Neoadjuvant Management of Stage III Melanoma

Joel Baumgartner, MD

Surgical Oncology

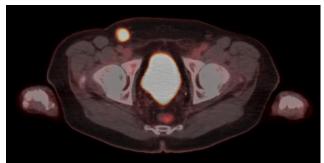
May 8, 2024



Case

- 64 y/o M with 1 yr h/o R flank lesion
- Shave biopsy: 7 mm thick nodular melanoma, + deep margin, + ulceration, 6 MF/mm2
- Had palpable R inguinal LN on PE
- PET/CT with FDG-avid R inguinal LN, no other metastatic disease
- R inguinal LN needle biopsy: metastatic melanoma
- Next steps
 - WLE + R inguinal LN dissection?
 - WLE + isolated R inguinal LN excision?
 - Systemic therapy alone?
 - Systemic therapy followed by WLE, R ILND?
 - Systemic therapy followed by WLE, R inguinal LN excision?

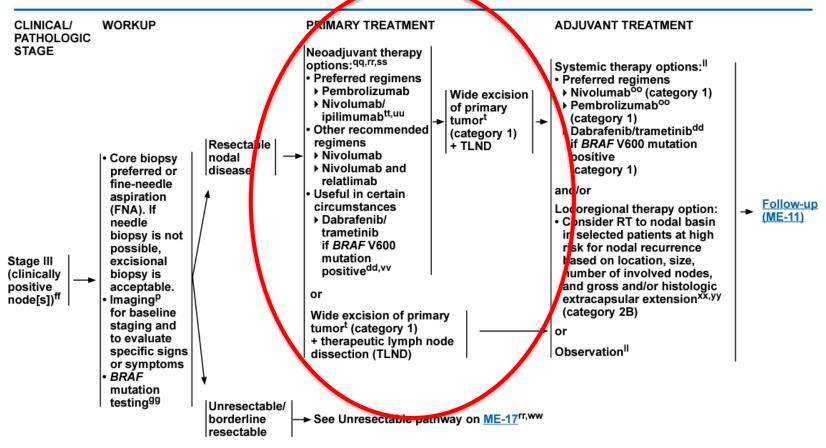




SWOG 1801

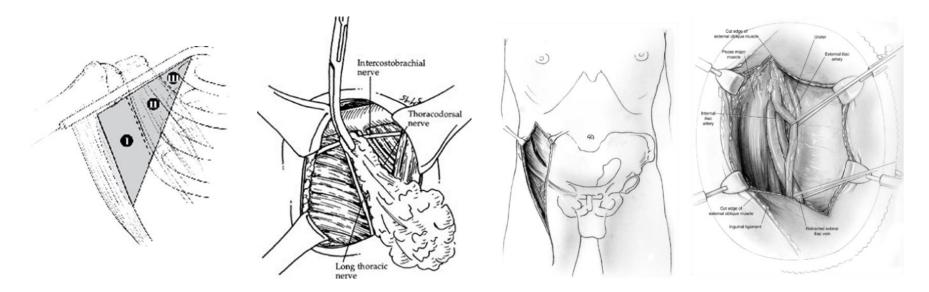


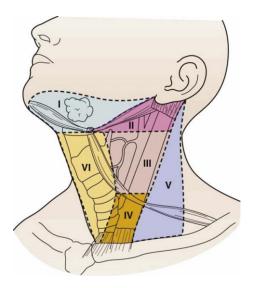
NCCN Guidelines Version 2.2024 Melanoma: Cutaneous NCCN Guidelines Index
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Discussion





Lymph Node Dissection

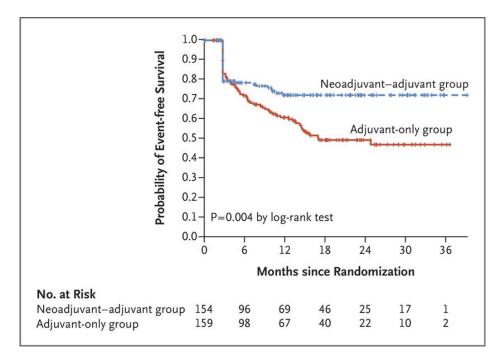






SWOG 1801

- Phase II RCT of stage IIIb-IVc melanoma patients to 3c neoadjuvant pembro + LND + 15c adjuvant pembro v. LND + 18c adjuvant pembro
- Primary endpoint: event-free survival disease progression or toxicity precluding surgery, inability to resect all gross disease, complications/toxicity precluding adjuvant therapy (w/i 84d), recurrence, death



