



# Neoadjuvant and adjuvant systemic therapy of ovarian cancer

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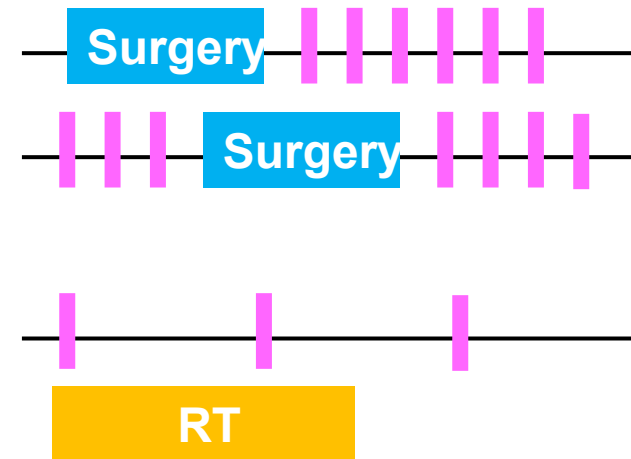
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# DECLARATION OF INTERESTS

Nothing to declare

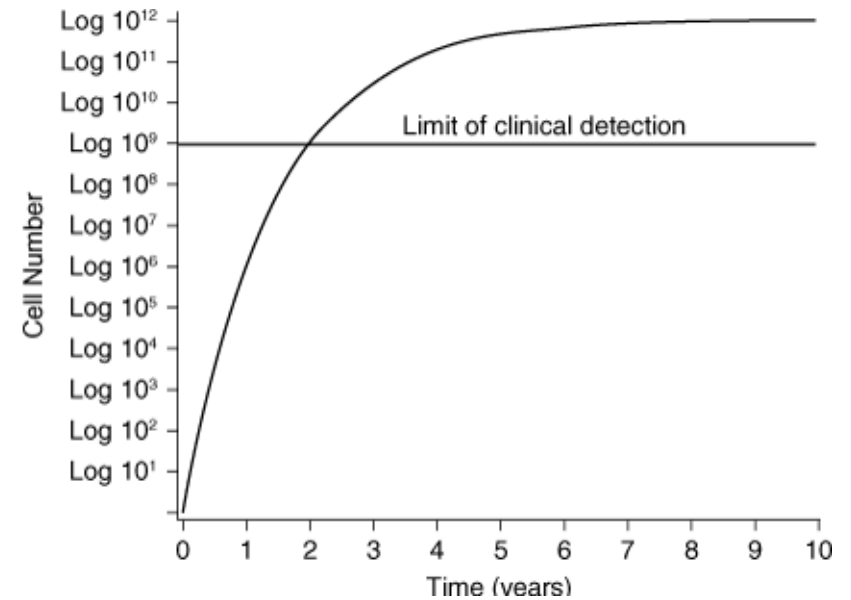
# Principles of chemotherapy

- Surgery (local)
- Radiotherapy (regional)
- Chemotherapy (systemic)
  - Adjuvant chemotherapy
  - Neoadjuvant chemotherapy
  - Palliative/salvage chemotherapy
  - Concurrent chemoradiation



# Principles of chemotherapy

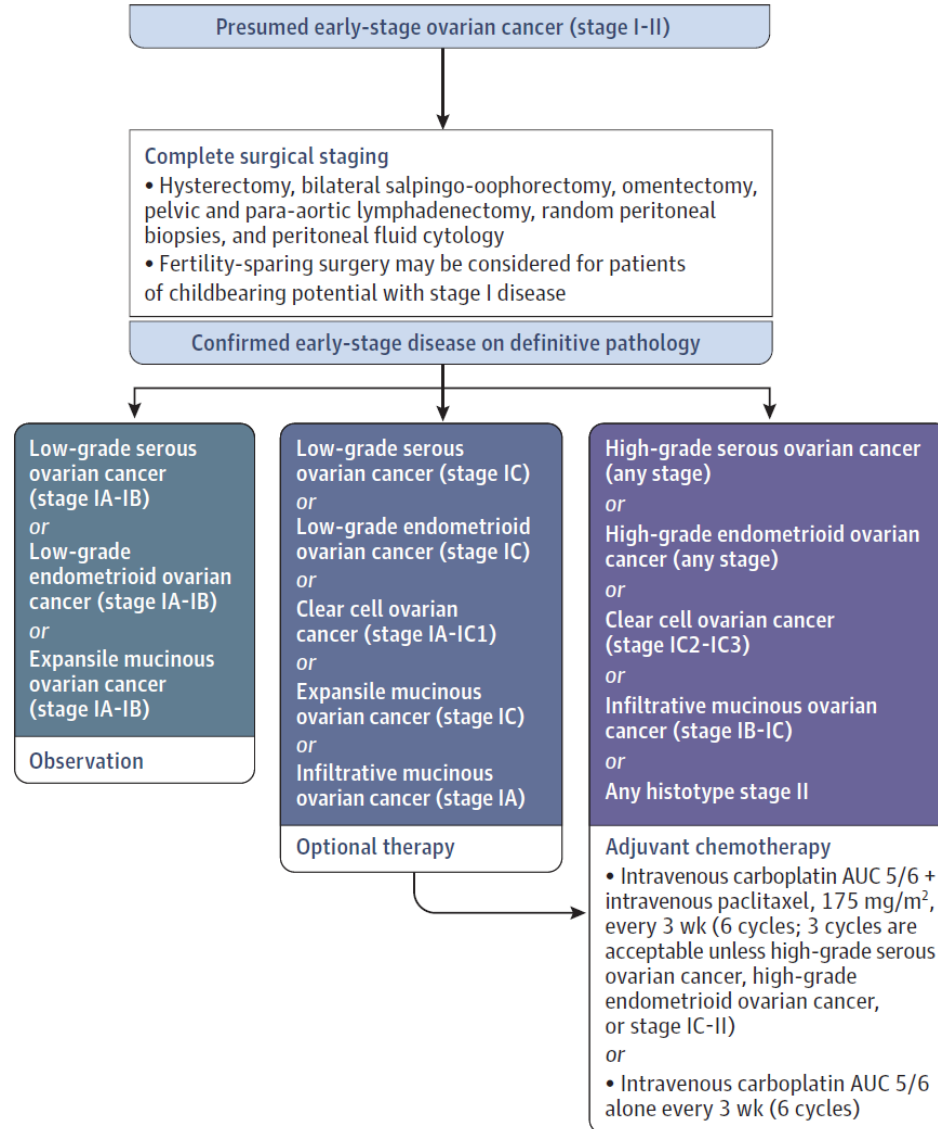
- 가능한 빠른 시기에
  - 암존재량이 적을수록 항암치료의 효과 ↑↑
- 복합항암화학요법
  - 2-3 가지 약제  
(다른 기전 및 부작용 종류)  
→ 순응도↑, 독성↓, 재발↓



