

Avons nous les moyens de nous payer l'immunothérapie?

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CHU Limoges



U707



Liens d'intérêt



Participation à des boards,
Consultant,
Voyages pour des congrès scientifiques,
Actions de formation
Subventions pour des essais cliniques

- Amgen
- Astra Zeneca
- BMS
- Boehringer
- Lilly
- MSD
- Roche
- Teva

Epidémiologie UE

CBP

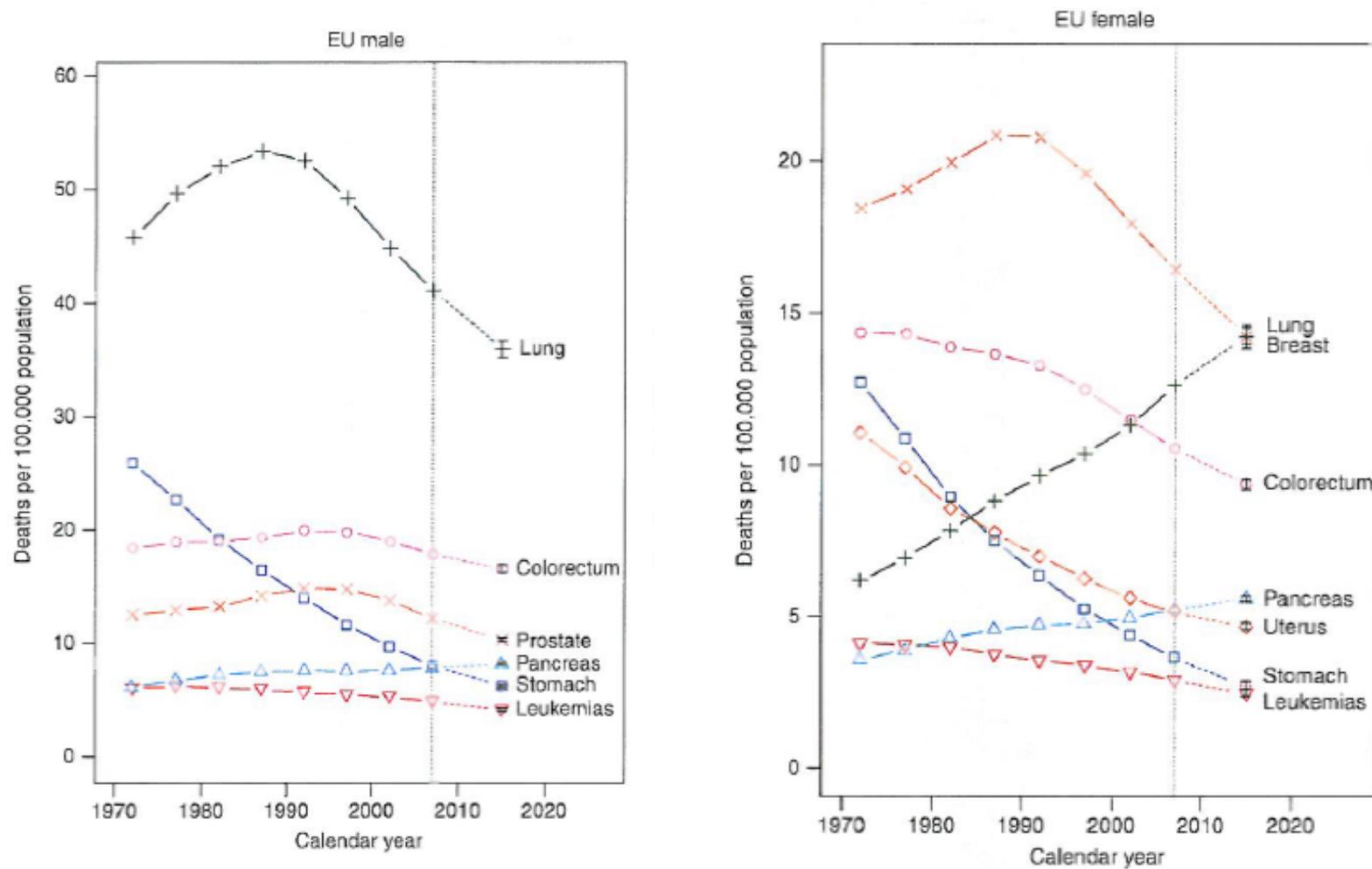
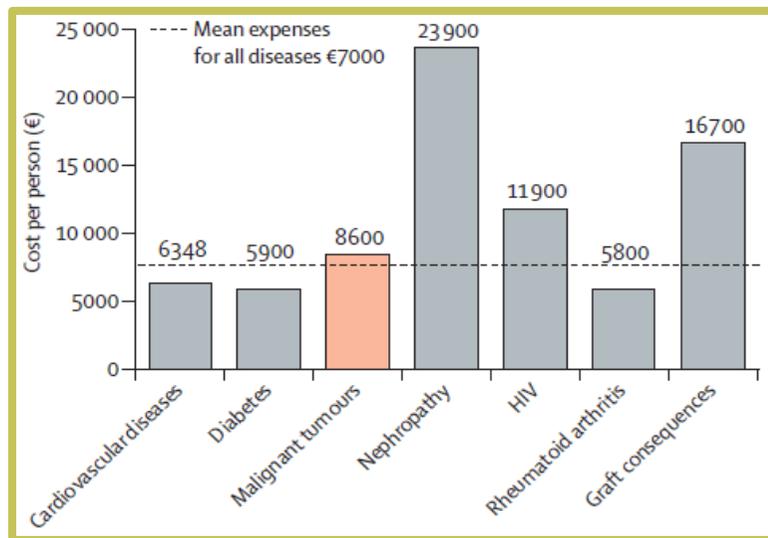
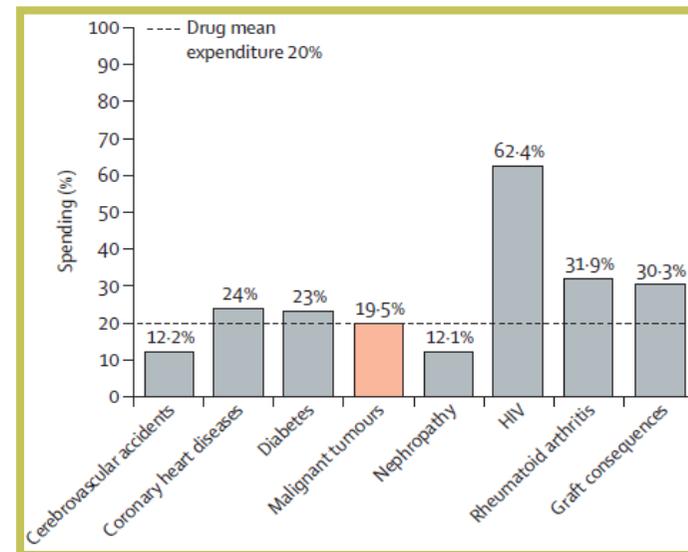


Figure 3. Age-standardised (world population) EU male and female cancer mortality rate trends in quinquennia from 1970–1974 to 2005–2009 and predicted rates for 2015 with 95% prediction intervals (PIs). Men: stomach (squares), colorectum (circles), pancreas (triangles), lung (crosses), prostate (xs) and leukaemias (inverted triangles). Women: stomach (squares), colorectum (circles), pancreas (triangles), lung (crosses), breast (xs), uterus (diamonds) and leukaemias (inverted triangles).

Le contexte



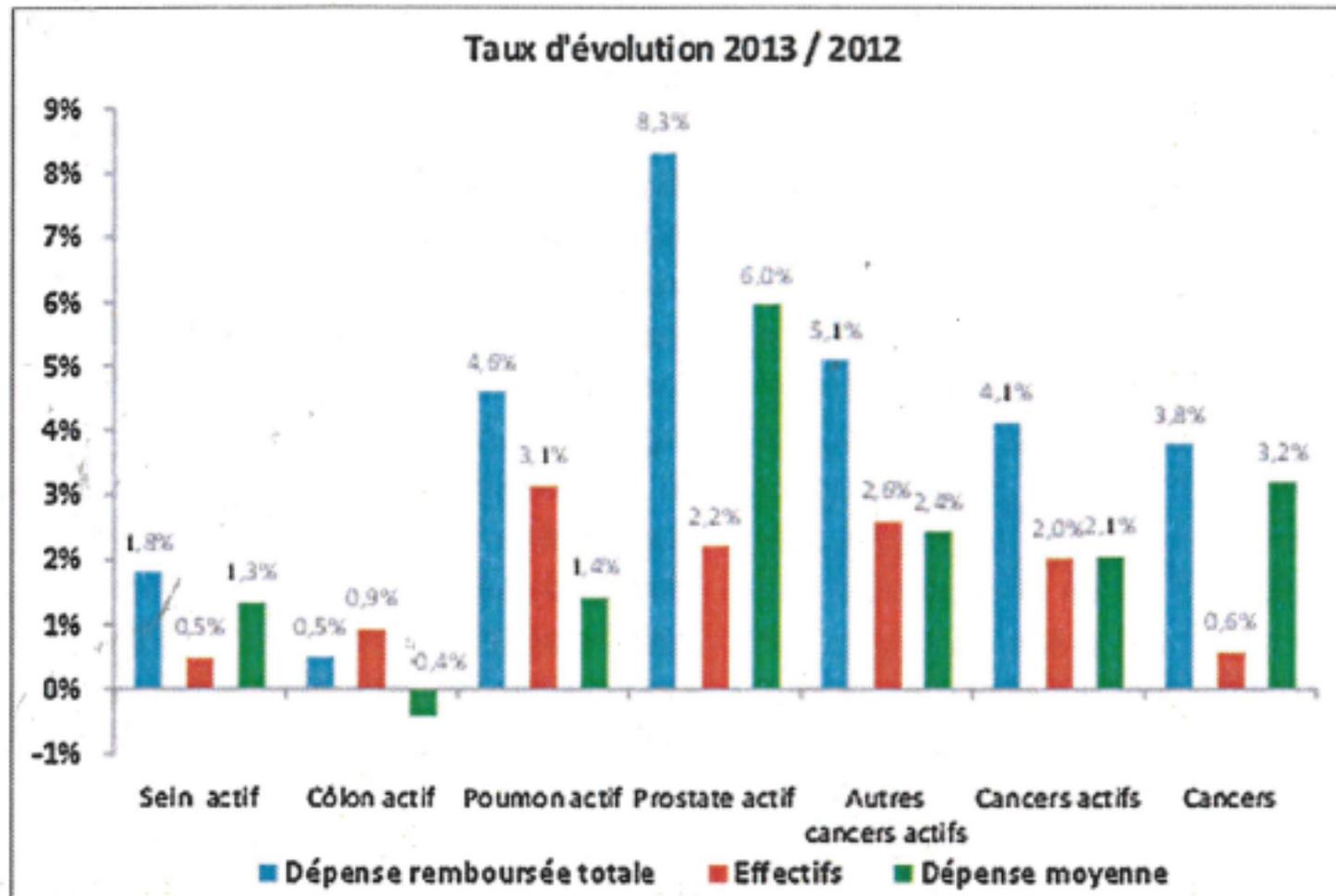
Le coût du cancer en France



La part des médicaments dans le coût des maladies chroniques en France

FIGURE 2

Évolution des dépenses attribuables au cancer et des effectifs (2012-2013 et 2013-2014), selon le type de cancer, pour les bénéficiaires du régime général d'assurance maladie



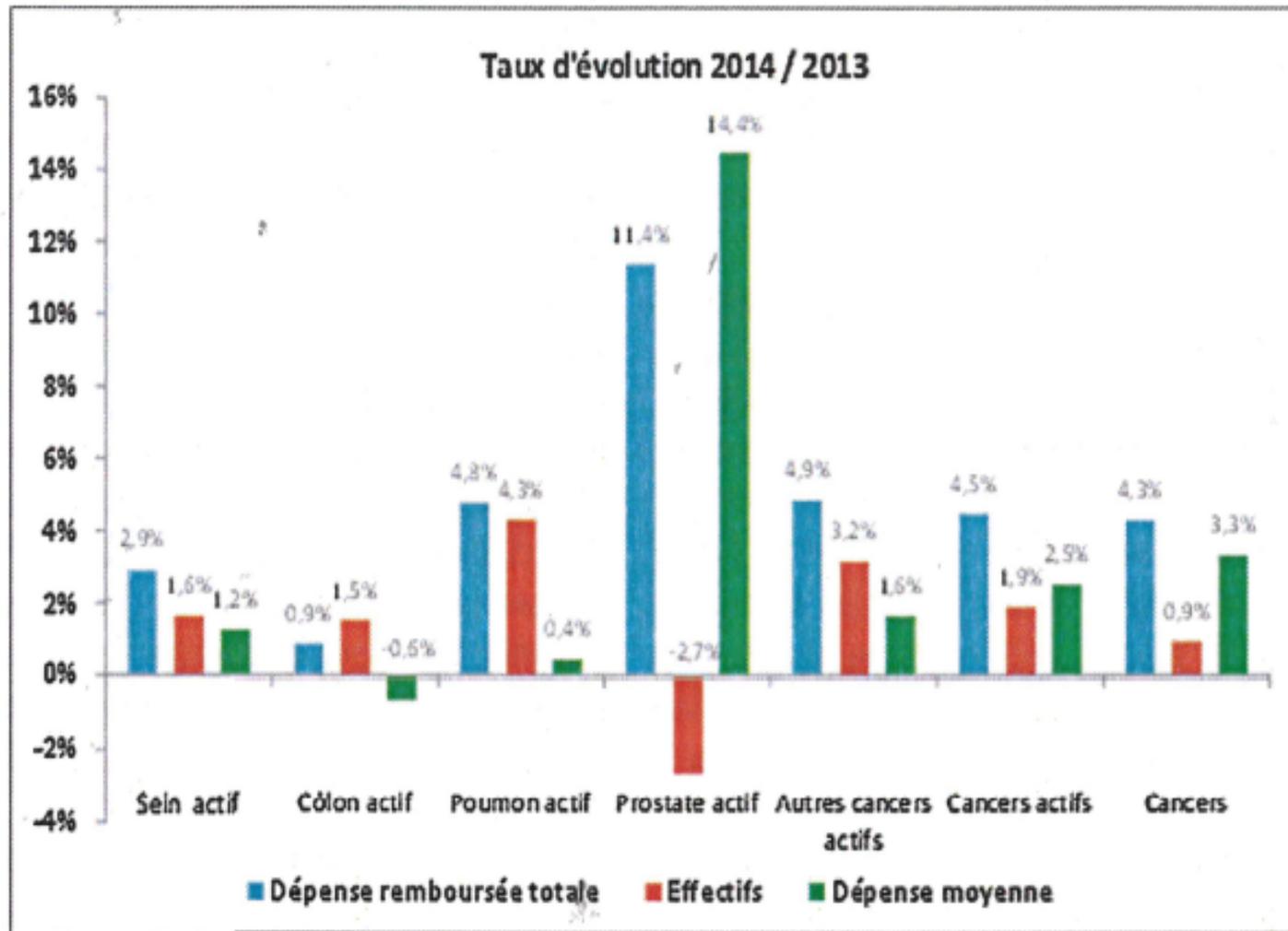


FIGURE 2

Évolution des dépenses attribuables au cancer et des effectifs (2012-2013 et 2013-2014), selon le type de cancer, pour les bénéficiaires du régime général d'assurance maladie

Les innovations thérapeutiques

- Distribution of (A) patients and (B) insurance payments by type of chemotherapy agent from 2001 to 2011. Analysis conducted using life-link health plan claims database from January 2001 to september 2011. IV,intravenous.

