

# Was gab's Neues im 2015 ? eine Auswahl...



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# Urticaria



**Indications for the Treatment of Chronic Urticaria with Spontaneous Onset**

Chronic urticaria is a common skin condition characterized by recurrent, self-limiting wheals (hives) and/or angioedema. The condition is often associated with an underlying allergic reaction, but in many cases, the cause is unknown (spontaneous onset). Treatment is aimed at relieving symptoms and preventing further episodes.

Indications for treatment include:

- Recurrent wheals and/or angioedema lasting for more than 6 weeks.
- Significant impact on quality of life.
- Failure of self-treatment with over-the-counter antihistamines.

First-line treatment is usually with a second-generation H1-antihistamine. If symptoms persist, higher doses or combination therapy with other medications may be considered.



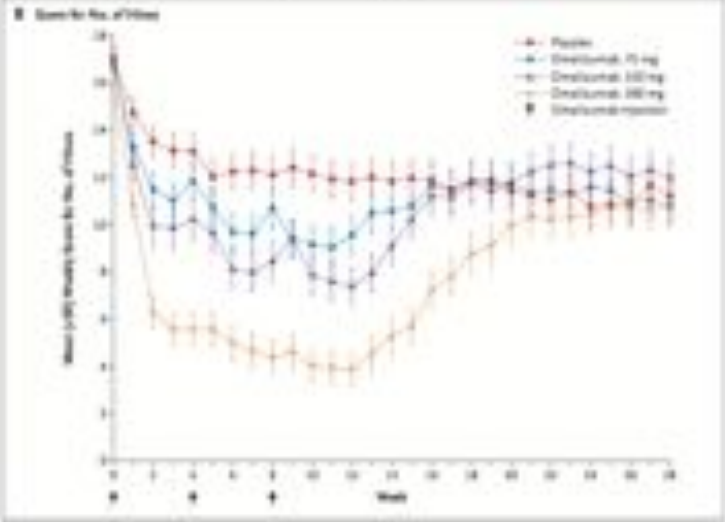
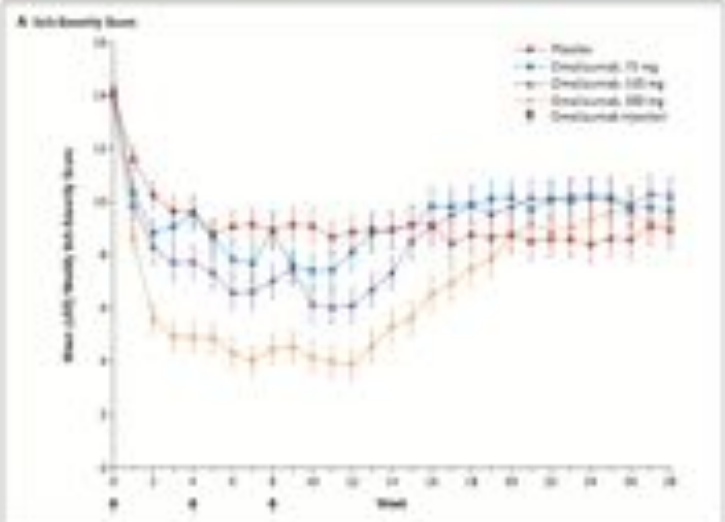
ORIGINAL ARTICLE

## Omalizumab for the Treatment of Chronic Idiopathic or Spontaneous Urticaria

Marcus Maurer, M.D., Karin Rosén, M.D., Ph.D., Hsin-Ju Hsieh, Ph.D.,  
Sarbjit Saini, M.D., Clive Grattan, M.D., Ana Giménez-Arnau, M.D., Ph.D.,  
Sunil Aganwal, M.D., Ramona Doyle, M.D., Janice Carwin, M.D.,  
Allen Kaplan, M.D., and Thomas Casale, M.D.

ABSTRACT







# Psoriasis



New kids on the block

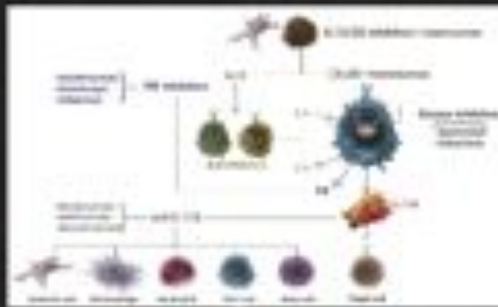
- Apremilast (Otelza)
- Secukinumab (Cosentyx)
- und.....