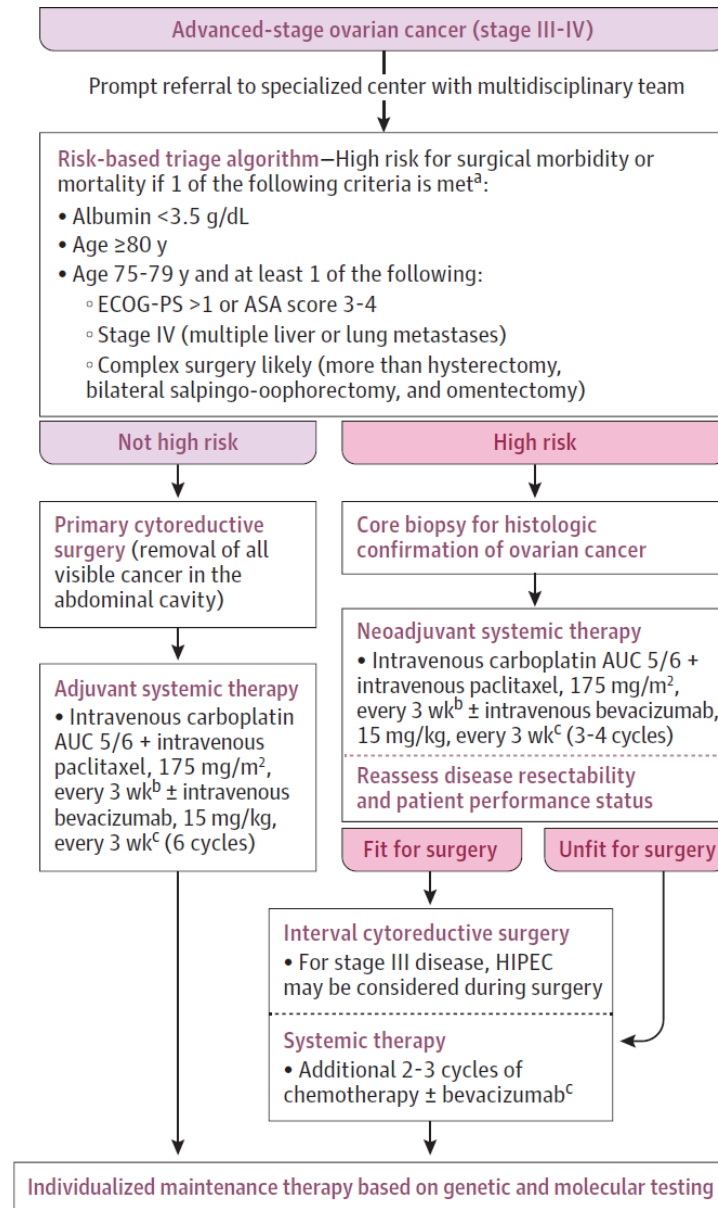
# Principles of chemotherapy

- Administration route
  - **IV: most common**
  - IM: methotrexate
  - PO: etoposide
  - IP (intraperitoneal): cisplatin, paclitaxel
  - IT (intrathecal): methotrexate
- 투약 간격
  - **3주 / 1주** (weekly regimens)
- 용량
  - **BSA (m<sup>2</sup>)**
    - DuBois & DuBois formula:  $BSA = 0.007184 \times weight (kg)^{0.425} \times height (cm)^{0.725}$
    - Mosteller formula:  $\sqrt{\frac{weight (kg) \times height (cm)}{3600}}$