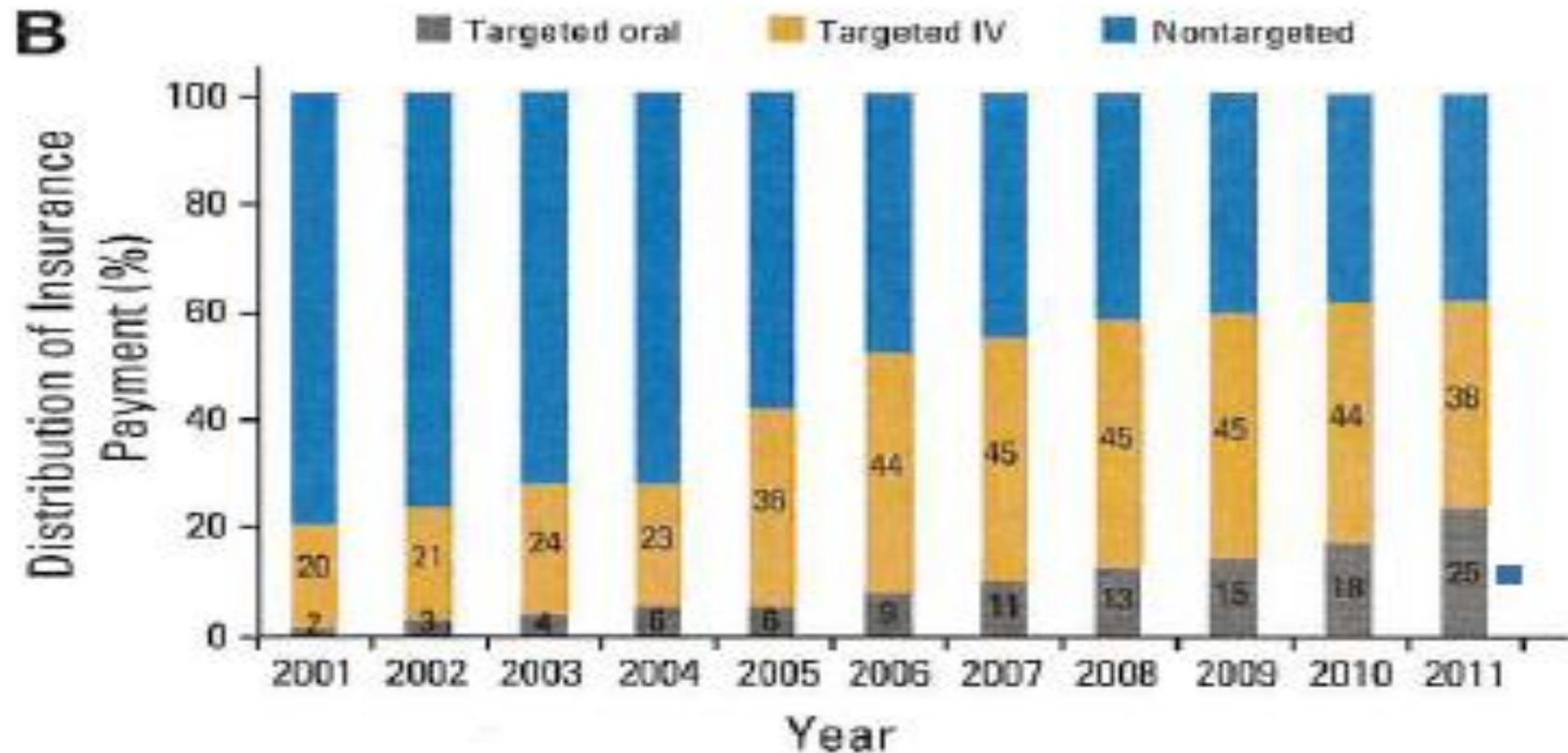
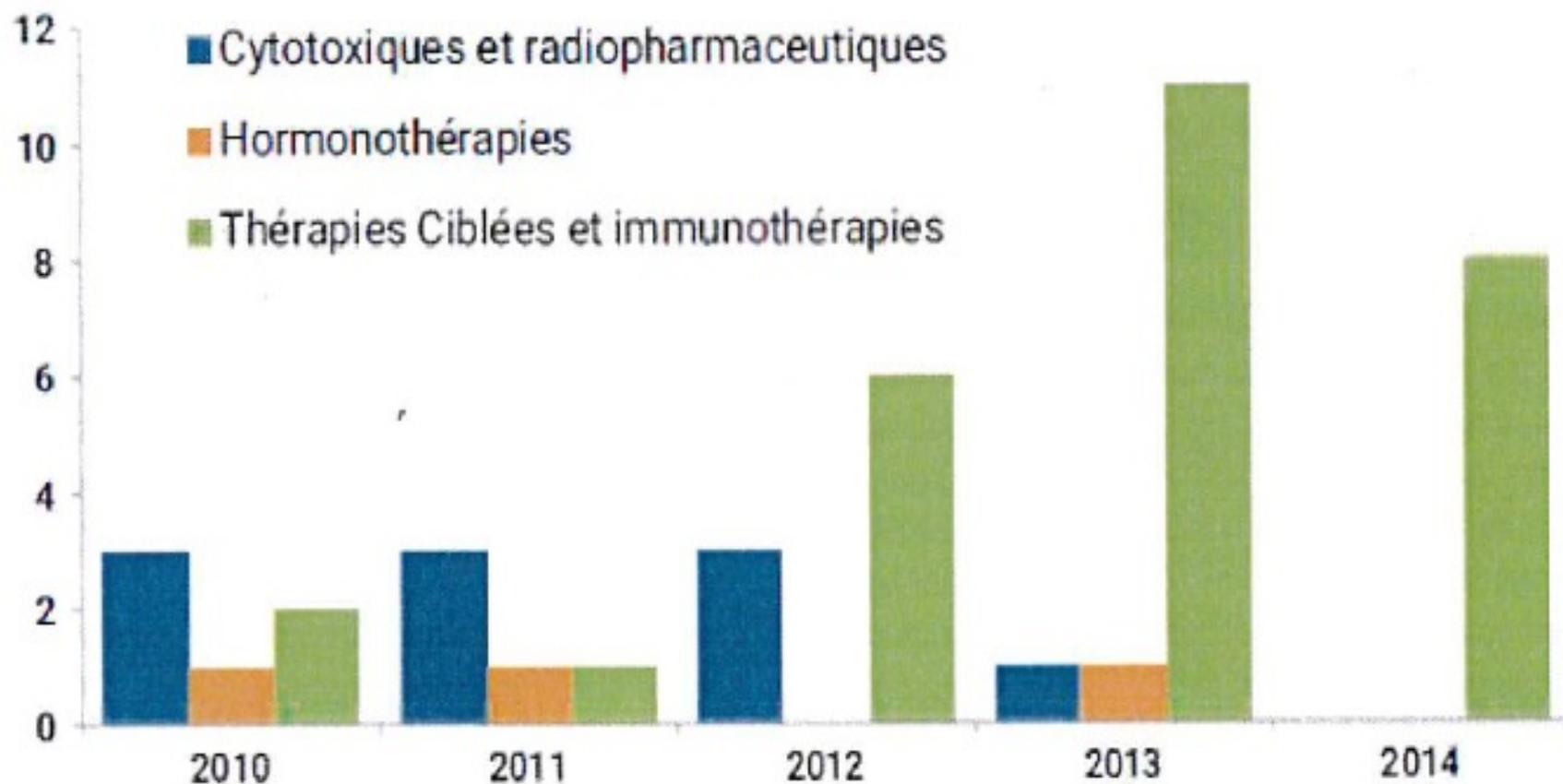


Figure : classes des molécules mises sur le marché pour la première fois sur la période 2010-2014.



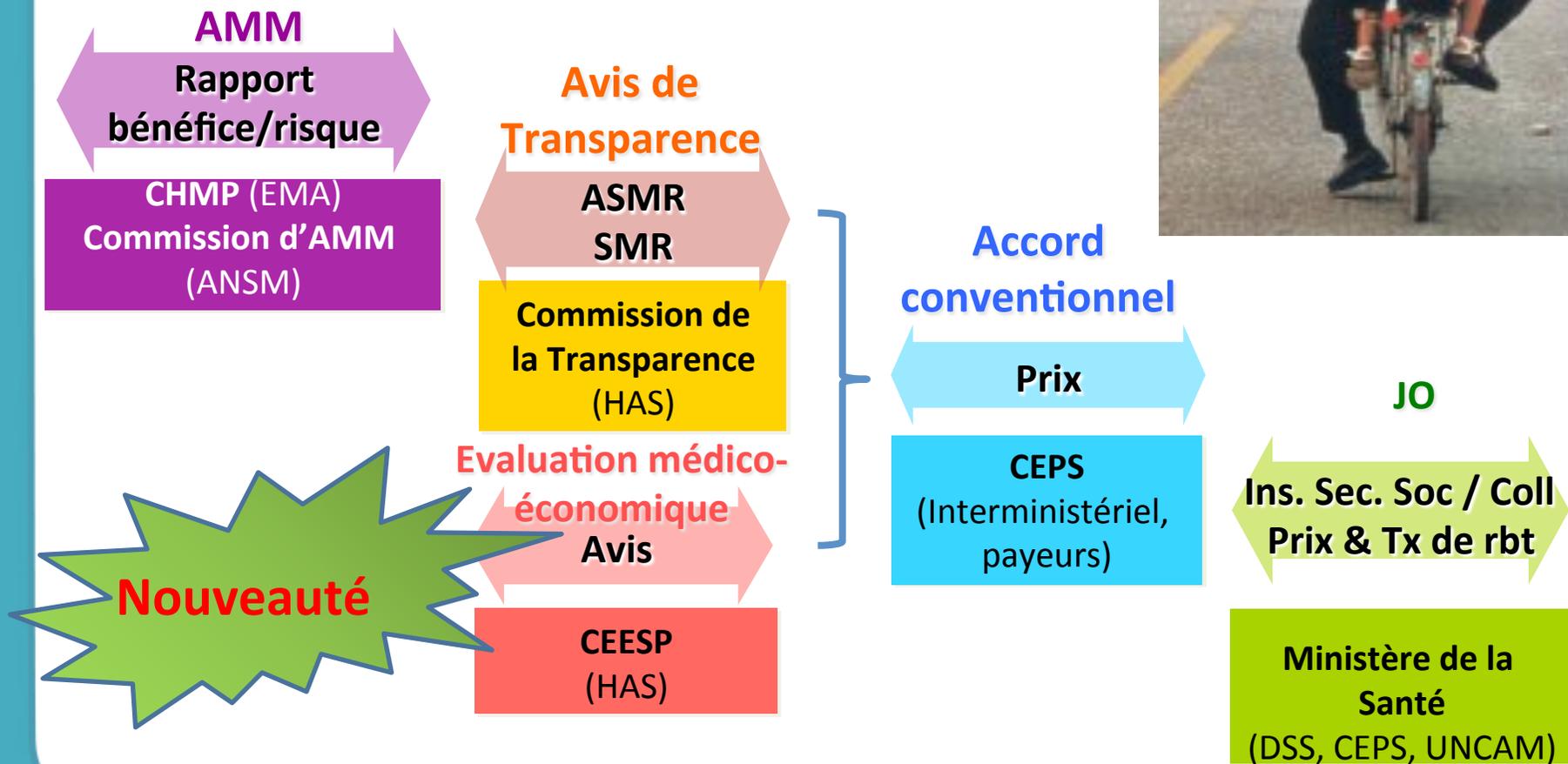
Source : INCa (la chimiothérapie orale du cancer en 2014 – note d'analyse, décembre 2015).