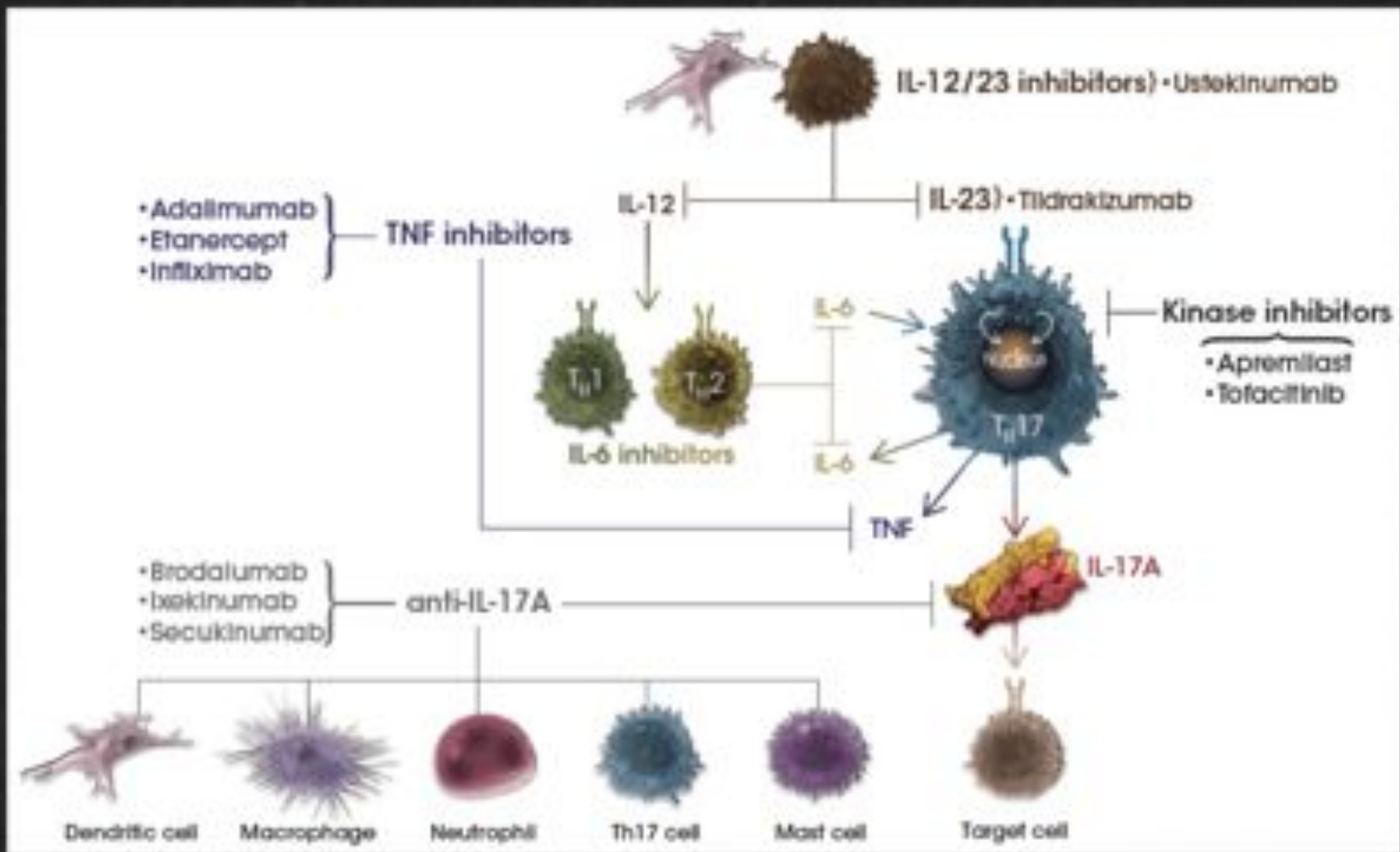


ORIGINAL ARTICLE

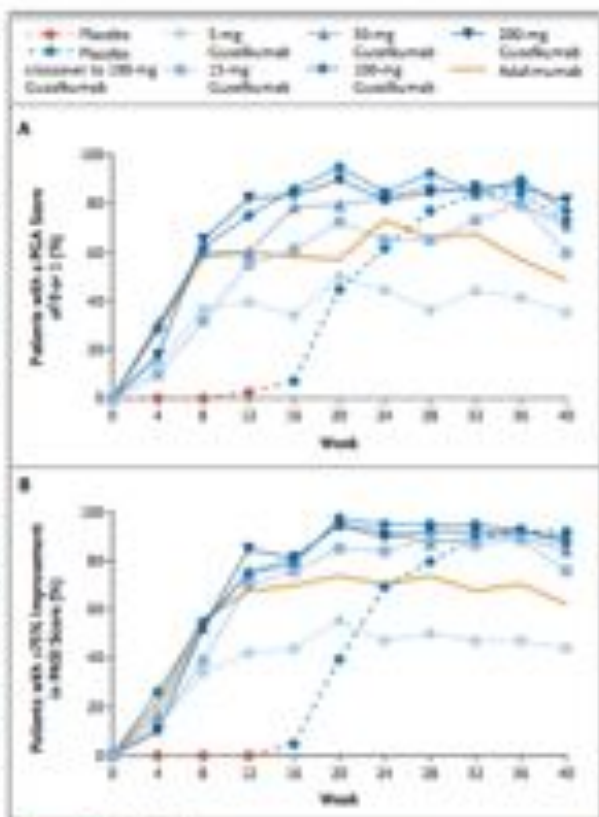
## A Phase 2 Trial of Guselkumab versus Adalimumab for Plaque Psoriasis