# Primary Treatment for Early-Stage Ovarian Cancer

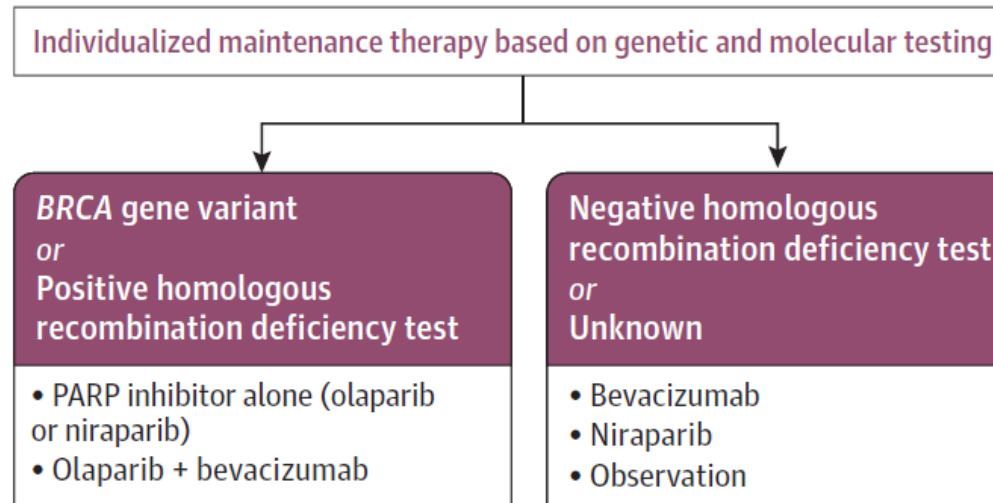


# Primary Treatment for Advanced-Stage EOC

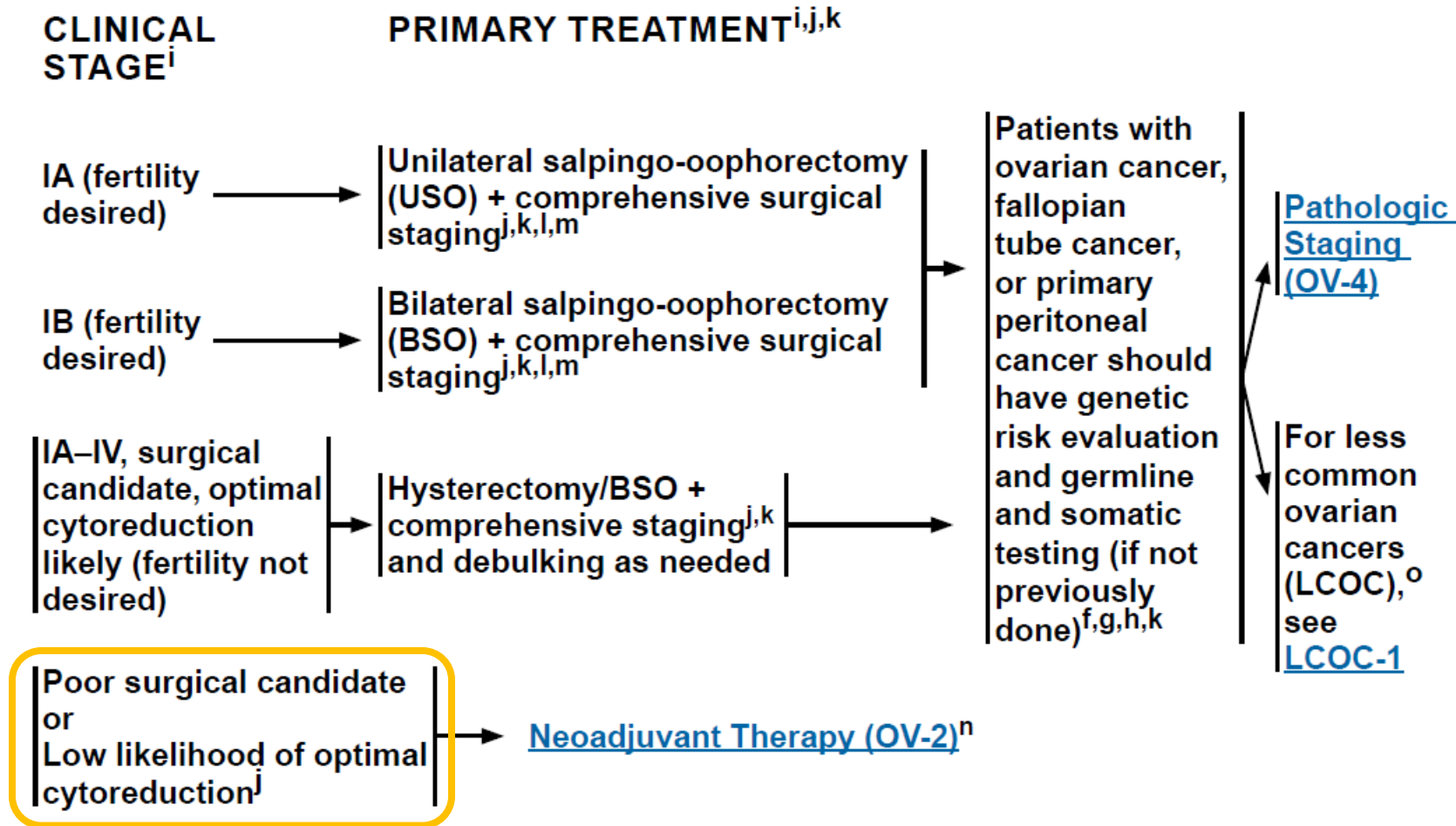


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# Primary Treatment for Advanced-Stage EOC