Accessibilité des nouvelles drogues en Europe

LUNG CANCER : Formulary and cost				
Country:	Erlotinib	Gefitinib	Afatinib	Crizotinib
Austria	Free	Free	Free	Free
Belgium	Free	Free	Full cost	Free
Cyprus	Free	Free	Full cost	Full cost
Denmark	Free	Free	Free	Free
Finland	Free	Free	Discount >50% and <100%	Free
France	Free	Free	Free	Free
Germany	Free	Free	Free	Free
Greece	Free	Free	Full cost	Free
Holland	Free	Free	Discount >50% and <100%	Free
Iceland	Free	Free	Free	Free
Ireland	Free	Free	<25% cost	Free
Israel	Free	Free	Free	Free
Italy	Free	Free	Full cost	Free
Luxembourg	Free	Free	Free	Free
Norway	Free	Free	Free	Free
Portugal	<25% cost	<25% cost	Free	Free
Spain	Free	Free	Free	Free
Sweden	Free	Free	Discount >50% and <100%	Free
Switzerland	<25% cost	<25% cost	<25% cost	<25% cost
Turkey	Free	Full cost	Full cost	Free
United Kingdom	Free	Free	<25% cost	Free

Free
<25% cost
25–50% cost
Discount >50% and <100%
Full cost
Not available
Missing data

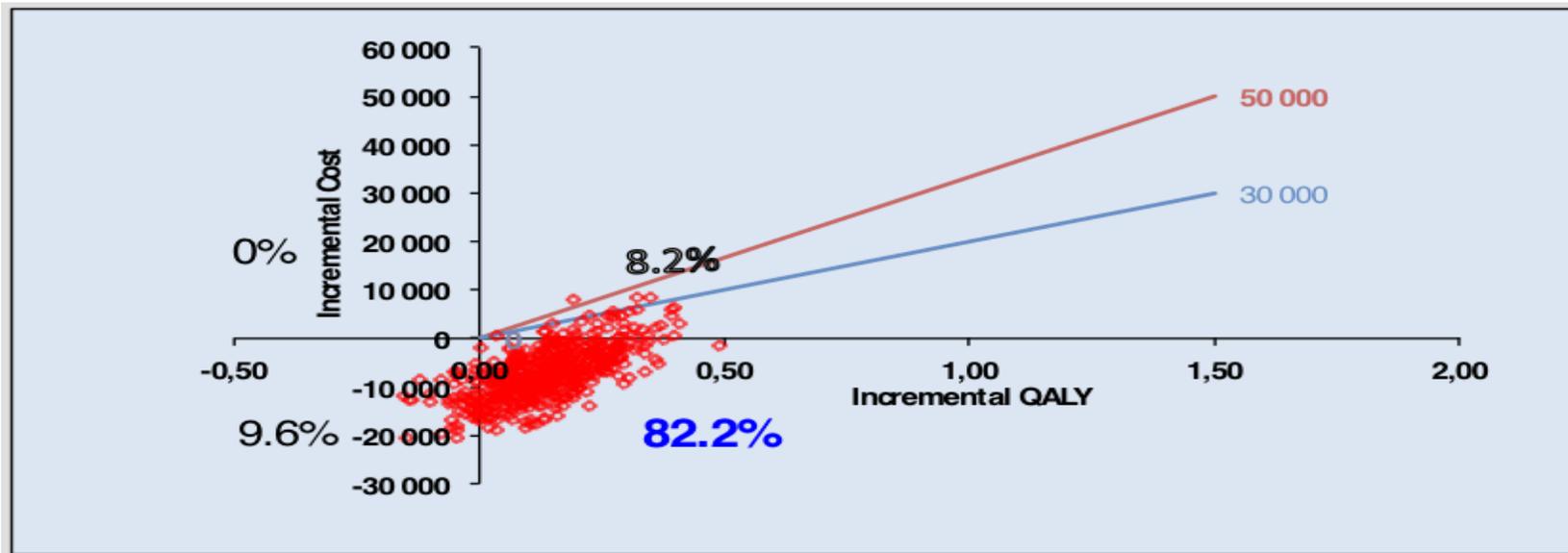
L'accès au marché du médicament En 2015



Incremental cost/utility ratio according to the 3 countries

	France	Italy	Spain
Cost difference	-9,498	-7,806	-7,466
Utility difference	0,117	0,117	0,117
ICER	negative	negative	negative

Results: scatter plot of ICER, probability to be under € 30000 = 90,2%



erlotinib *versus* chemo first line

Corrélation bénéfiques, autorisations, prix des drogues en cancérologie

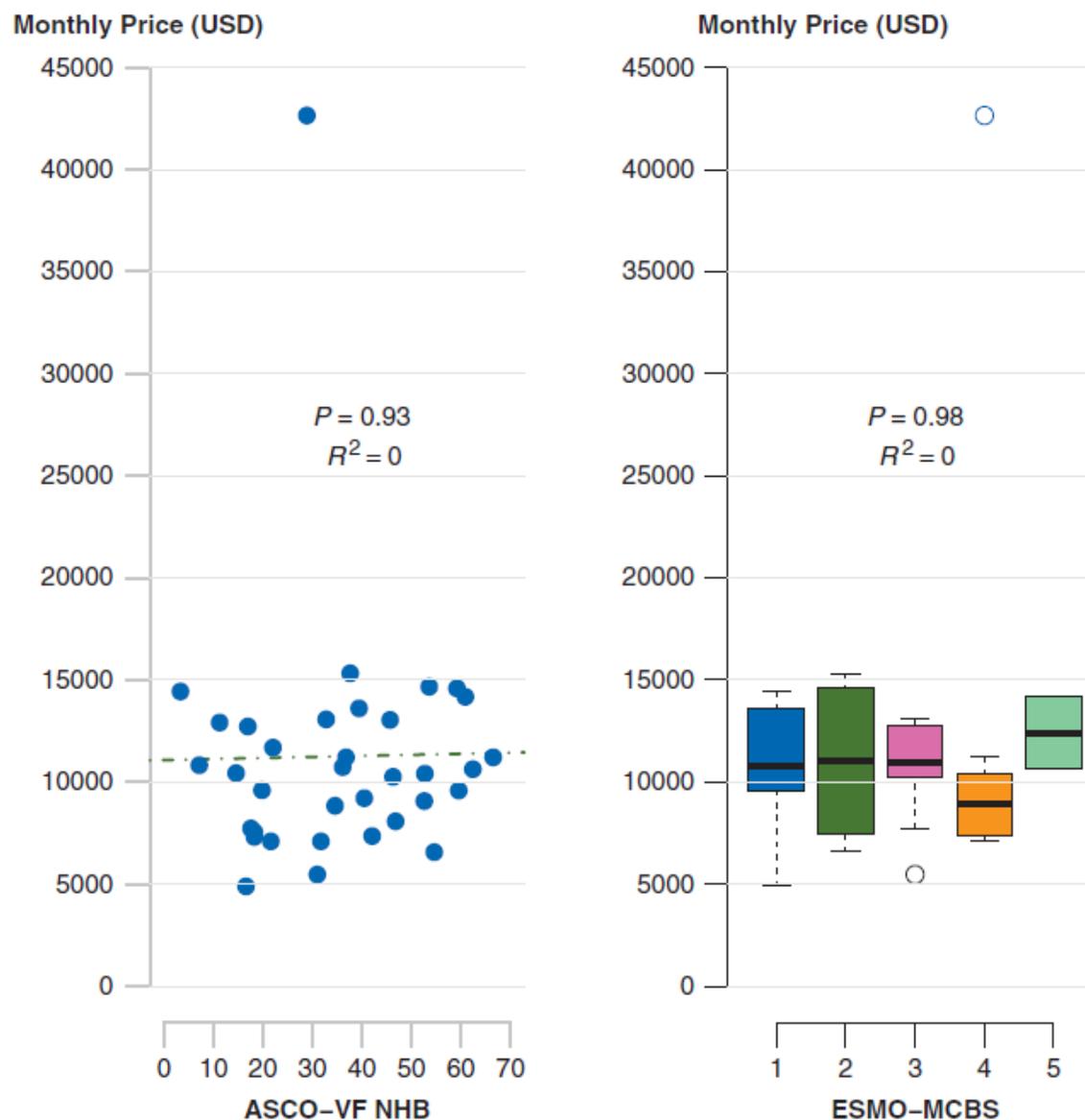
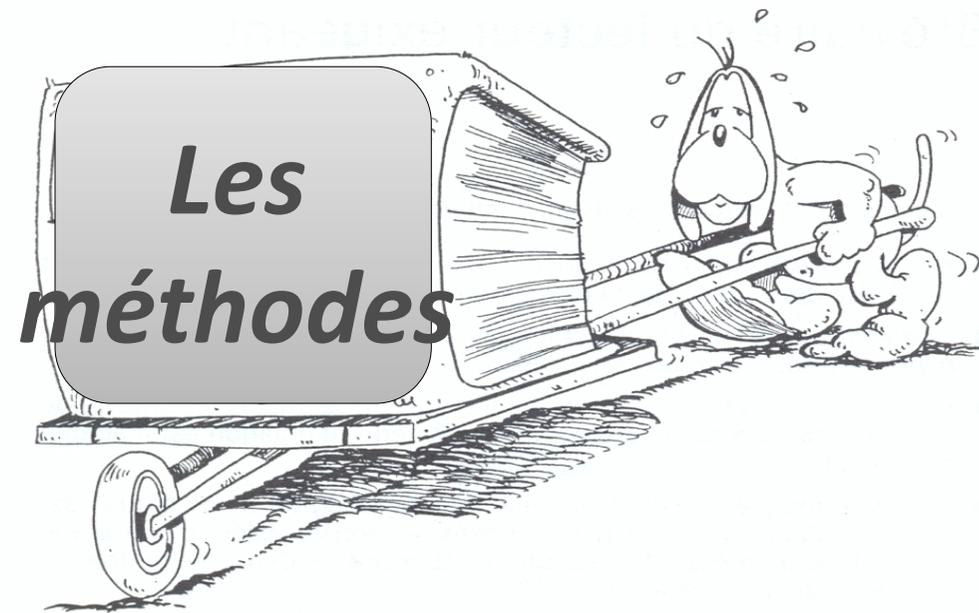


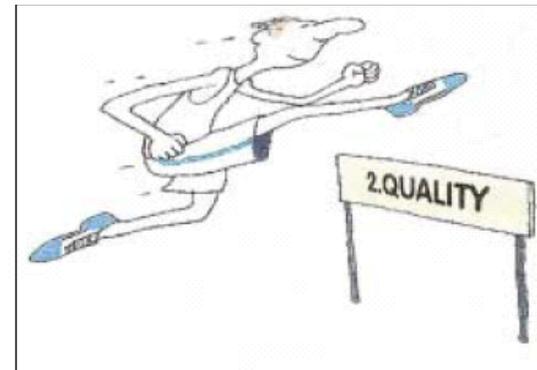
Figure 2. Relationship between the clinical benefit of the 37 anticancer drugs approved by the FDA from 2000 to 2015 as evaluated by the 2016 update of the ASCO-VF NHB and the ESMO-MCBS and the price according to US Medicare (data on prices retrieved from DrugAbascus).