Kenneth B. Gordon, M.D., Kristina Callis Duffin, M.D., Robert Bissonnette, M.D., Jörg C. Prinz, M.D., Yasmine Wasfi, M.D., Ph.D., Shu Li, Ph.D., Yaung-Kaung Shen, Ph.D., Philippe Szapary, M.D., M.S.C.E., Bruce Randazzo, M.D., Ph.D., and Kristian Reich, M.D., Ph.D.









**Figure 2. Efficacy Outcomes.**

Panel A shows the proportion of patients with a score on the Physician's Global Assessment (PGA) of 0 (indicating cleared psoriasis) or 1 (indicating minimal psoriasis). Panel B shows the proportion of patients with at least a 75% improvement from baseline in the Psoriasis Area and Severity Index (PASI) score. The 1 mg, 30 mg, and 200 mg guceflumab groups received doses at weeks 0 and 4 and every 12 weeks thereafter, and the 15 mg and 100 mg guceflumab groups received doses every 8 weeks. At week 16, patients in the placebo group crossed over to receive guceflumab at a dose of 200 mg every 8 weeks.

## "meine" Bilder des Jahres



Mein Beruf ist Diagnose | Ich bin Schafshüter

19.06.2015, geschrieben von [Doktor Schmidt](#)



[Bildquelle](#)

[Aktuelle Infos](#)



Ein Zeichen, die in bestimmten Antriebsmodellen der Welt verbreitet werden

# Ein nicht alltägliches türkisches Souvenir

Mehmet Gültekin, Mehmet Eğin, Nakiye Söğüt, Mustafa Laleli, Fatma Gültekin, Nurettin...



Abbildung 2: Verlauf der skleralen Läsion bei einem Patienten über zwei Tage nach Therapiebeginn





**Figure 1. Thoracic, Axillary/Brachial Lymph  
node Swelling/lymphadenitis**



**Figure 2. Subclavicular Lymphadenitis Example  
in the Distribution of a Rash**

A bilateral, lateral edge of parapsoriasis (rash) continues  
— up to axilla or the base of the neck ascending into  
the region of thoracic lymphadenitis (L1 over) (L)