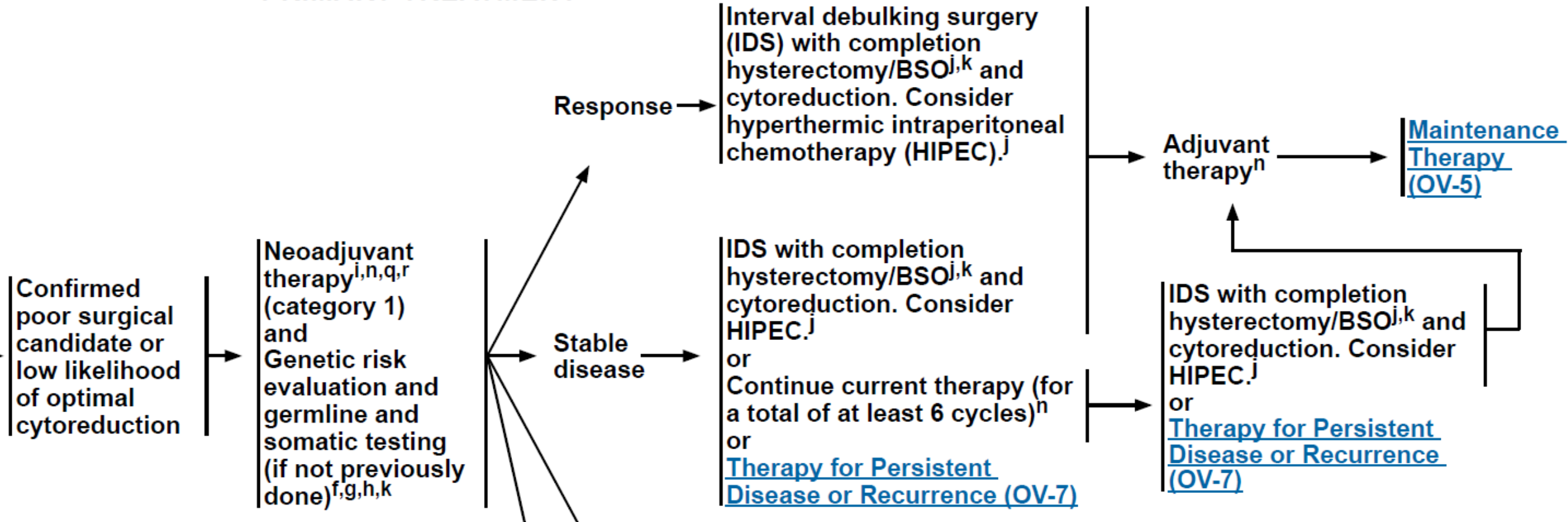


# Primary Treatment for Ovarian Cancer



# Primary Treatment for Ovarian Cancer

## PRIMARY TREATMENT



# Adjuvant CTx for advanced EOC

- Adjuvant therapy: following cancer surgery, to decrease the risk of disease recurrence or to primarily treat residual disease
- Paclitaxel/Carboplatin q 3wks x 6 cycles
- +/- Bevacizumab (stage IV, >1cm residual ds after surgery)
- Starting within 6 weeks after surgery

# Adjuvant CTx for advanced EOC

- IV Paclitaxel 175 mg/m<sup>2</sup> / Carboplatin AUC 5-6 on D1, q 3wks x 6 cycles (preferred)
- Other IV regimens
  - Docetaxel 60-75 mg/m<sup>2</sup> / Carboplatin AUC 5-6 on D1, q 3wks x 6 cycles (SCOTROC1)
  - Weekly paclitaxel 80 mg/m<sup>2</sup> (D1,8,15) / Carboplatin AUC 5-6 on D1, q 3wks x 6 cycles (JGOG-3016)
  - Weekly paclitaxel 60 mg/m<sup>2</sup> / Carboplatin AUC 2 on D1,8,15, q 3wks x 6 cycles (MITO-7, ICON8)
  - Liposomal doxorubicin 30 mg/m<sup>2</sup> / Carboplatin AUC 5 on D1, q 4wks x 6 cycles

