytokine therapy
- Cytokine + cytokine combination
- Cytokine + chemotherapy
- Autologous tumor cells + BCG

Keynote 564 : nouveau paradigme?



- 2020' : Décennie du basculement des immunothérapies en stade précoce ?
 - Cet ASCO 2021 semble l'annoncer (également mélanome, poumon, vessie)