2 yr EFS

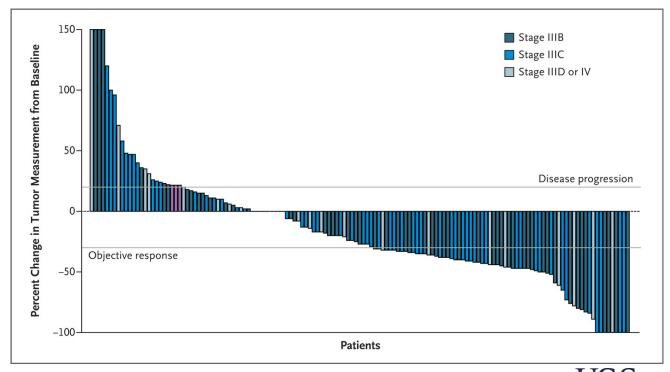
-NA group: 72%

-adjuvant group: 49%



SWOG 1801

- Among patients receiving NA treatment (n=142)
 - 6% radiologic CR, 41% PR
 - 21% pathologic CR (pCR)



OpACIN trials

- Clinical stage III melanoma
- OpACIN: RCT of 20 pts to LND + 4c adjuvant ipi/nivo v. 2c NA ipi/nivo + LND + 2c adjuvant ipi/nivo
- OpACIN-neo: RCT of 86 pts to 2c NA ipi1/nivo3 v. ipi3/nivo1 v. ipi3/nivo3 (sequential), followed by LND
- Pathologic response per INMC criteria
 - pCR: 0% residual viable tumor
 - near-pCR: 0-10% residual viable tumor
 - pPR: 10-50% residual viable tumor
 - pNR: > 50% residual viable tumor

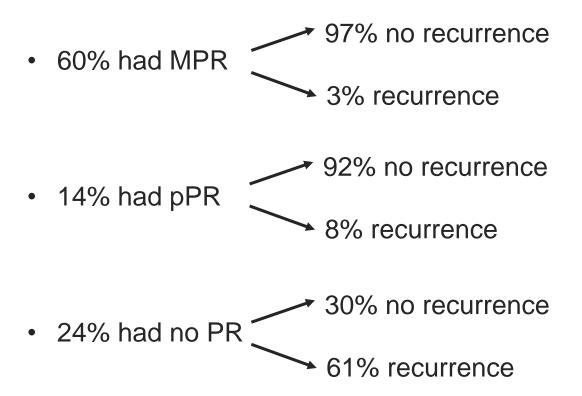


major pathologic

response (MPR)