# Adjuvant CTx for advanced EOC

## 2. 수술후보조요법 (adjuvant)

| 연번 | 항암요법                                                                                                               | 투여대상                                                                                                 |
|----|--------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------|
| 1  | docetaxel + carboplatin<br>(제2007-7호: 2007.11.20, 개정 제2022-190호: 2022.8.1.)                                        | 가. stage IA (Grade 2, 3)<br>나. stage IB (Grade 2, 3)<br>다. stage IC 이상                               |
| 2  | docetaxel + cisplatin<br>(제2006-3호: 2006.1.9, 개정 제2007-7호: 2007.11.20., 개정 제2022-190호: 2022.8.1.)                  |                                                                                                      |
| 3  | paclitaxel + cisplatin<br>(제2006-3호: 2006.1.9, 개정 제2007-7호: 2007.11.20., 개정 제2022-190호: 2022.8.1.)                 |                                                                                                      |
| 4  | paclitaxel + carboplatin<br>(제2007-7호: 2007.11.20, 개정 제2015-291호: 2015.12.1, 개정 제2022-190호: 2022.8.1.)             | 가. stage IA (Grade 2, 3)<br>나. stage IB (Grade 2, 3)<br>다. stage IC 이상<br>라. Sex Cord-Stromal Tumors |
| 5  | paclitaxel + cisplatin (Intra-peritoneal)<br>(제2006-4호: 2006.5.1, 개정 제2006-6호: 2006.8.1, 개정 제2022-190호: 2022.8.1.) | stage III인 환자에서 수술 후 잔류종양이 1cm 이하인 경우                                                                |

# Adjuvant CTx for advanced EOC

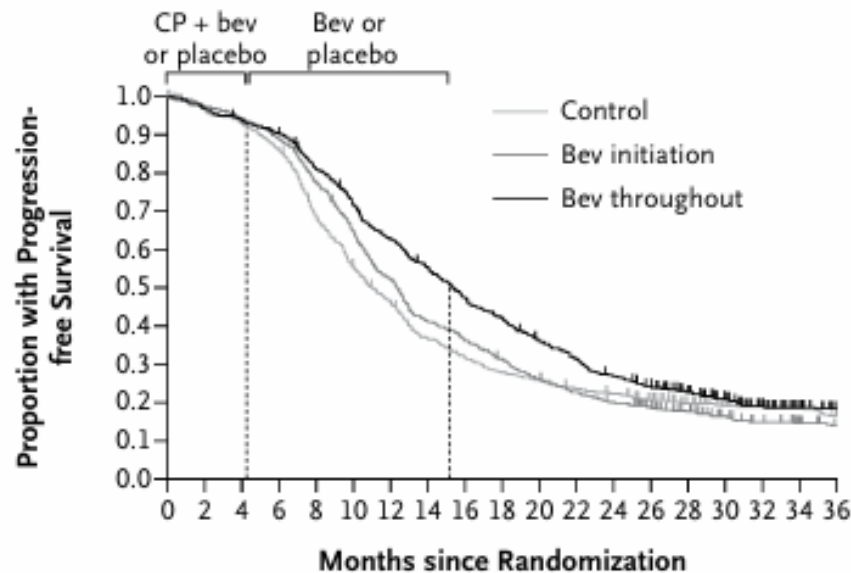
- Paclitaxel 175 mg/m<sup>2</sup> / Carboplatin AUC 5-6  
vs. Docetaxel 60-75 mg/m<sup>2</sup> / Carboplatin AUC 5-6
- No significant difference in efficacy (PFS/OS)
- Paclitaxel: more neuropathy, alopecia
- Docetaxel: more GI, peripheral edema, nail change  
→ considered for patients at high risk for neuropathy (e.g. diabetes)

# Adjuvant CTx for advanced EOC

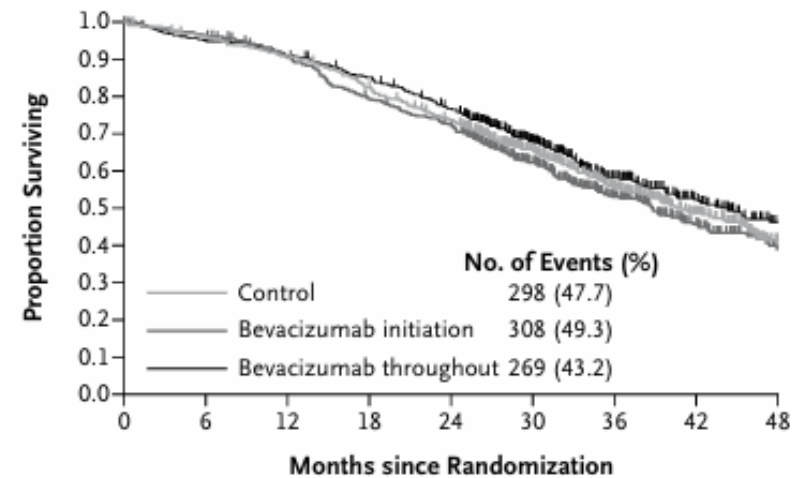
- Number of cycles
  - Stage I : 3 cycles (clear cell, endometrioid, mucinous)  
or 6 cycles for high-grade serous ca (GOG-157)
  - Stage II-IV : 6 cycles vs. >6 cycles
    - no evidence of additional clinical benefit of treatment beyond 6 cycles, increased neurotoxicity