Vivot et al, Ann Oncol 2017;28:1111-6

Pharmaco-économie

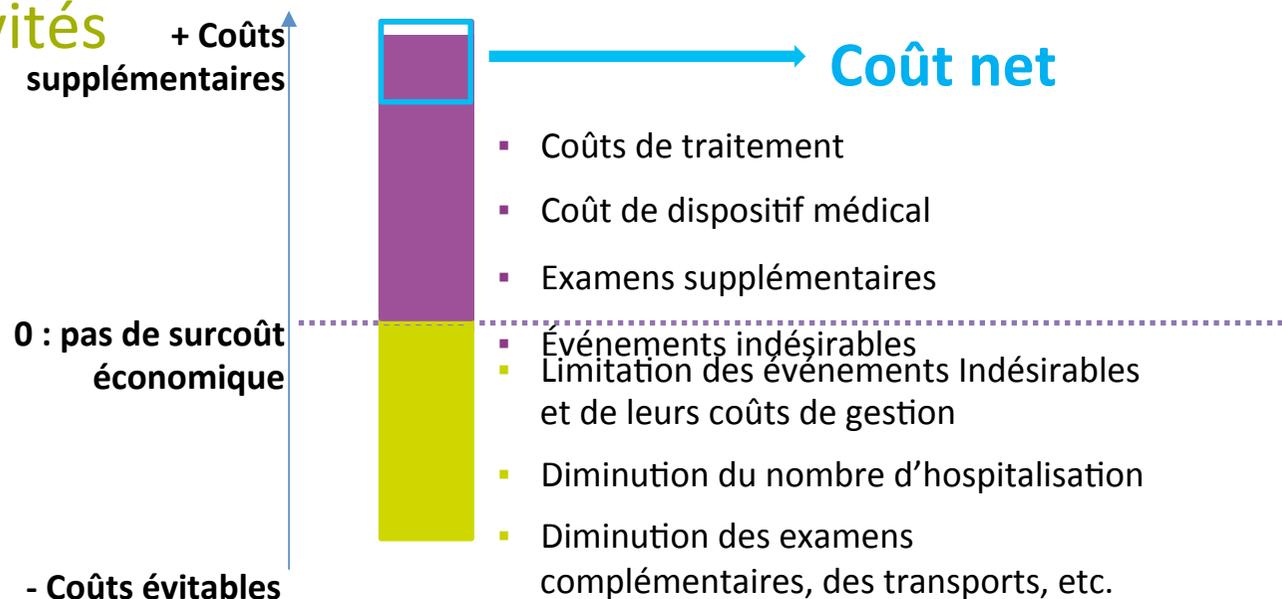


Décision médicale

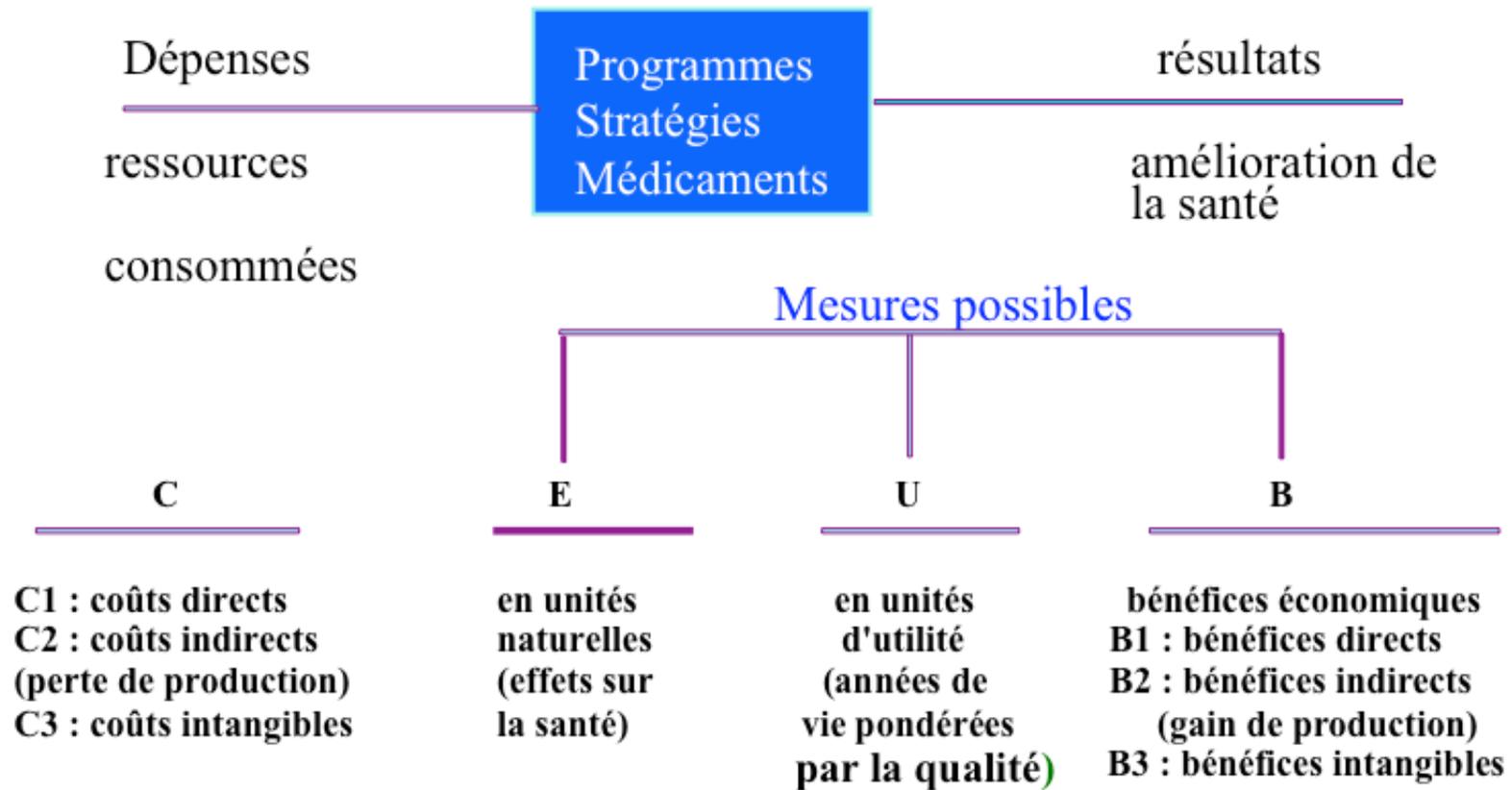


Raisonnement en coût net

Approche en **coût net** = coûts supplémentaires –
coûts évités

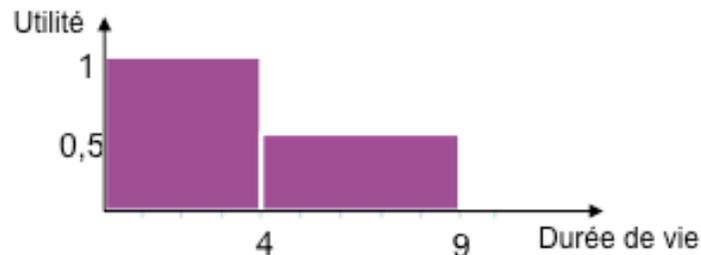


Les différentes techniques de coût efficacité



Qu'est-ce qu'un Qaly?

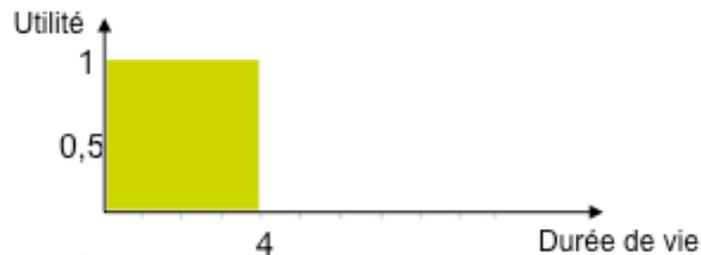
- Le but des QALYs est d'évaluer non seulement la durée de vie mais aussi sa qualité.
- Dans le calcul du QALY, la qualité de vie vaut autant que la quantité de vie.



Ici le patient vit durant 9 ans en pleine santé, puis subit un événement qui altère fortement sa qualité de vie. Il meurt la 9^e année.

Calcul du nombre de QALYs :

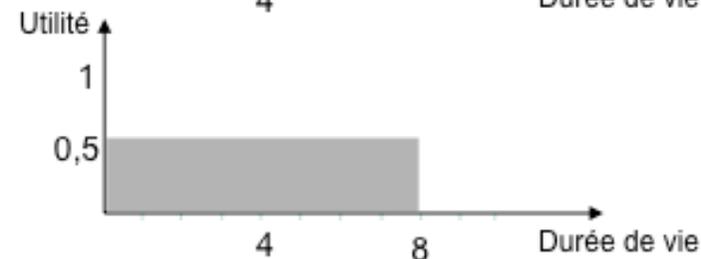
$$4 \times 1 + (0,5 \times 5) = 6,5$$



Le patient vit durant 8 ans en pleine santé puis meurt brusquement.

Calcul de nombre de QALYs :

$$4 \times 1 = 4$$



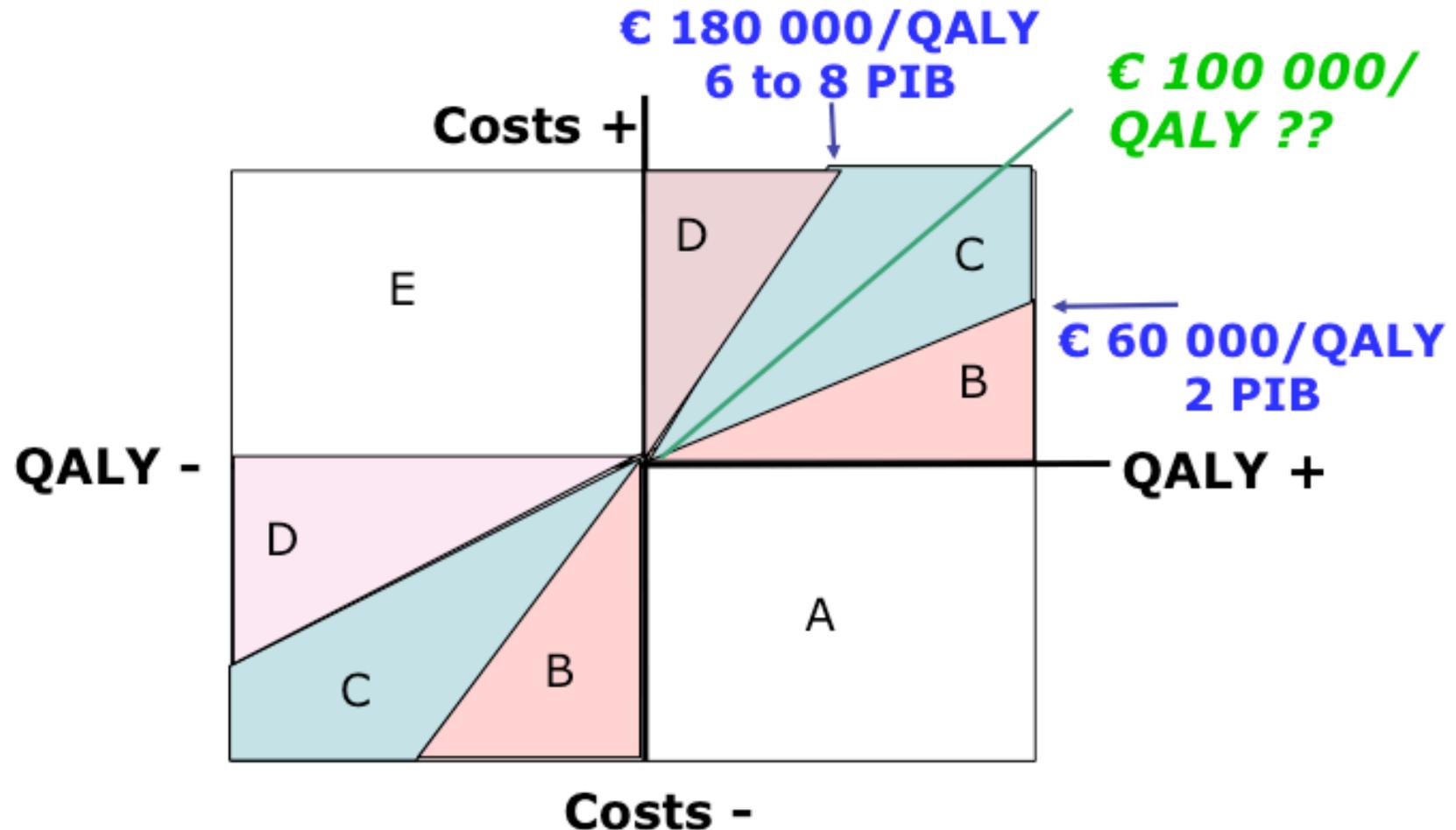
Le patient vit durant 8 ans avec une qualité de vie dégradée puis meurt brusquement.

Calcul de nombre de QALYs :

$$8 \times 0,5 = 4$$

1 QALY = 1 année de vie en parfaite santé

Quels seuils pour une analyse coûts/Qaly en France?



Interprétation de la mesure entre 2 stratégies ou traitements :

A : plus efficace et moins chère ; B : forte préférence ; C : préférence modeste ; D : faible préférence ; E : rejet (moins efficace et plus chère), adapté à la France avec un PIB/h de 32 700€



Le produit en lui-même n'a pas d'effets adverses
mais le prix peut entraîner vertiges et syncope