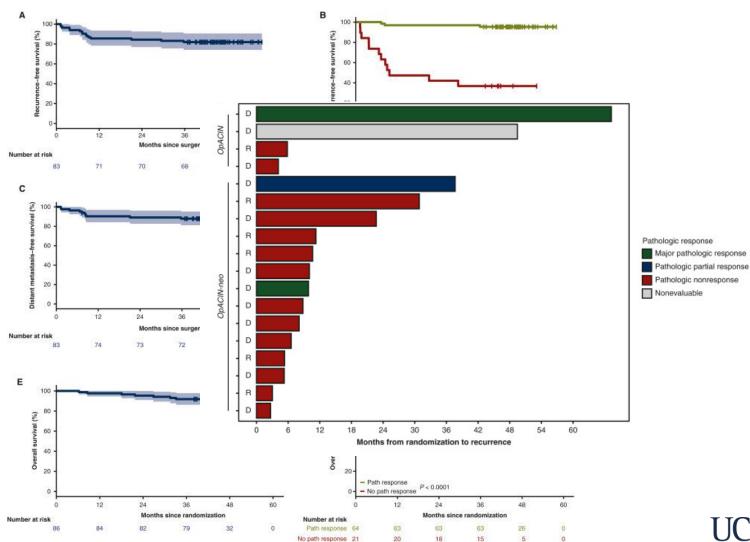
OpACIN trials

Among 96 pts who received NA treatment



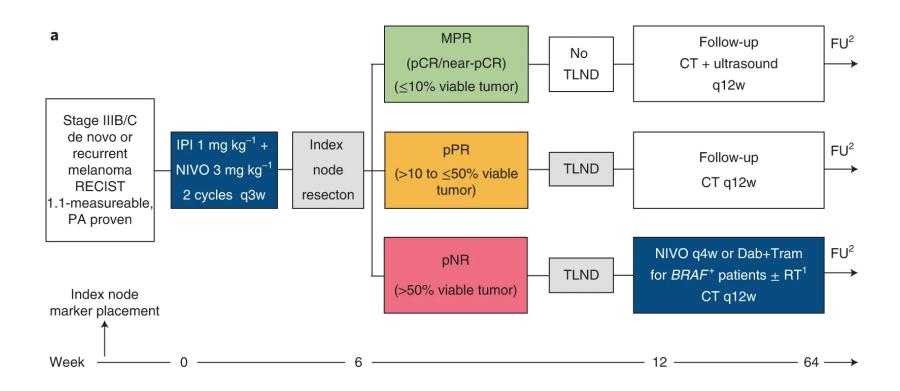


OpACIN trials



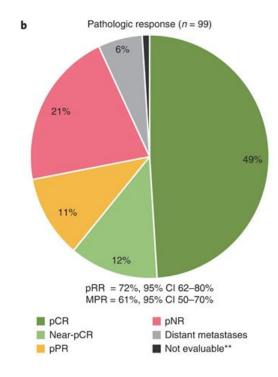
PRADO Trial

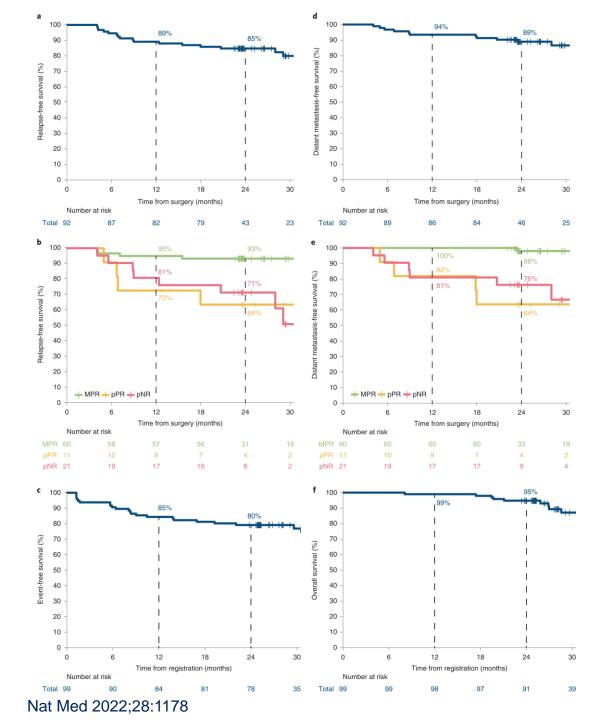
Extension cohort of the OpACIN-neo trial in 99 clinical stage III pts



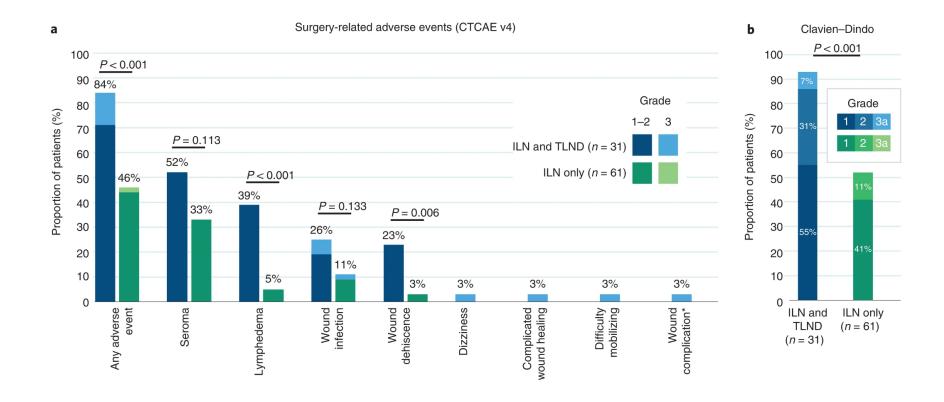


PRADO Trial





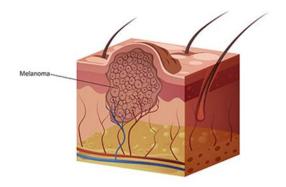
PRADO Trial





Neoadjuvant treatment summary for stage III melanoma

- Improved event-free survival v. adjuvant treatment alone
- Pathologic response is major predictor of outcome
- Pathologic response is greater with ipi/nivo than single agent pembro
- Resection of index LN is feasible, and can direct further surgical and systemic therapy (including de-escalation), and is less morbid than LND
- Unanswered questions
 - Is there a survival benefit with NA treatment?
 - Is tailored LN excision sufficient in those receiving NA treatment?







Thank You!

j1baumgartner@health.ucsd.edu