# Adjuvant CTx for advanced EOC

- Addition of bevacizumab (GOG-218, ICON7)
- Paclitaxel/Carboplatin/Bevacizumab → Bevacizumab maintenance



| No. at Risk    |     |     |     |     |     |    |    |
|----------------|-----|-----|-----|-----|-----|----|----|
| Control        | 625 | 535 | 283 | 169 | 133 | 78 | 49 |
| Bev initiation | 625 | 552 | 319 | 190 | 121 | 67 | 40 |
| Bev throughout | 623 | 559 | 386 | 256 | 162 | 97 | 56 |



| No. at Risk            |     |     |     |     |     |     |     |     |    |
|------------------------|-----|-----|-----|-----|-----|-----|-----|-----|----|
| Control                | 625 | 595 | 558 | 506 | 446 | 322 | 200 | 116 | 56 |
| Bevacizumab initiation | 625 | 598 | 557 | 486 | 440 | 304 | 191 | 108 | 54 |
| Bevacizumab throughout | 623 | 587 | 561 | 519 | 463 | 321 | 201 | 114 | 62 |

# Adjuvant CTx for advanced EOC

- Addition of bevacizumab (GOG-218, ICON7)

| Trial   | Patients                                                        | First-line CTx<br>→ Maintenance       | FU<br>(mo) | PFS (median, mo)<br>HR [95% CI], p-value | OS (median, mo)<br>HR [95% CI], p-value |
|---------|-----------------------------------------------------------------|---------------------------------------|------------|------------------------------------------|-----------------------------------------|
| GOG-218 | Stage III<br>incompletely<br>resected or stage IV<br>85% serous | Arm 1: carbo/pac/placebo<br>→ placebo | 19.4       | 10.3                                     | 39.3                                    |
|         |                                                                 | Arm 2: carbo/pac/bev<br>→ placebo     |            | 11.2<br>0.908 [0.795-1.040], $P=.16$     | 38.7<br>1.036 [0.827-1.297], $P=.76$    |
|         |                                                                 | Arm 3: carbo/pac/bev<br>→ bev         |            | 14.1<br>0.717 [0.625-0.824], $P<.001$    | 39.7<br>0.915 [0.727-1.152], $P=.45$    |
| ICON7   | Stage I-IIA high-risk<br>or IIB-IV<br>69% serous                | Arm 1: carbo/pac<br>→ none            | 48.6       | 17.5                                     | 58.6                                    |
|         |                                                                 | Arm 2: carbo/pac/bev<br>→ bev         | 48.8       | 19.9<br>0.93 [0.83-1.05], $P=.25$        | 58.0<br>0.99 [0.85-1.14], $P=.25$       |

\* Greater benefit for patients with poor prognosis  
: inoperable stage III, suboptimally debulked (residual >1cm), stage IV

# Adjuvant CTx for advanced EOC

- Addition of bevacizumab (GOG-218, ICON7)

| Trial   | Treatments                                                                                                                                                                                                                        |
|---------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| GOG-218 | Paclitaxel 175 mg/m <sup>2</sup> IV + Carboplatin AUC 6 IV, q 3wks x cycles 1-6<br>+ bevacizumab 15 mg/kg q 3wks x cycles 2-6<br>→ maintenance bevacizumab 15 mg/kg q 3wks x cycles 7-22                                          |
| ICON7   | Paclitaxel 175 mg/m <sup>2</sup> IV + Carboplatin AUC 5-6 IV, q 3wks x 6 cycles<br>+ bevacizumab 7.5 mg/kg q 3wks x cycles 5-6 (omitted cycle 1 if <4 wks from surgery)<br>→ maintenance bevacizumab 7.5 mg/kg q 3wks x 12 cycles |