Economie

CBP immunottt

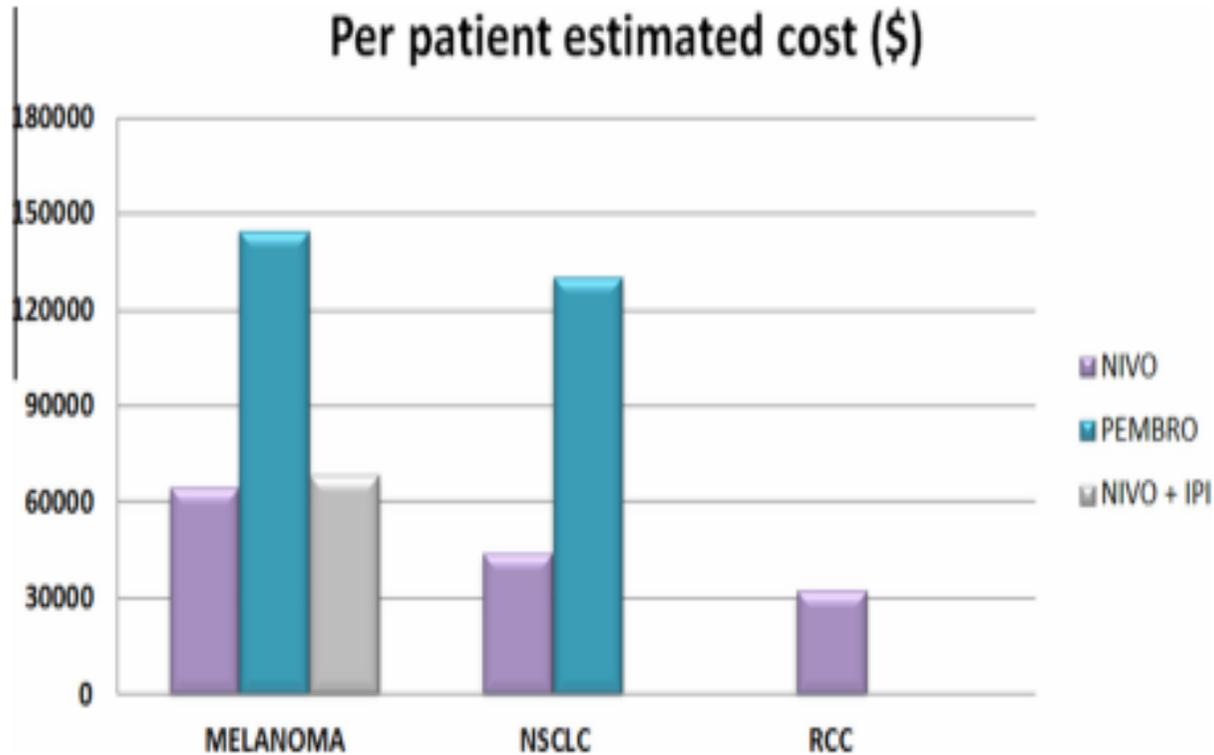


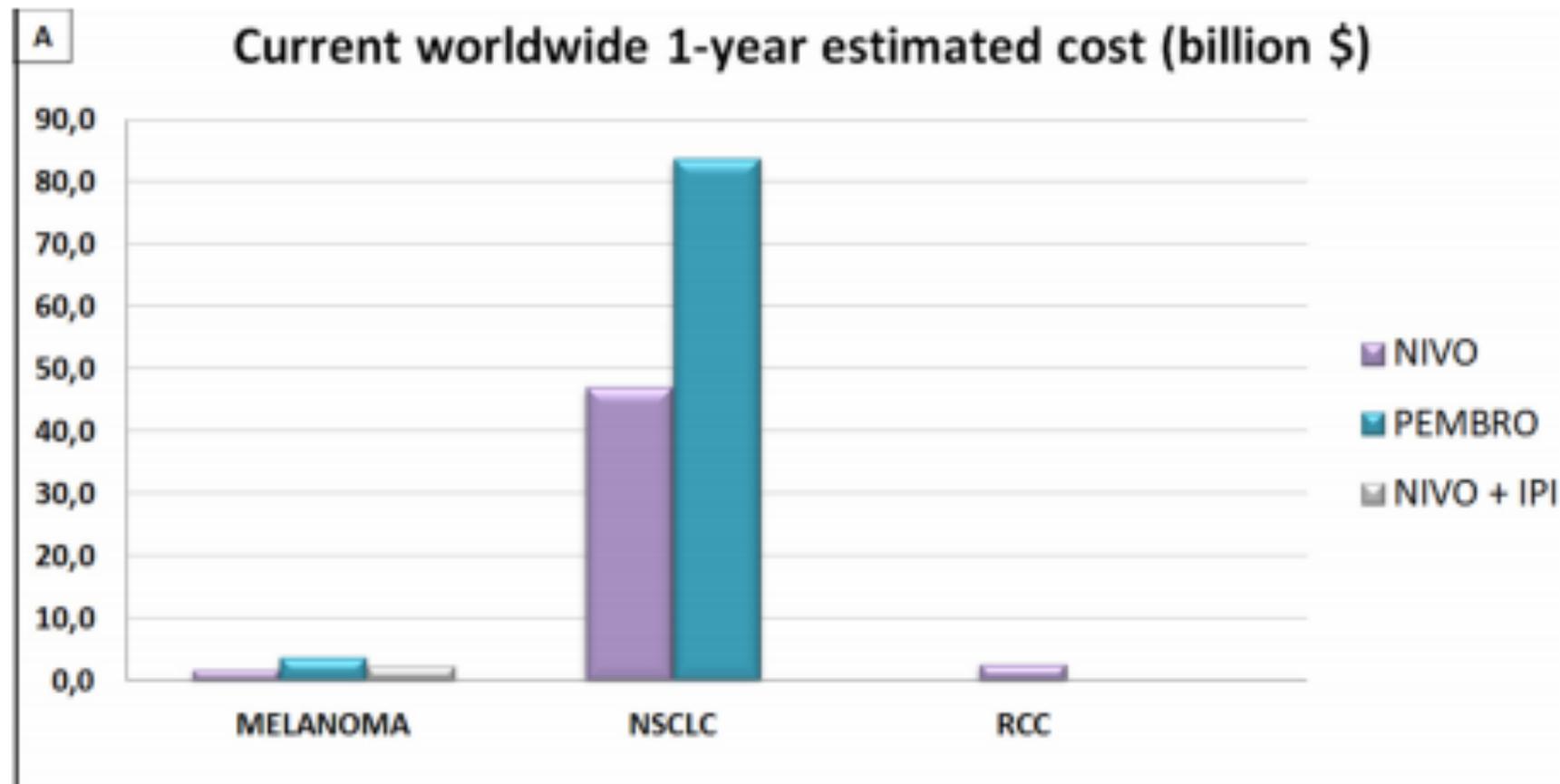
Fig. 1. Per patient estimated cost (\$) of nivolumab and pembrolizumab for the treatment of patients with advanced melanoma, non-small cell lung cancer and renal cell carcinoma. IPI = ipilimumab; Nivo = nivolumab; Pembro = pembrolizumab; NSCLC = non-small cell lung cancer; RCC = renal cell carcinoma.



Economie

CBP immunottt

F. Tartari et al./Cancer Treatment Reviews 48 (2016) 20–24



Tartari et al, 2016;48:20-24,

Economie

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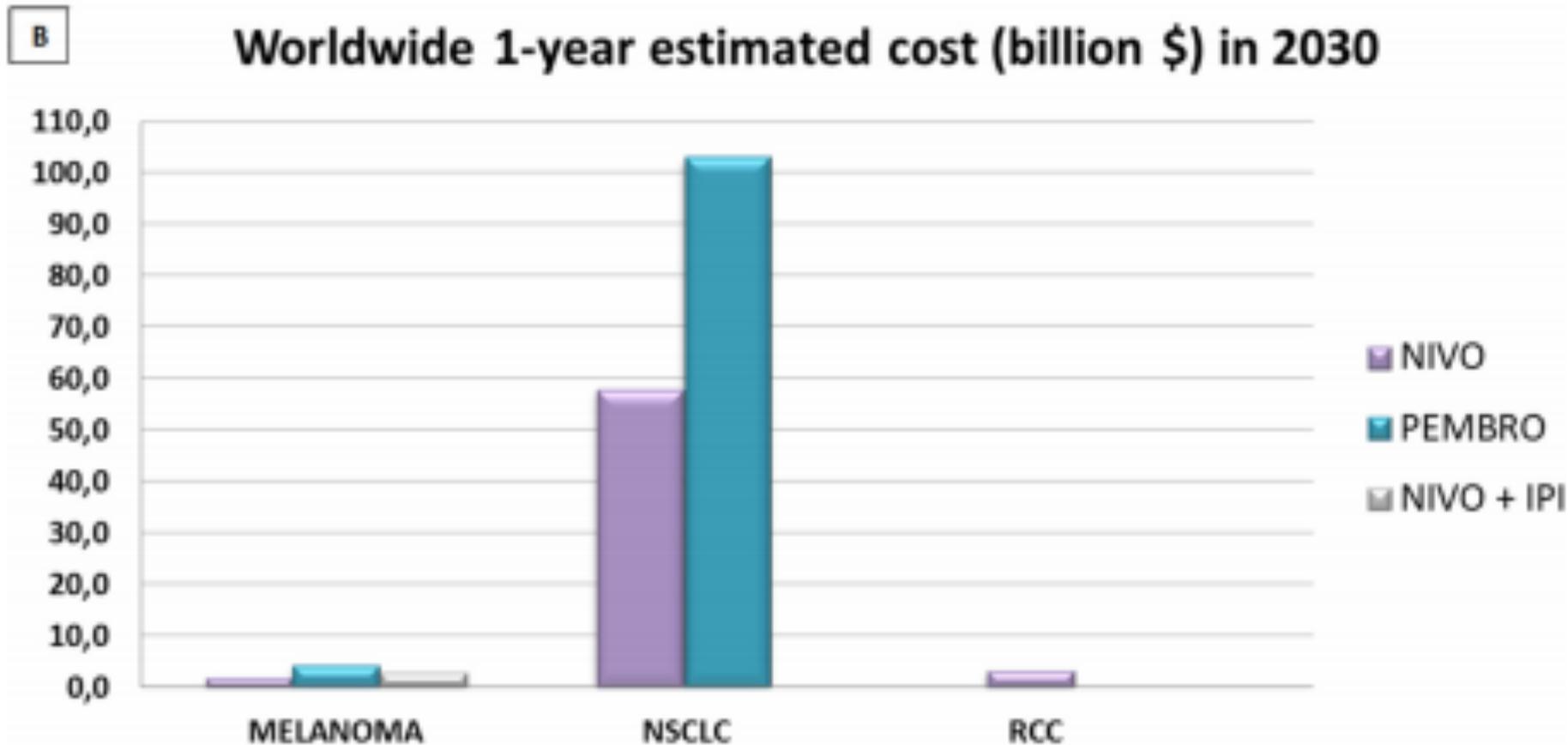
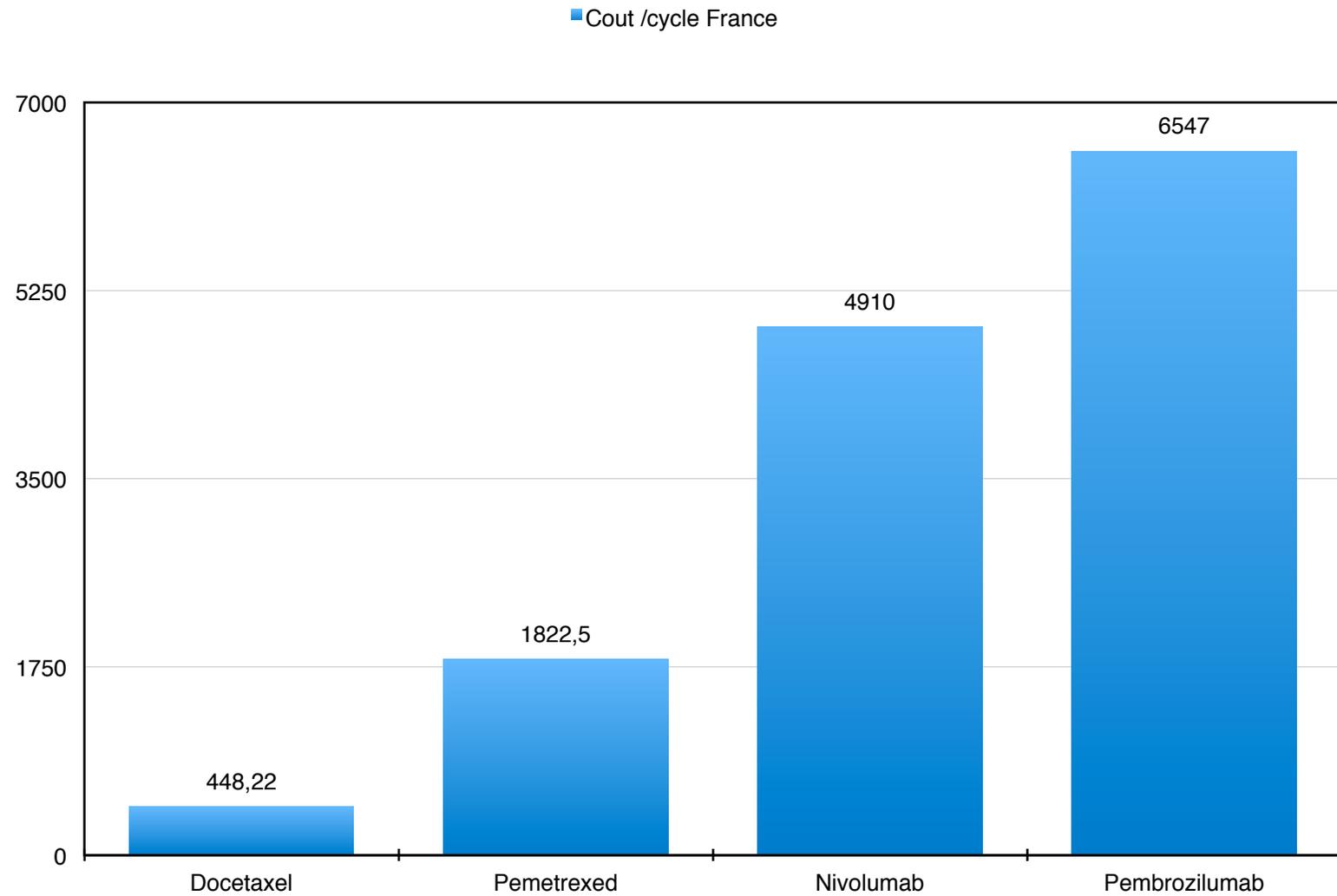


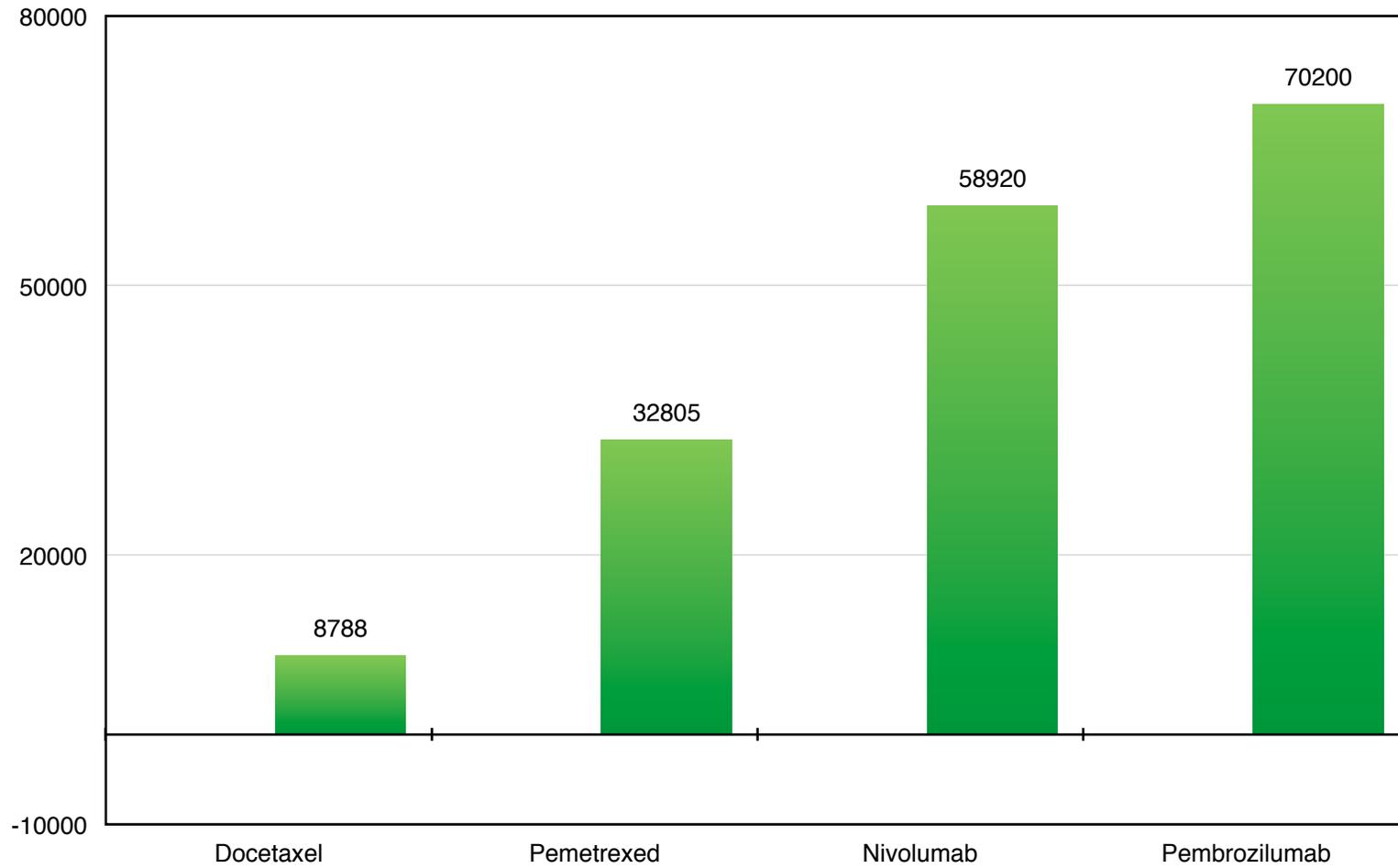
Fig. 2. Worldwide 1-year estimated cost (billion \$) of nivolumab and pembrolizumab for the treatment of patients with advanced melanoma, non-small cell lung cancer and renal cell carcinoma according to WHO data for 2012 (A) and 2030 (B). IPI = ipilimumab; Nivo = nivolumab; Pembro = pembrolizumab; NSCLC = non-small cell lung cancer; RCC = renal cell carcinoma; WHO = World Health Organization.