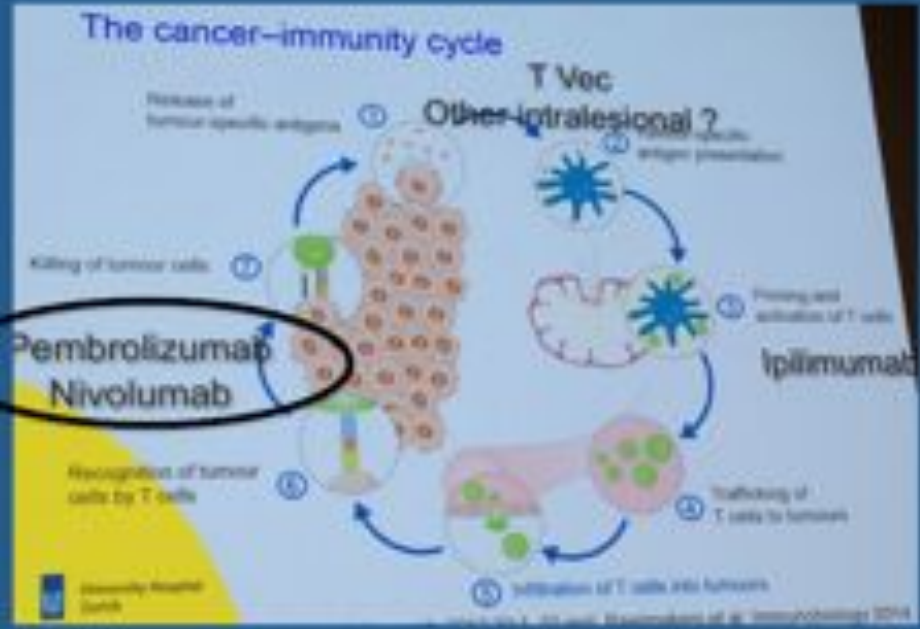






# Melanom - Jahr des PD-1s

## Melanom - Immunologische Therapieansätze



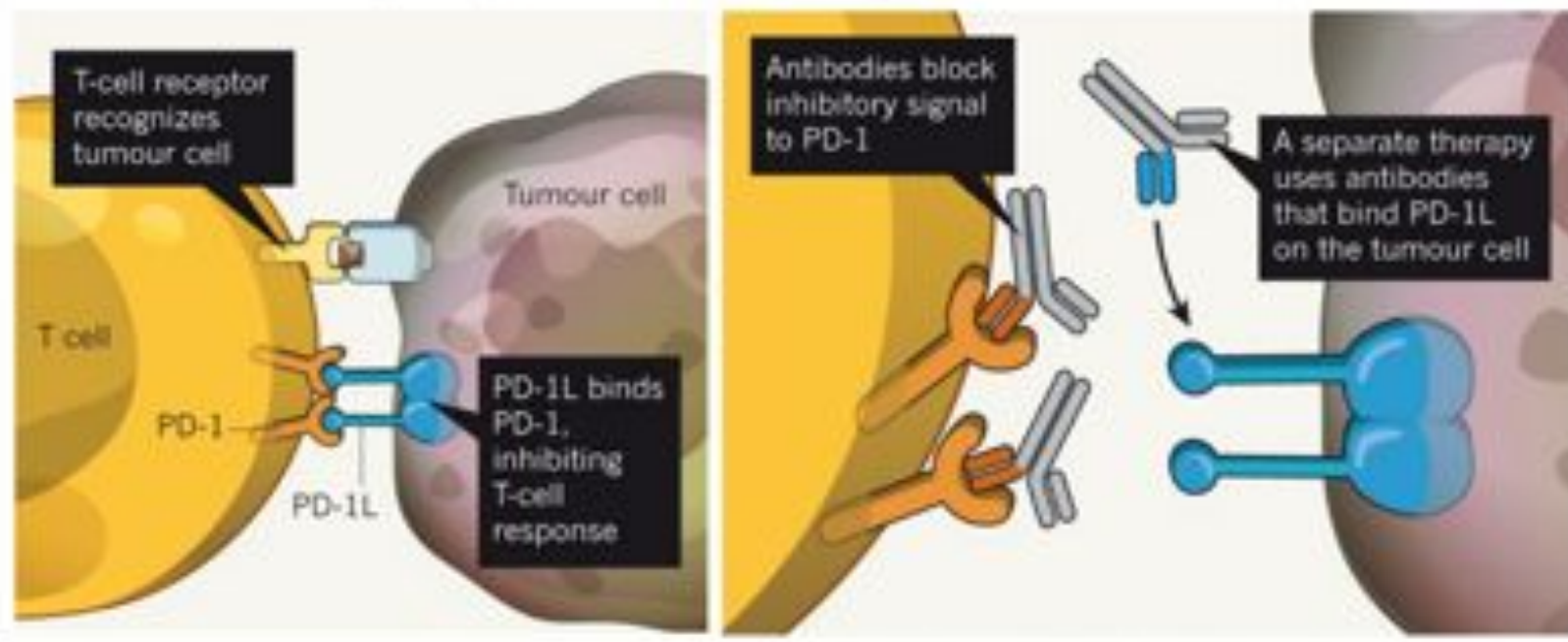
## Programmed cell death 1



- Programmed cell death protein 1, also known as PD-1 and CD279
- PD-1 is a cell surface receptor that belongs to the immunoglobulin superfamily and is expressed on T cells and pro-B cells.
- PD-1 binds two ligands, PD-L1 and PD-L2.
- PD-1, functioning down regulates the immune system
- Prevents the activation of T-cells, which in turn reduces autoimmunity and promotes self-tolerance.