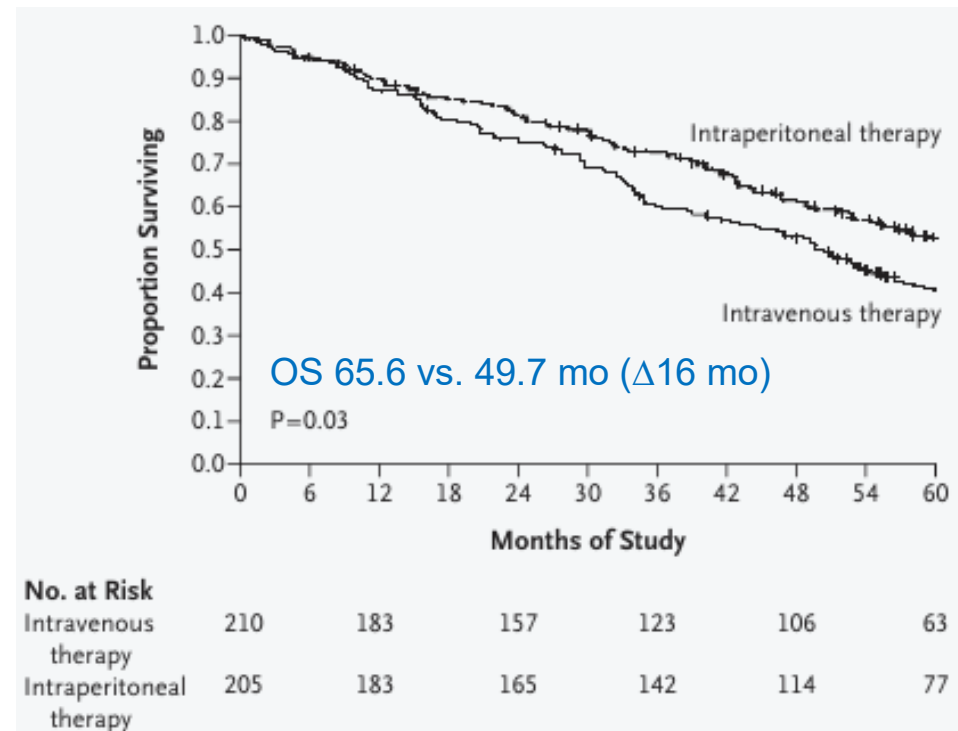
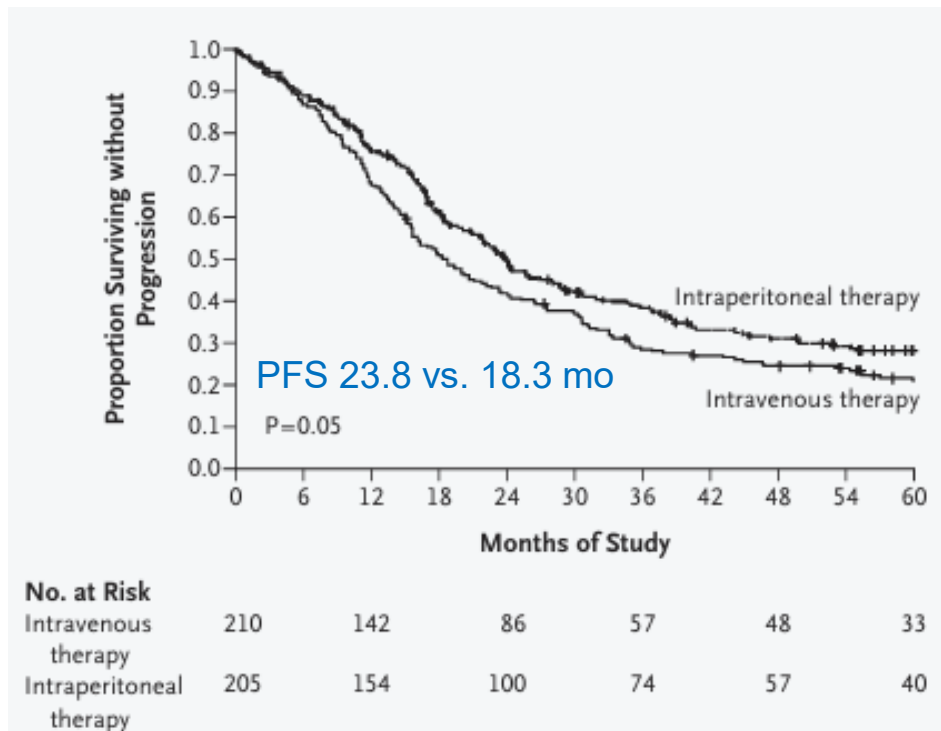
| 연번 | 항암요법                                      | 투여대상                                                                                                                                                                                                                                                                                                                           | 투여단계 |
|----|-------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------|
|    | bevacizumab<br>+ paclitaxel + carboplatin | <p>가. FIGO stage IIIb, IIIc 환자중 suboptimally debulked(잔존종양크기 &gt;1cm)이거나 stage IV 상피성 난소암, 난관암 또는 원발성 복막암</p> <p>1) bevacizumab은 7.5mg/kg로 투여하며, 화학요법 제2주기에서 제6주기까지 bevacizumab을 병용 투여하고, 이후 단독투여하며 최대 18주기까지 급여인정함. (단, 질병진행시 투여중단)</p> <p>2) 허가사항 범위 내에서 상기에 해당 하지 않는 front-line 요법의 경우 'bevacizumab'은 약값 전액을 본인이 부담토록 함</p> | 1차   |

# Adjuvant CTx for advanced EOC

- S/E of bevacizumab
  - Bleeding, hypertension, thromboembolic events
  - Proteinuria
  - GI perforation
  - Wound healing complications

# Adjuvant CTx for advanced EOC

- IP/IV chemotherapy (GOG-172)



