

# Clinical trials of immune checkpoint inhibition for melanoma in first line

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## 1 anti-CTLA-4

Trial	Treatments	Patients	Trials design and methods
<b>ipilimumab + dacarbazine vs dacarbazine</b>			
<a href="#">Robert , 2011</a> [NCT00324155] n=NA follow-up:	ipilimumab (10 mg per kilogram) plus dacarbazine (850 mg per square meter of body-surface area) versus dacarbazine (850 mg per square meter)	patients with previously untreated metastatic melanoma (stage III (unresectable) or stage IV)	Parallel groups double blind

## References

### Robert, 2011:

Robert C, Thomas L, Bondarenko I, O'Day S, Weber J, Garbe C, Lebbe C, Baurain JF, Testori A, Grob JJ, Davidson N, Richards J, Maio M, Hauschild A, Miller WH Jr, Gascon P, Lotem M, Harmankaya K, Ibrahim R, Francis S, Chen TT, Humphrey R, Hoos A, Wolchok JD Ipilimumab plus dacarbazine for previously untreated metastatic melanoma. N Engl J Med 2011;364:2517-26 [21639810]

## 2 anti-PD-1

Trial	Treatments	Patients	Trials design and methods
<b>nivolumab vs dacarbazine</b>			
<a href="#">CheckMate 066 (Robert) , 2015</a> [NCT01721772] n=210/208 follow-up:	nivolumab at a dose of 3 mg per kilogram of body weight every 2 weeks versus dacarbazine at a dose of 1000 mg per square meter of body-surface area every 3 weeks	previously untreated patients who had unresectable metastatic melanoma without a BRAF mutation (stage III or IV)	Parallel groups double-blind
<b>nivolumab vs ipilimumab</b>			
<a href="#">CheckMate 067 (nivo vs ipi) , 2015</a> [NCT01844505] n=316/315 follow-up:	3 mg of nivolumab per kilogram of body weight every 2 weeks versus 3 mg of ipilimumab per kilogram every 3 weeks for 4 doses	Previously Untreated Advanced Melanoma	Parallel groups double-blind
<b>nivolumab + ipilimumab vs ipilimumab</b>			

continued...

<b>Trial</b>	<b>Treatments</b>	<b>Patients</b>	<b>Trials design and methods</b>
<b>CheckMate 067 (nivo + ipi vs ipi) , 2015</b> [NCT01844505] n=314/315 follow-up:	1mg of nivolumab per kilogram every 3 weeks plus 3 mg of ipilimumab per kilogram every 3 weeks for 4 doses, followed by 3 mg of nivolumab per kilogram every 2 weeks for cycle 3 and beyond versus 3 mg of ipilimumab per kilogram every 3 weeks for 4 doses	Previously Untreated Advanced Melanoma	Parallel groups double-blind
<b>pembrolizumab (every 2W) vs ipilimumab</b>			
<b>KEYNOTE-006 (every 2W) , 2015</b> [NCT01866319] n=NA follow-up:	pembrolizumab (at a dose of 10 mg per kilogram of body weight) every 2 weeks or every 3 weeks versus four doses of ipilimumab (at 3 mg per kilogram) every 3 weeks	patients with advanced melanoma who had received no more than one previous systemic therapy for advanced disease	Parallel groups open-label
<b>pembrolizumab (every 3W) vs ipilimumab</b>			
<b>KEYNOTE-006 (every 3W) , 2015</b> [NCT01866319] n=277/278 follow-up:	Pembrolizumab Every 3 Weeks versus Ipilimumab (Participants receive ipilimumab, 3 mg/kg IV, once every 3 weeks for a total of Pembrolizumab Every 2 Weeks (Participants receive pembrolizumab, 10 mg intravenously (IV), once every 2 weeks for up to 2 years) 2/ Pembrolizumab Every 3 Weeks (P	patients with unresectable stage III or IV advanced melanoma and who had received no more than one previous systemic therapy for advanced disease	Parallel groups open label
<b>nivolumab + ipilimumab vs nivolumab</b>			
<b>CheckMate 067 (nivo + ipi vs nivo) , 2015</b> [NCT01844505] n=314/316 follow-up:	Nivolumab + ipilimumab versus nivolumab alone	Previously Untreated Advanced Melanoma	Parallel groups double-blind