Analyse économique globale F



Analyse économique globale F

■ Coût/an F (répondeur)



Analyses Médico-économiques



€



Nivolumab seconde ligne, US

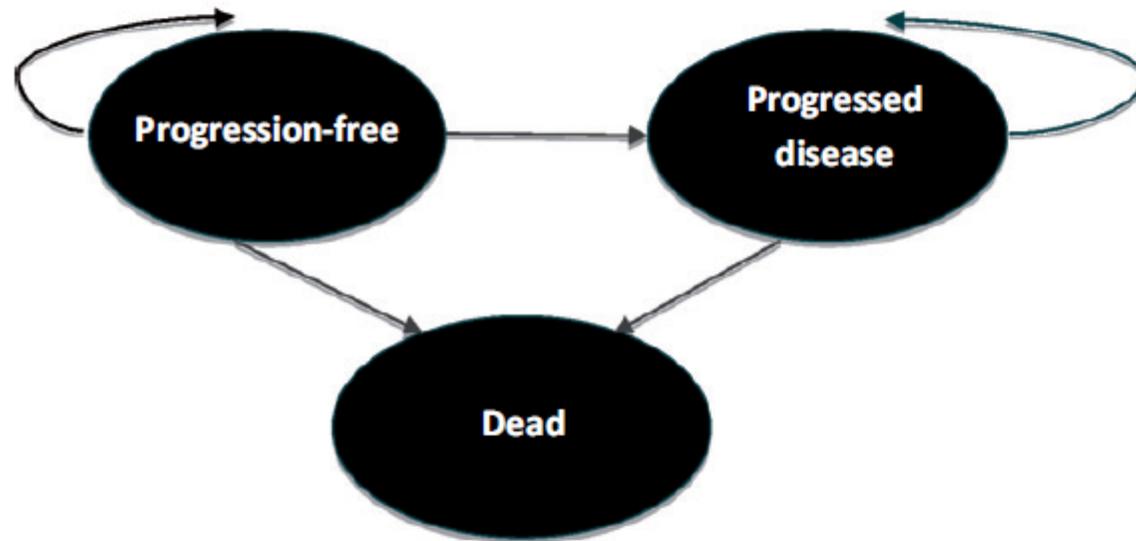


Figure 3. Health state structure for the economic models.

Nivolumab seconde ligne, US

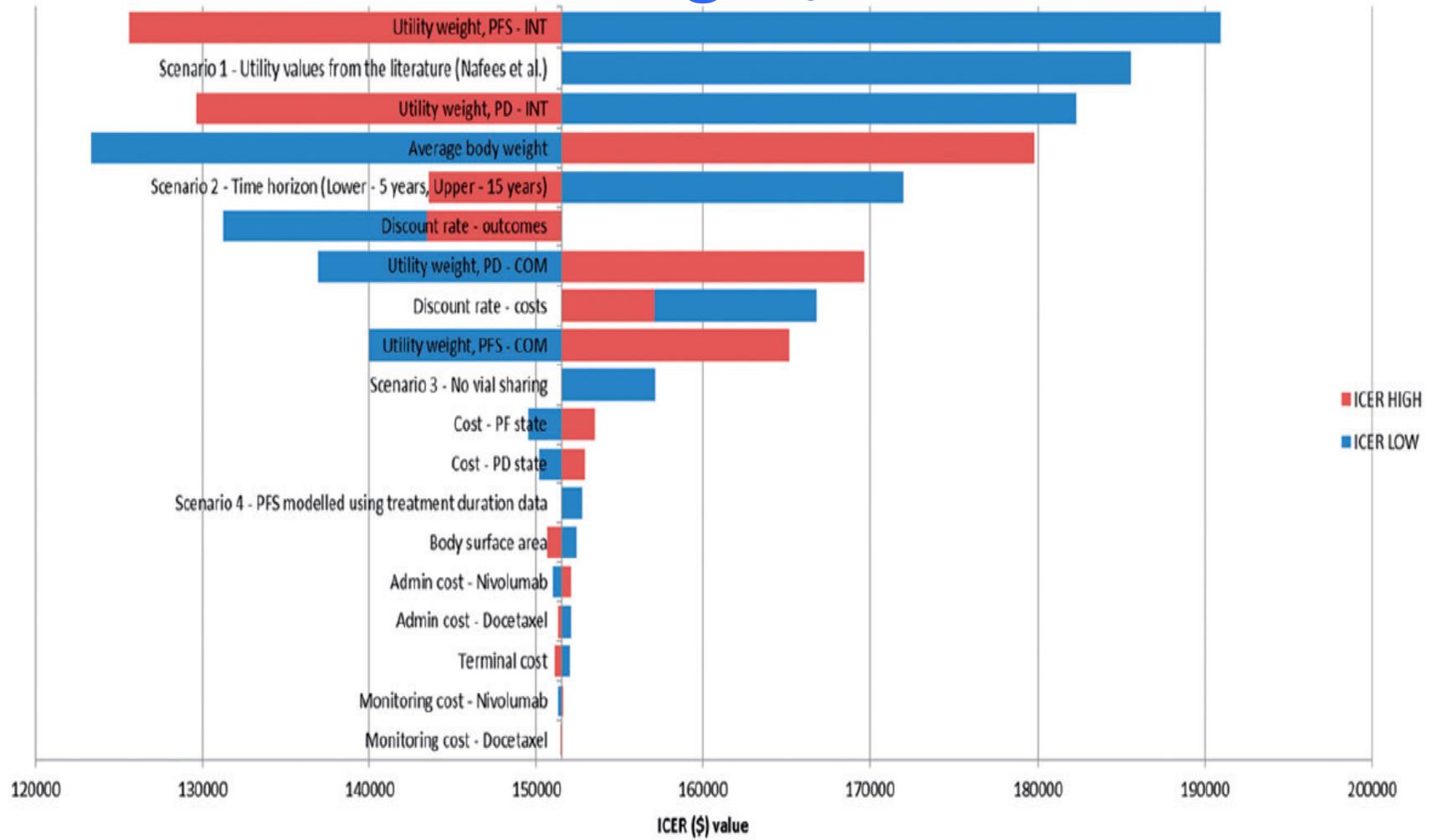


Figure 4. Tomado diagram of one-way sensitivity analyses results for nivolumab compared to docetaxel based on the PS model. COM, comparator; INT, intervention; PD, progressed disease; PF, progression-free; PFS, progression-free survival.

Goree et al, J Med Economics 2017;19:630-44

Nivolumab seconde ligne, US

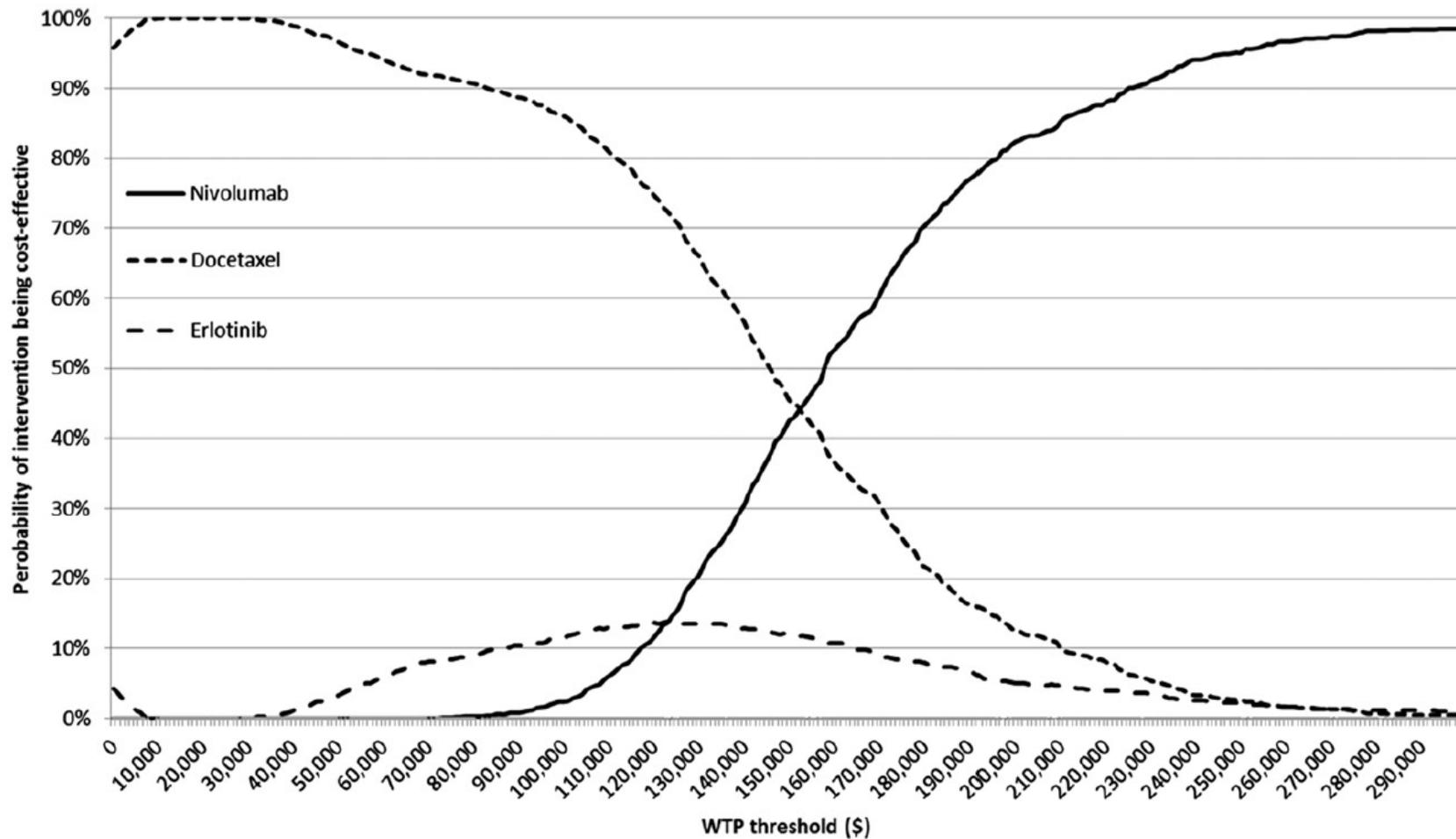


Figure 7. Multiple cost-effectiveness acceptability curves. WTP, willingness-to-pay per quality adjusted life year (QALY) gained.

Goree et al, J Med Economics 2017;19:630-44

Nivolumab seconde ligne, Suisse

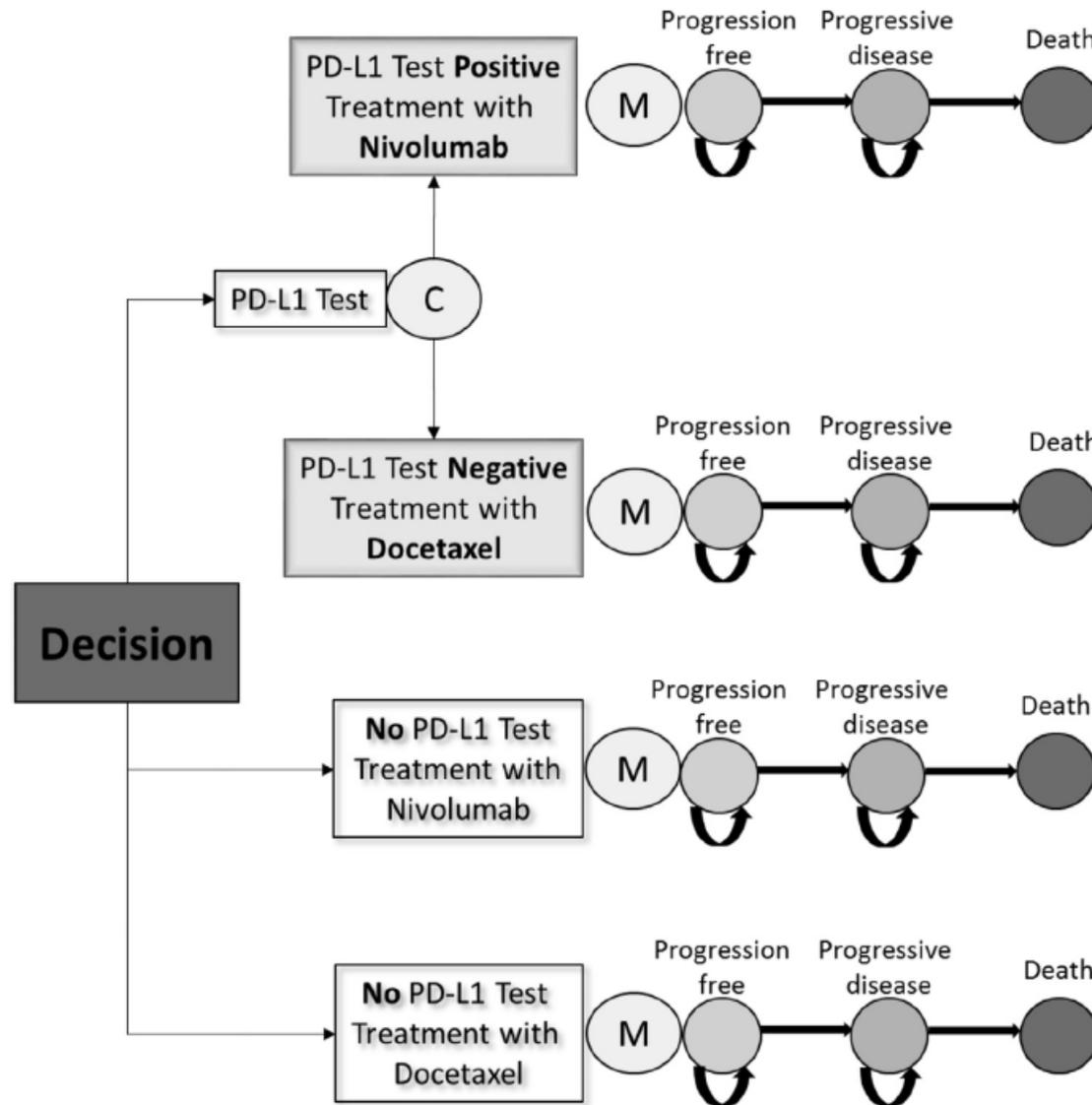


Figure 1. Model structure. PD-L1, programmed death ligand 1; C, chance node; M, Markov node.