## WAKING UP THE BODY'S DEFENCES

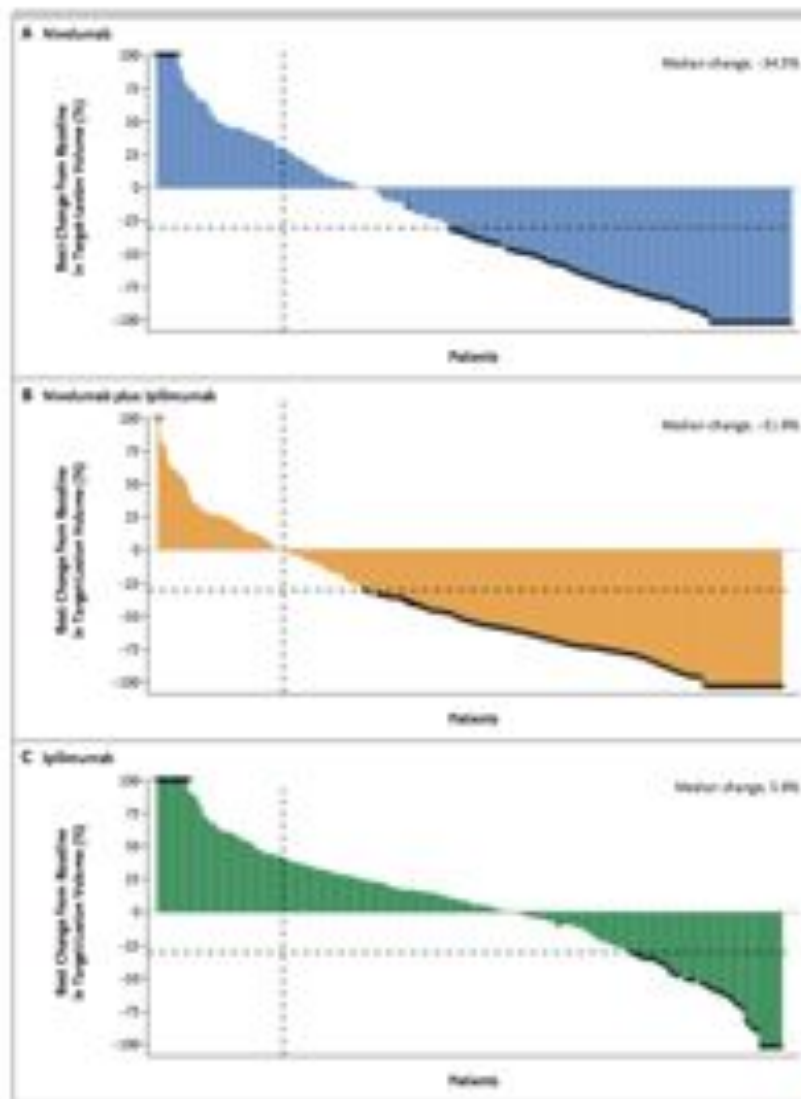
Tumour cells can inhibit the body's immune response by binding to proteins, such as PD-1, on the surface of T cells. Antibody therapies that block this binding reactivate the immune response.





## Combined Nivolumab and Ipilimumab or Monotherapy in Untreated Melanoma

J. Larkin, V. Chiarion-Sileni, R. Gonzalez, J.J. Grob, C.L. Cowey, C.D. Lao, D. Schadendorf, R. Dummer, M. Smylie, P. Rutkowski, P.F. Ferrucci, A. Hill, J. Wagstaff, M.S. Carlino, J.B. Haanen, W. Maio, I. Marquez-Rodas, G.A. McArthur, P.A. Ascierto, G.V. Long, M.K. Callahan, M.A. Postow, K. Grossmann, M. Szniol, B. Dreno, L. Bastholt, A. Yang, L.M. Rollin, C. Horiak, F.S. Hodi, and J.D. Wolchok





### Rapid Eradication of a Bulky Melanoma Mass with One Dose of Immunotherapy

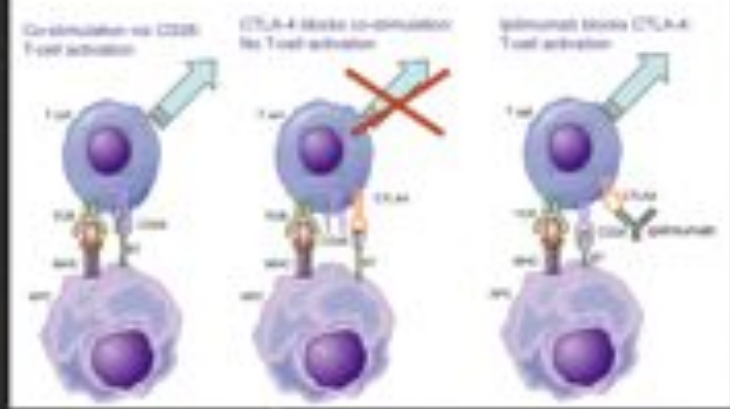
**TO THE EDITOR:** Both ipilimumab<sup>1</sup> (anti-CTLA-4) and nivolumab<sup>2</sup> (anti-PD-1) are immunotherapies that have shown promising results in the treatment of melanoma. We report the rapid eradication of a bulky melanoma mass with one dose of ipilimumab.



**YERVOY<sup>®</sup>**  
(ipilimumab)  
Injection for intravenous infusion

**INDICATION**  
YERVOY is indicated for the treatment of unresectable or metastatic melanoma. The indication for YERVOY includes both previously treated and treatment-naïve patients.