## References

### CheckMate 066 (Robert), 2015:

Robert C, Long GV, Brady B, Dutriaux C, Maio M, Mortier L, Hassel JC, Rutkowski P, McNeil C, Kalinka-Warzocha E, Savage KJ, Hernberg MM, Lebb C, Charles J, Mihalciou C, Chiarion-Sileni V, Mauch C, Cognetti F, Arance A, Schmidt H, Schadendorf D, Gogas H, Nivolumab in previously untreated melanoma without BRAF mutation. *N Engl J Med* 2015 Jan 22;372:320-30 [25399552] [10.1056/NEJMoa1412082](https://doi.org/10.1056/NEJMoa1412082)

### CheckMate 067 (nivo vs ipi), 2015:

Larkin J, Chiarion-Sileni V, Gonzalez R, Grob JJ, Cowey CL, Lao CD, Schadendorf D, Dummer R, Smylie M, Rutkowski P, Ferrucci PF, Hill A, Wagstaff J, Carlino MS, Haanen JB, Maio M, Marquez-Rodas I, McArthur GA, Ascierto PA, Long GV, Callahan MK, Postow MA, Combined Nivolumab and Ipilimumab or Monotherapy in Untreated Melanoma. *N Engl J Med* 2015;373:23-34 [26027431]

### CheckMate 067 (nivo + ipi vs ipi), 2015:

Larkin J, Chiarion-Sileni V, Gonzalez R, Grob JJ, Cowey CL, Lao CD, Schadendorf D, Dummer R, Smylie M, Rutkowski P, Ferrucci PF, Hill A, Wagstaff J, Carlino MS, Haanen JB, Maio M, Marquez-Rodas I, McArthur GA, Ascierto PA, Long GV, Callahan MK, Postow MA, Combined Nivolumab and Ipilimumab or Monotherapy in Untreated Melanoma. *N Engl J Med* 2015 May 31; [26027431] [10.1056/NEJMoa1504030](https://doi.org/10.1056/NEJMoa1504030)

**KEYNOTE-006 (every 2W), 2015:**

Robert C, Schachter J, Long GV, Arance A, Grob JJ, Mortier L, Daud A, Carlino MS, McNeil C, Lotem M, Larkin J, Lorigan P, Neyns B, Blank CU, Hamid O, Mateus C, Shapira-Frommer R, Kosh M, Zhou H, Ibrahim N, Ebbinghaus S, Ribas A Pembrolizumab versus Ipilimumab in Advanced Melanoma. N Engl J Med 2015 Apr 19;: [25891173] [10.1056/NEJMoa1503093](https://doi.org/10.1056/NEJMoa1503093)

Schachter J, Ribas A, Long GV, Arance A, Grob JJ, Mortier L, Daud A, Carlino MS, McNeil C, Lotem M, Larkin J, Lorigan P, Neyns B, Blank C, Petrella TM, Hamid O, Zhou H, Ebbinghaus S, Ibrahim N, Robert C Pembrolizumab versus ipilimumab for advanced melanoma: final overall survival results of a multicentre, randomised, open-label phase 3 study (KEYNOTE-006). Lancet 2017;: [28822576]

**KEYNOTE-006 (every 3W), 2015:**

Robert C, Schachter J, Long GV, Arance A, Grob JJ, Mortier L, Daud A, Carlino MS, McNeil C, Lotem M, Larkin J, Lorigan P, Neyns B, Blank CU, Hamid O, Mateus C, Shapira-Frommer R, Kosh M, Zhou H, Ibrahim N, Ebbinghaus S, Ribas A Pembrolizumab versus Ipilimumab in Advanced Melanoma. N Engl J Med 2015 Apr 19;: [25891173] [10.1056/NEJMoa1503093](https://doi.org/10.1056/NEJMoa1503093)

Schachter J, Ribas A, Long GV, Arance A, Grob JJ, Mortier L, Daud A, Carlino MS, McNeil C, Lotem M, Larkin J, Lorigan P, Neyns B, Blank C, Petrella TM, Hamid O, Zhou H, Ebbinghaus S, Ibrahim N, Robert C Pembrolizumab versus ipilimumab for advanced melanoma: final overall survival results of a multicentre, randomised, open-label phase 3 study (KEYNOTE-006). Lancet 2017;: [28822576]

**CheckMate 067 (nivo + ipi vs nivo), 2015:**

Larkin J, Chiarion-Sileni V, Gonzalez R, Grob JJ, Cowey CL, Lao CD, Schadendorf D, Dummer R, Smylie M, Rutkowski P, Ferrucci PF, Hill A, Wagstaff J, Carlino MS, Haanen JB, Maio M, Marquez-Rodas I, McArthur GA, Ascierto PA, Long GV, Callahan MK, Postow MA, Combined Nivolumab and Ipilimumab or Monotherapy in Untreated Melanoma. N Engl J Med 2015;373:23-34 [26027431]

### 3 About TrialResults-center.org

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