Nivolumab seconde ligne, Suisse

Icer 177,478 CHF,
PDL-1>1% 133,267,
PDL-1>10% 124,891

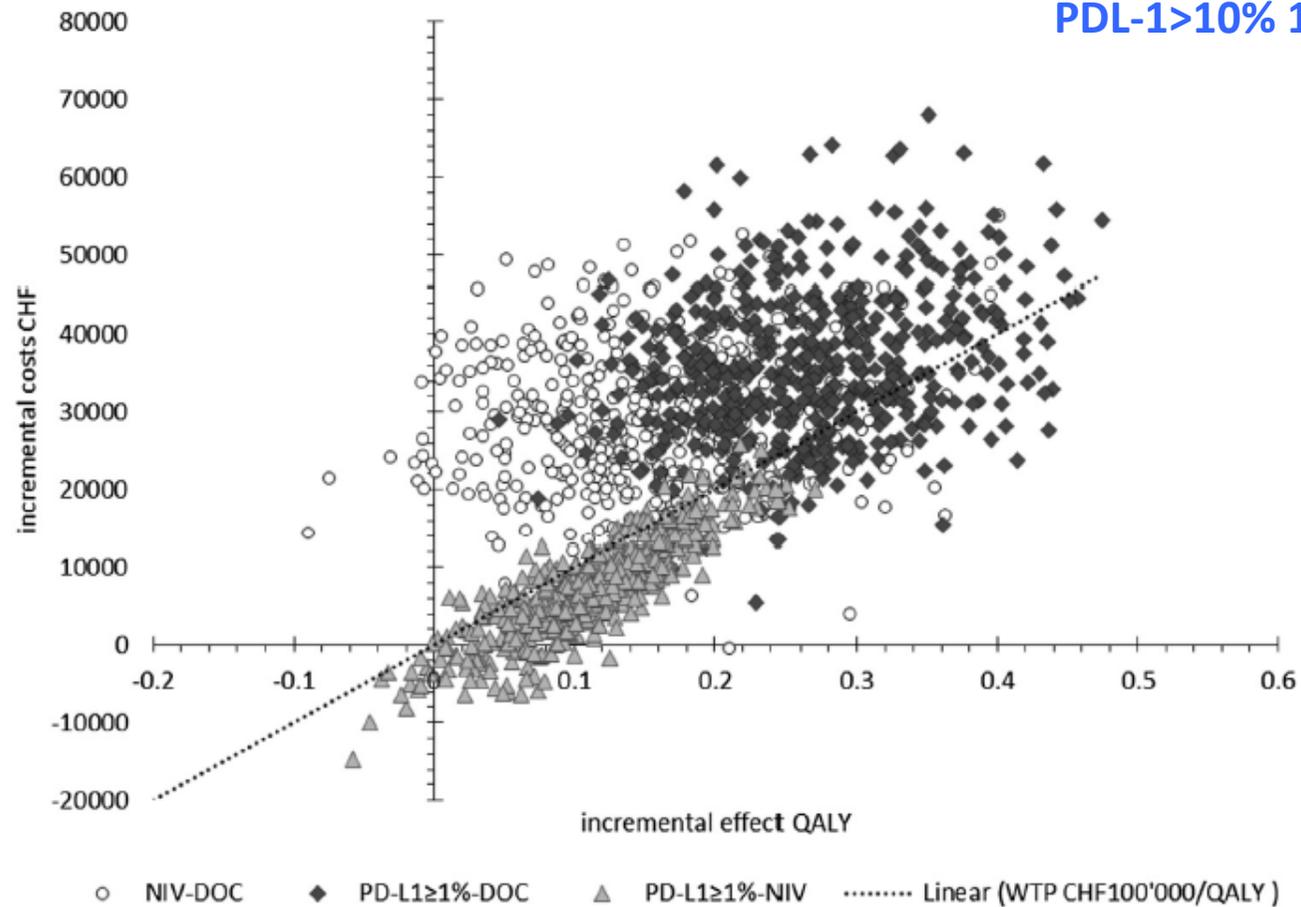


Figure 2. Probabilistic sensitivity analyses for nivolumab (NIV) versus docetaxel (DOC) and by programmed death ligand 1 (PD-L1) testing with a cutoff of 1% or more positive cells (PD-L1 \geq 1%). PD-L1 \geq 1% testing versus docetaxel (patients with positive test results receive nivolumab, patients with negative test results receive docetaxel) and PD-L1 \geq 1% test versus nivolumab. QALY, quality-adjusted life-year; WTP, willingness-to-pay threshold.

Matter-Walstra et al., JTO 2016;11,1846-55

Nivolumab seconde ligne, Suisse

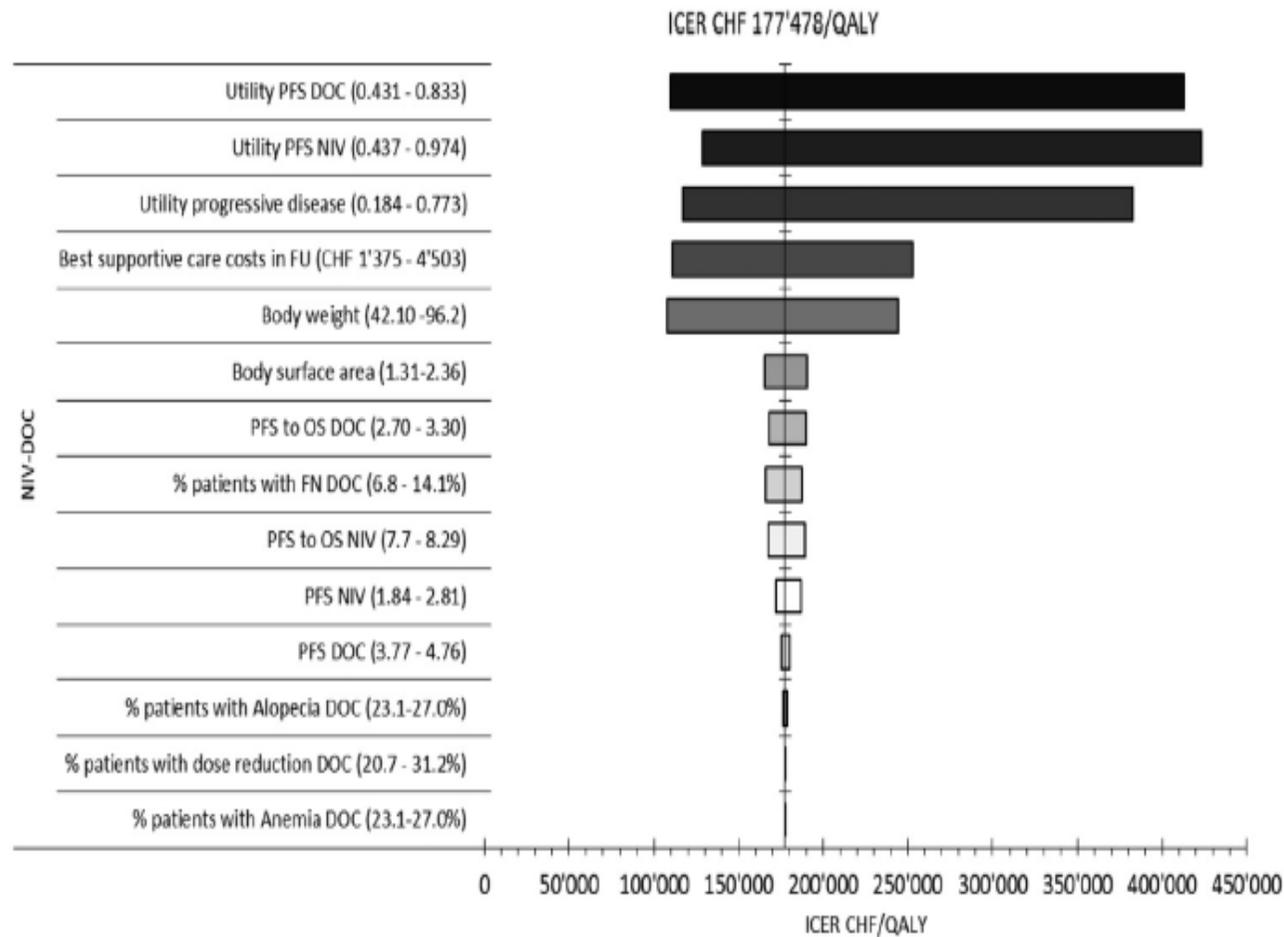


Figure 3. Tornado plot of the univariate sensitivity analyses for nivolumab (NIV) versus docetaxel (DOC). ICER, incremental cost-effectiveness ratio; QALY, quality-adjusted life-year; PFS, progression-free survival; OS, overall survival; FU, follow-up phase; FN, febrile neutropenia.

Matter-Walstra et al., JTO 2016;11,1846-55

Pembrolizumab, Seconde ligne, US

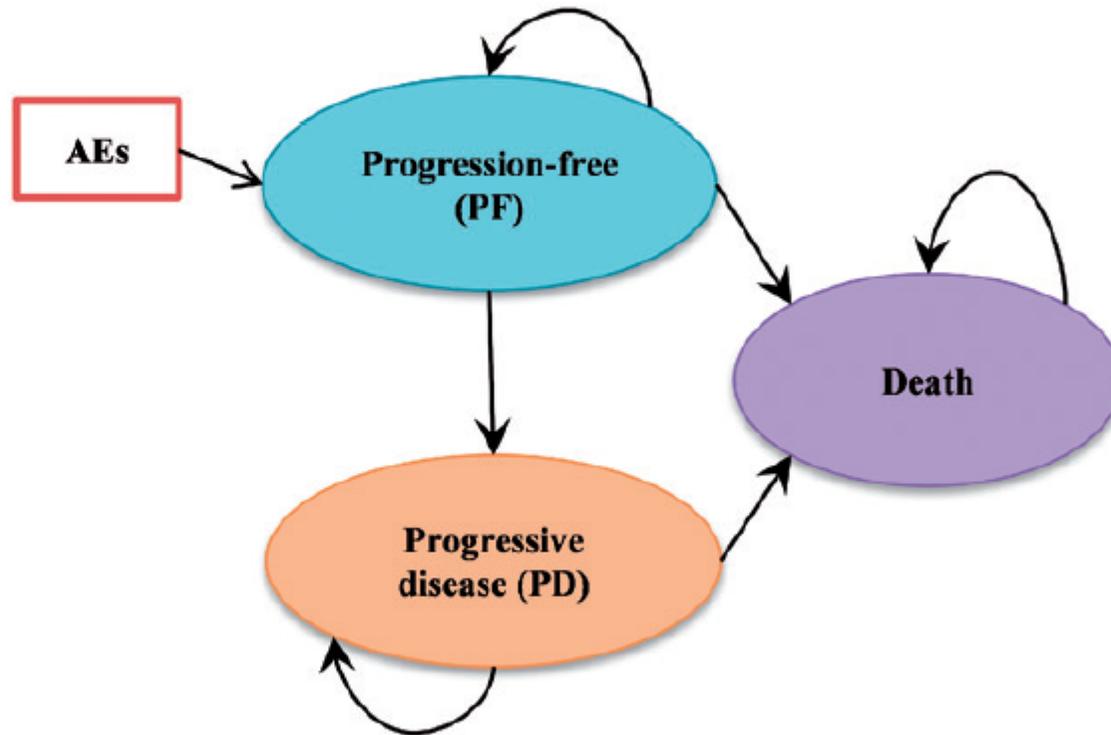


Figure 1. Transition diagram for the cohort simulation model.