### Ipilimumab Blocks Negative Signaling From CTLA-4



ORIGINAL ARTICLE

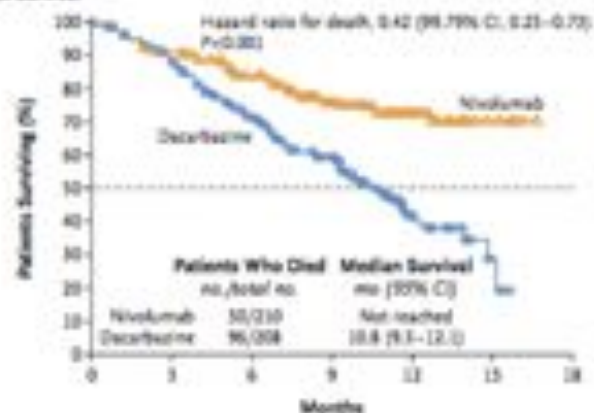
## Nivolumab in Previously Untreated Melanoma without BRAF Mutation

Caroline Robert, M.D., Ph.D., Georgina V. Long, M.D., Ph.D., Benjamin Brady, M.D., Caroline Durrain, M.D., Michele Maio, M.D., Laurent Mortier, M.D., Jessica C. Hassel, M.D., Piotr Rutkowski, M.D., Ph.D., Carina Mitchell, M.D., Ph.D., Eva Kalinka-Warocha, M.D., Ph.D., Kerry J. Savage, M.D., Mikaela M. Hemborg, M.D., Ph.D., Celeste LaSota, M.D., Ph.D., Julie Charles, M.D., Ph.D., Catalin Mihaleciuc, M.D., Vanna Chiarion-Sileni, M.D., Cornelia Mauch, M.D., Ph.D., Francesco Cognetti, M.D., Ana Arance, M.D., Ph.D., Henrik Schmidt, M.D., D.M.Sc., Dirk Schadendorf, M.D., Helen Cogswell, M.D., Lotta Lundgren-Eriksson, M.D., Christine Horak, Ph.D., Brian Sharkey, Ph.D., Ian M. Waxman, M.D., Victoria Atkinson, M.D., and Paolo A. Ascierto, M.D.

ABSTRACT

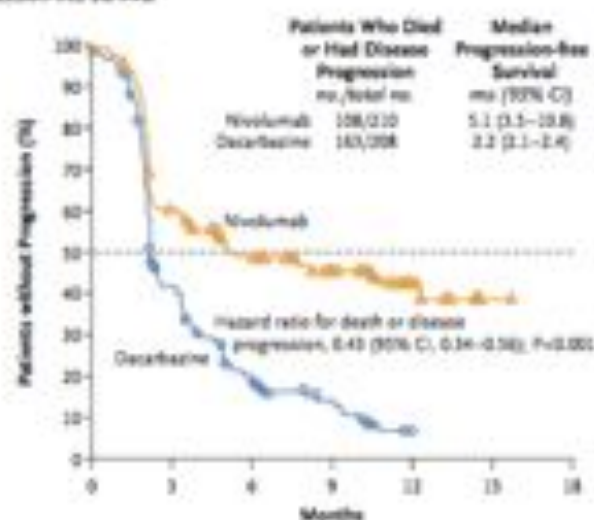


### A. Overall Survival



No. at Risk	0	3	6	9	12	15	18
Nivolumab	210	183	150	105	43	8	0
Docetaxine	208	177	125	62	21	1	0