Pembrolizumab, Seconde ligne, US

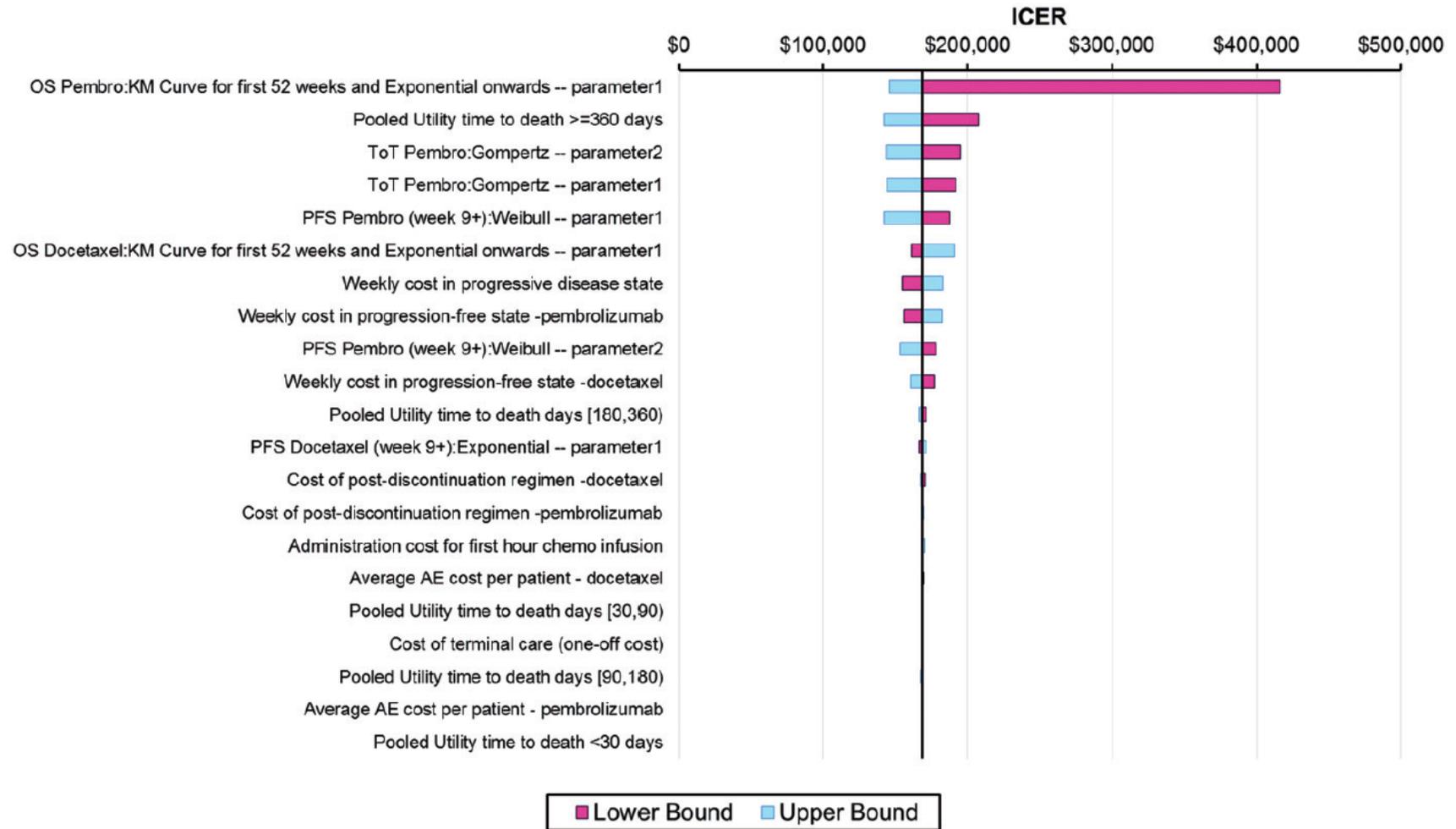


Figure 4. Tornado diagram for one-way sensitivity analyses.

Pembrolizumab, Seconde ligne, US

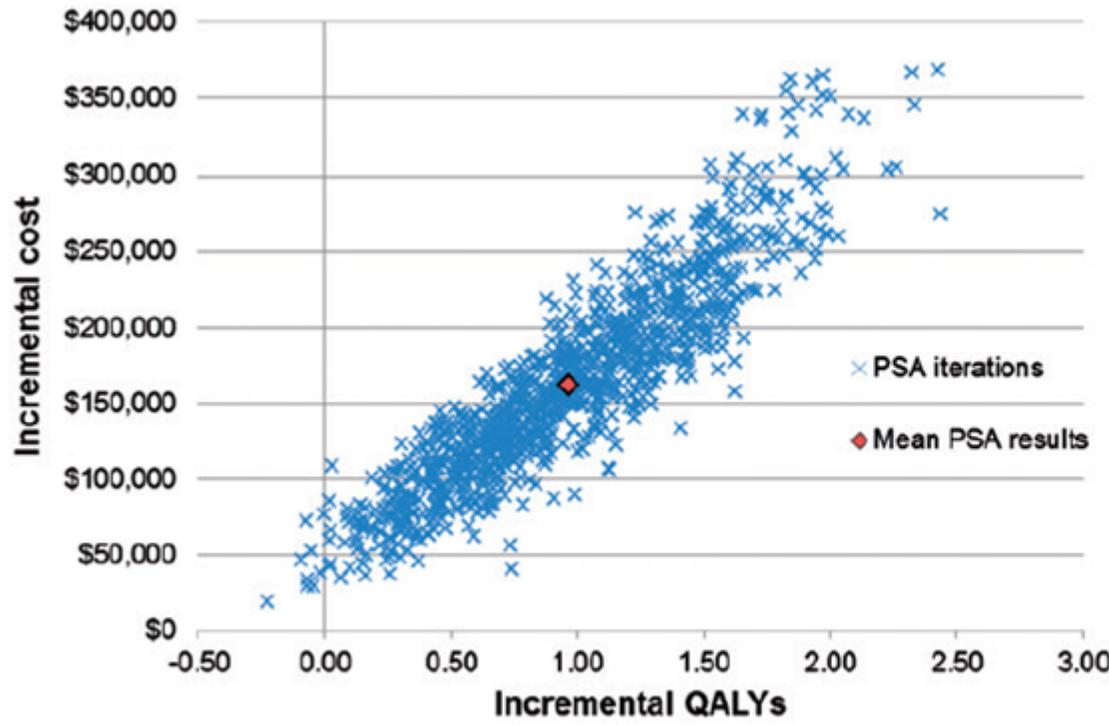


Figure 5. Probabilistic sensitivity analysis scatter plot and cost-effectiveness acceptability curve for base case.

Pembrolizumab, Seconde ligne, US

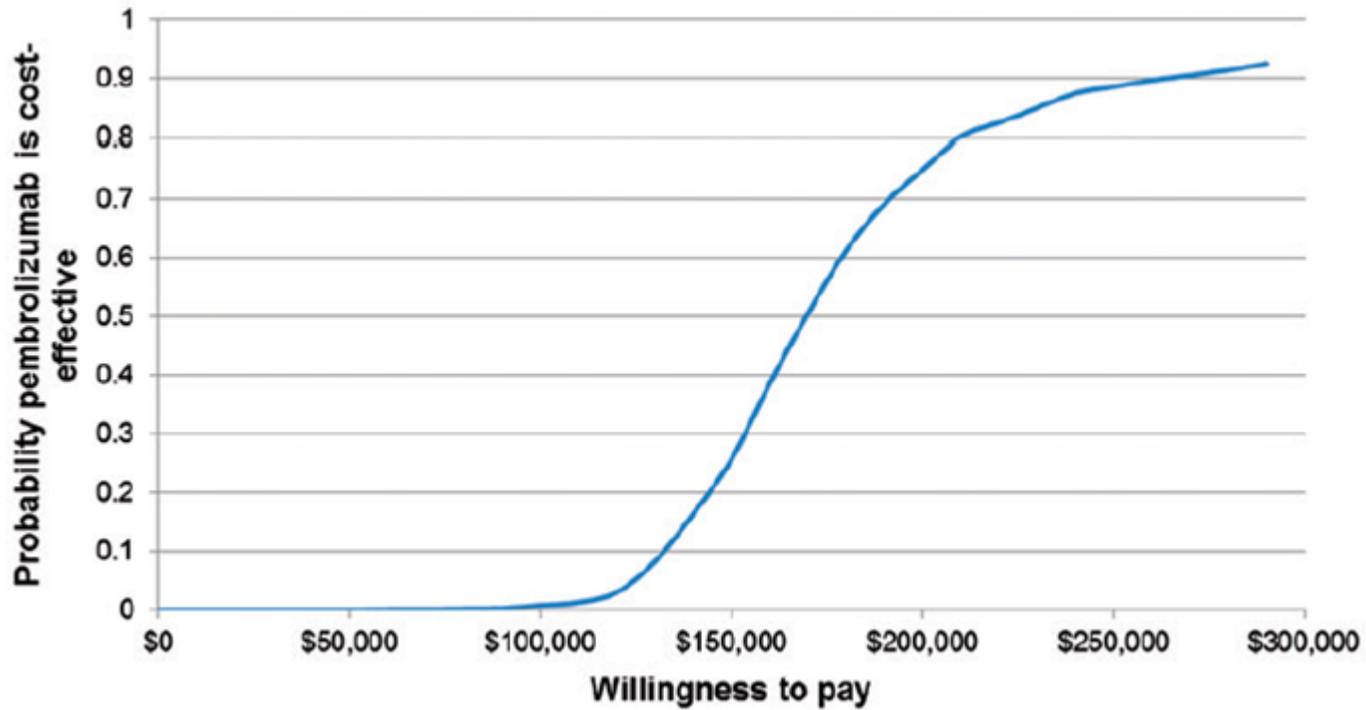
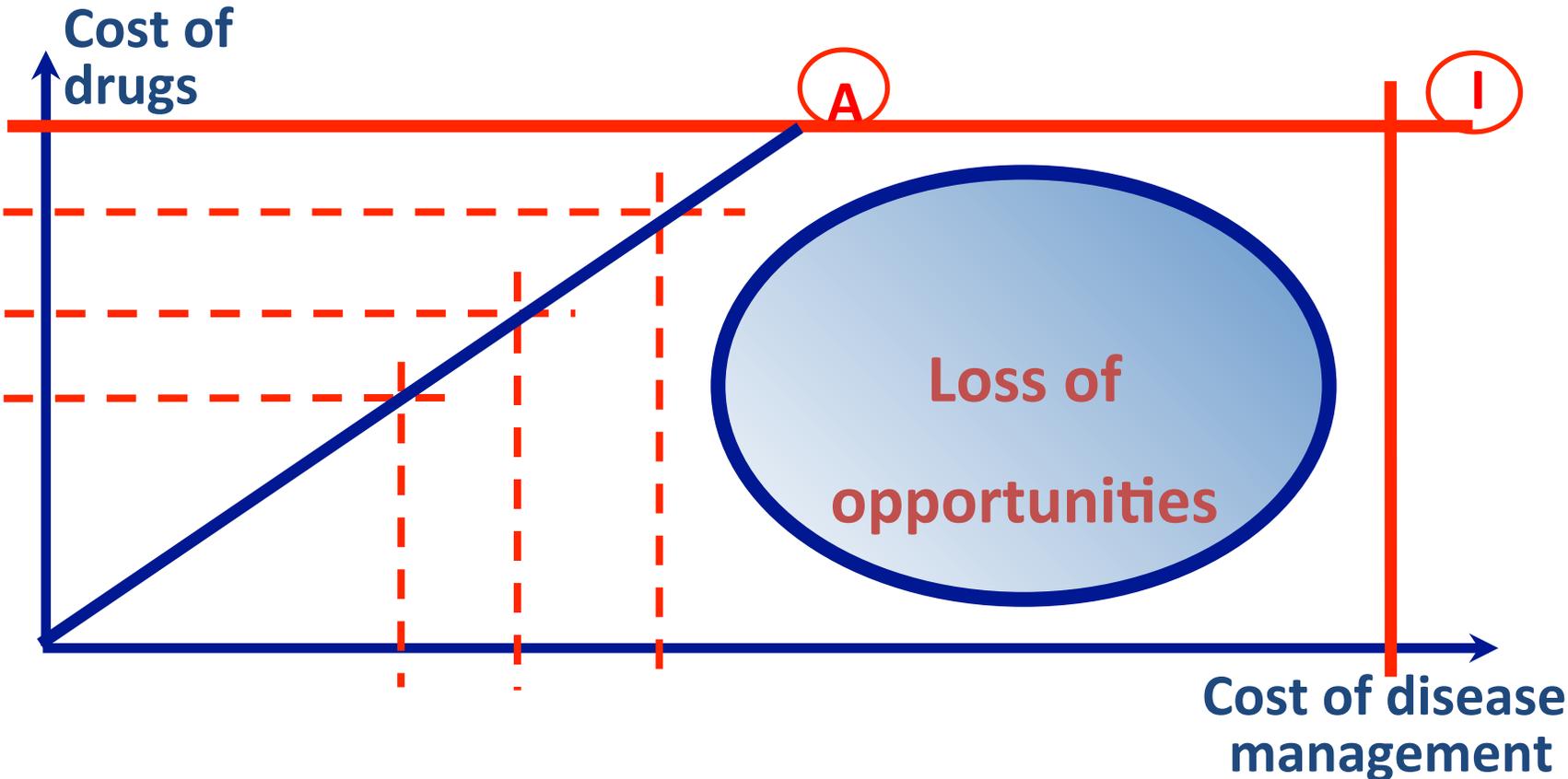


Figure 5. Probabilistic sensitivity analysis scatter plot and cost-effectiveness acceptability curve for base case.

Huang et al, 2017, J Med Economics 2017;20:40-150

Coût du parcours vs coût des drogues



CONCLUSION

- **Coût K US : \$120 billions 2010, \$158 billion 2020**
- **Prix des médicaments 300% en 15 ans \$8,000 à \$10,000 par mois**
- **Aucune corrélation (critères Asco), avec le bénéfice clinique ou les critères du Nice, les Qaly, critères Esmo**
- **Faire des études en France**
- **Analyser les parcours complets, et pas seulement les molécules et leur ligne de traitement**