### B. Progression-Free Survival

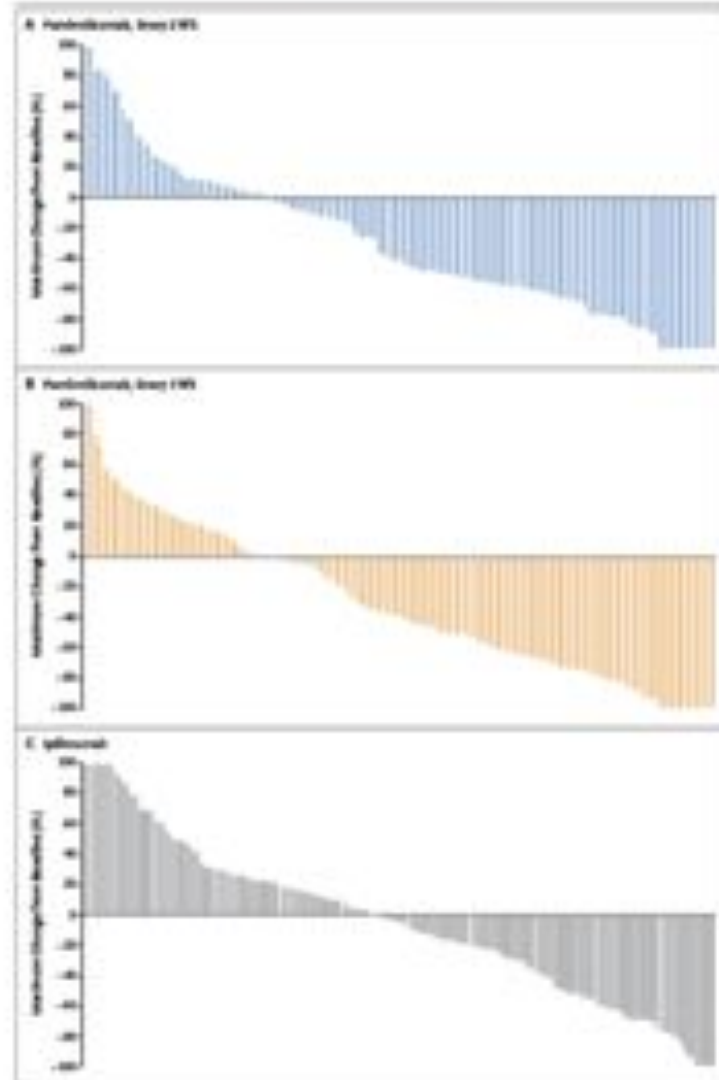


ORIGINAL ARTICLE

## Pembrolizumab versus Ipilimumab in Advanced Melanoma

Caroline Robert, M.D., Ph.D., Jacob Schachner, M.D., Georgina V. Long, M.D., Ph.D., Ana Arance, M.D., Ph.D., Jean-Jacques Grob, M.D., Ph.D., Laurent Mortier, M.D., Ph.D., Adil Daud, M.D., Matteo S. Carlino, M.B., B.S., Catriona McNeil, M.D., Ph.D., Michael Larson, M.D., James Larkin, M.D., Ph.D., Paul Lorigan, M.D., Bart Neyns, M.D., Ph.D., Christian U. Blank, M.D., Ph.D., Omid Hamid, M.D., Christine Mattus, M.D., Ronnie Shapira-Frommer, M.D., Michele Kosh, R.N., B.S.N., Honghong Zhou, Ph.D., Nagesh Ibrahim, M.D., Scot Ebbinghaus, M.D., and Antoni Ribas, M.D., Ph.D., for the KEYNOTE-006 Investigators\*

ABSTRACT



# PD-1s für andere Tumore

## *The* NEW ENGLAND JOURNAL *of* MEDICINE

ESTABLISHED IN 1812

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VOL. 371 NO. 4

### **PD-1 Blockade with Nivolumab in Relapsed or Refractory Hodgkin's Lymphoma**

Stephen M. Ansell, M.D., Ph.D., Alexander M. Leshchkin, M.D., Ivan Bernaldo, M.D., Ahmad Halwani, M.D.,  
Emma C. Scott, M.D., Martin Gutierrez, M.D., Stephen J. Schuster, M.D., Mikhael M. Millenson, M.D.,  
Deepika Cetty, M.S., Gordon J. Freeman, Ph.D., Scott J. Rodig, M.D., Ph.D., Bjørn Chapuis, M.D., Ph.D.,  
Azra H. Ligon, Ph.D., Lei Zhu, M.S., Joseph F. Grossi, Ph.D., Se Young Kim, M.D., Ph.D.,  
John M. Timmerman, M.D., Margaret A. Shipp, M.D., and Philippe Armand, M.D., Ph.D.

ABSTRACT



# BRAF Inhibitoren für andere Tumore

ORIGINAL ARTICLE

## Vemurafenib in Multiple Nonmelanoma Cancers with BRAF V600 Mutations

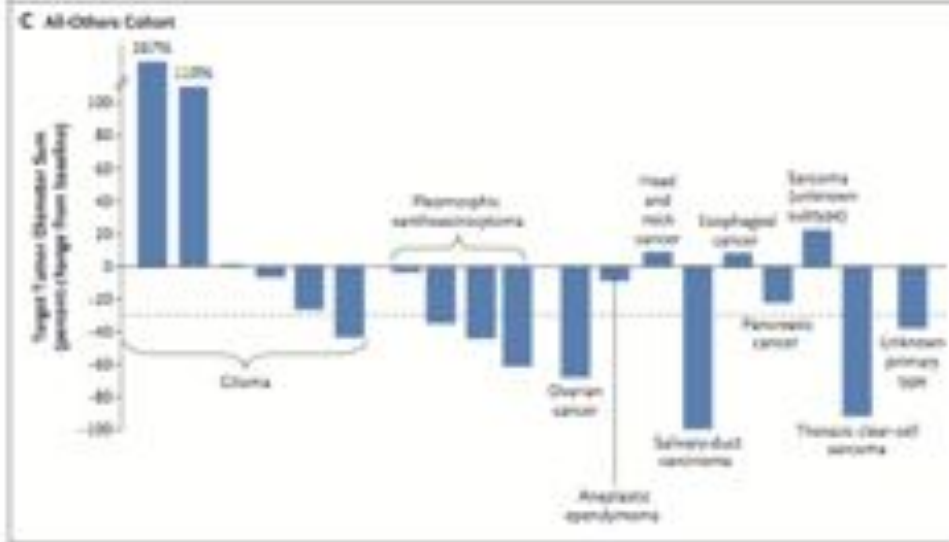
David M. Hyman, M.D., Igor Puzanov, M.D., Vivek Subbiah, M.D., Jason E. Falta, M.D., Ian Chau, M.D., Jean-Yves Blay, M.D., Ph.D., Jürgen Wolf, M.D., Ph.D., Neelam S. Raja, M.D., Eli L. Diamond, M.D., Antoine Hellebecque, M.D., Raji Gonzalez, M.D., Maria Elena Das-Fernandez, M.D., Antoine Italiano, M.D., Ph.D., Raf-Dieser Hoffmeier, M.D., Manuel Hidalgo, M.D., Ph.D., Emily Chan, M.D., Ph.D., Martin Schuler, M.D., Susan Franco Lescotte, M.Sc., Martina Makrutzki, M.D., Florin Siron, M.D., Ph.D., Maria Luisa Veronesi, M.D., Josep Tabernero, M.D., Ph.D., and Joel Baselga, M.D., Ph.D.

Table 1. Overall Survival Results by Population

Population	Median OS (months)	95% CI
All patients	11.5	10.5-12.5
Non-melanoma	11.5	10.5-12.5
Melanoma	11.5	10.5-12.5

Table 2. Overall Response Rate by Population

Population	ORR (%)	95% CI
All patients	48.1	44.1-52.1
Non-melanoma	48.1	44.1-52.1
Melanoma	48.1	44.1-52.1





**Table 1. Recently Approved Agents for Melanoma**

Drug Name	Drug Class	Year Approved
Peginterferon alfa-2b	Immunotherapy	2011
Ipilimumab	Immunotherapy (anti-CTLA-4)	2011
Vemurafenib	BRAF inhibitor	2011
Dabrafenib	BRAF inhibitor	2013
Trametinib	MEK inhibitor	2013
Pembrolizumab	Immunotherapy (PD-1 blocker)	2014

**Table. Efficacy of BRAF Inhibitors**

Regimen	Response Rate, %	Median OS, mo	Median PFS, mo	12-Mo OS, %
Ipilimumab 3 mg/kg q3wk x 4	10.1	10.9	2.8	45.6
Vemurafenib 960 mg bid	48.4	13.6	5.3	56
Dacarbazine 1,000 mg/m <sup>2</sup>	5.4	9.7	1.6	44
Dabrafenib 150 mg bid	50	NR	5.1	NR

BR, twice daily; NR, no response; OS, overall survival; PFS, progression free survival  
NEJM 2011;364:2527-2536; NEJM 2010;362:76-77; Lancet 2012;380(9833):358-365.  
(Courtesy of Larkin-Horns, PhMD)

# Just one more thing.....



## Squamous Change in Basal-Cell Carcinoma with Drug Resistance

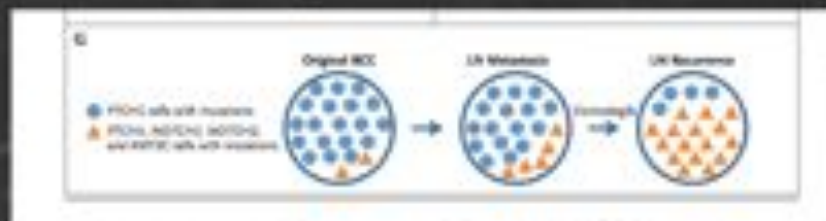
**TO THE EDITOR:** Basal-cell carcinomas are driven by activation of the hedgehog signaling pathway, commonly through mutations in genes encoding patched 1 protein (PTCH1) or smoothened, fil-

lial class receptor (SMO). Vismodegib inhibits SMO and is active in advanced basal-cell carcinoma. However, more than 50% of such lesions develop resistance to vismodegib, commonly through

N Engl J Med 2012;367:11-21. DOI: 10.1056/NEJL1201111

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Appendix, available with the full text of this letter at NEJM.org). Thus, we concluded that cells in a basal-cell carcinoma can switch to squamous cells under vismodegib selection, potentially as a mechanism of tumor escape.



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