# Adjuvant CTx for advanced EOC

- IP/IV chemotherapy

| Trial   | Patients                             | Regimen                                                                                                                                                                                                           | PFS<br>(median, mo) | OS<br>(median, mo) |
|---------|--------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------|--------------------|
| GOG-172 | Stage III<br>residual ds $\leq 1$ cm | IP/IV<br>Paclitaxel 135 mg/m <sup>2</sup> IV D1<br>+ cisplatin 100 mg/m <sup>2</sup> IP D2<br>+ paclitaxel 60 mg/m <sup>2</sup> IP D8, q 3wks x 6 cycles                                                          | 23.8                | 65.6               |
|         |                                      | IV<br>Paclitaxel 135 mg/m <sup>2</sup> IV D1<br>+ cisplatin 75 mg/m <sup>2</sup> IV D2, q 3wks x 6 cycles                                                                                                         | 18.3                | 49.7               |
| GOG-252 | Stage III/IV                         | IV/IP pac/carbo/bev<br>Paclitaxel 80 mg/m <sup>2</sup> IV D1,8,15<br>+ carboplatin AUC 6 IP D1, q 3wks x 6 cycles<br>Bevacizumab 15 mg/kg IV q 3wks cycles 2-22                                                   | 27.4                | 78.9               |
|         |                                      | IV/IP pac/cis/bev<br>Paclitaxel 135 mg/m <sup>2</sup> IV D1<br>+ cisplatin 75 mg/m <sup>2</sup> IP D2<br>+ paclitaxel 60 mg/m <sup>2</sup> IP D8, q 3wks x 6 cycles<br>Bevacizumab 15 mg/kg IV q 3wks cycles 2-22 | 26.2                | 72.9               |
|         |                                      | IV pac/carbo/bev<br>Paclitaxel 80 mg/m <sup>2</sup> IV D1,8,15<br>+ carboplatin AUC 6 IV D1, q 3wks x 6 cycles<br>Bevacizumab 15 mg/kg IV q 3wks cycles 2-22                                                      | 24.9                | 75.5               |

# Adjuvant CTx for advanced EOC

- IP chemotherapy
  - Stage III, optimally debulked (residual <1 cm) patients
  - Not recommended for stage I and IV
  - S/E: catheter complications, abdominal pain, (cisplatin-renal toxicity, neuropathy)
  - IP chemotherapy does not further improve outcomes on PFS benefit of adding bevacizumab (GOG-252)

# Primary Treatment for Ovarian Cancer

## Primary Systemic Therapy Regimens<sup>c</sup> - Epithelial Ovarian/Fallopian Tube/Primary Peritoneal

### Primary Therapy for Stage I Disease

|                                                                                                                                                                                             |                                                                                                                                                                                                                                                                                                                                            |                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           |                                                                                                                                                                                                                                                                                                                  |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <ul style="list-style-type: none"> <li>• High-grade serous</li> <li>• Endometrioid (grade 2/3)</li> <li>• Clear cell carcinoma<sup>d</sup></li> <li>• Carcinosarcoma<sup>d</sup></li> </ul> | <b>Preferred Regimens</b> <ul style="list-style-type: none"> <li>• Paclitaxel/carboplatin every 3 weeks<sup>g,h</sup></li> </ul>                                                                                                                                                                                                           | <b>Other Recommended Regimens</b> <ul style="list-style-type: none"> <li>• Carboplatin/liposomal doxorubicin</li> <li>• Docetaxel/carboplatin</li> </ul>                                                                                                                                                                                                                                                                                                                                  | <b>Useful in Certain Circumstances</b> <ul style="list-style-type: none"> <li>• Paclitaxel/cisplatin</li> </ul> <p>For carcinosarcoma:</p> <ul style="list-style-type: none"> <li>• Carboplatin/ifosfamide</li> <li>• Cisplatin/ifosfamide</li> <li>• Paclitaxel/ifosfamide (category 2B)<sup>g</sup></li> </ul> |
| <b>Mucinous carcinoma (stage IA, IB, and IC, grades 1–3)<sup>d</sup></b>                                                                                                                    | <b>Preferred Regimens</b> <ul style="list-style-type: none"> <li>• 5-FU/leucovorin/oxaliplatin</li> <li>• Capecitabine/oxaliplatin</li> <li>• Paclitaxel/carboplatin every 3 weeks<sup>g,h</sup></li> </ul>                                                                                                                                | <b>Other Recommended Regimens</b> <ul style="list-style-type: none"> <li>• Carboplatin/liposomal doxorubicin</li> <li>• Docetaxel/carboplatin</li> </ul>                                                                                                                                                                                                                                                                                                                                  | <b>Useful in Certain Circumstances</b> <ul style="list-style-type: none"> <li>• Paclitaxel/cisplatin</li> </ul>                                                                                                                                                                                                  |
| <b>Low-grade serous (stage IC)/Grade I endometrioid (stage IC)<sup>d,e,f</sup></b>                                                                                                          | <b>Preferred Regimens</b> <ul style="list-style-type: none"> <li>• Paclitaxel/carboplatin every 3 weeks<sup>g,h</sup> ± maintenance letrozole (category 2B<sup>f</sup>) or other hormonal therapy (category 2B)<sup>i</sup></li> <li>• Hormone therapy (aromatase inhibitors: anastrozole, letrozole, exemestane) (category 2B)</li> </ul> | <b>Other Recommended Regimens</b> <ul style="list-style-type: none"> <li>• Carboplatin/liposomal doxorubicin ± maintenance letrozole (category 2B<sup>f</sup>) or other hormonal therapy (category 2B)<sup>i</sup></li> <li>• Docetaxel/carboplatin ± maintenance letrozole (category 2B<sup>f</sup>) or other hormonal therapy (category 2B)<sup>i</sup></li> <li>• Hormone therapy (leuprolide acetate, goserelin acetate, tamoxifen,<sup>j</sup> fulvestrant) (category 2B)</li> </ul> | <b>Useful in Certain Circumstances</b> <ul style="list-style-type: none"> <li>• Paclitaxel/cisplatin</li> </ul>                                                                                                                                                                                                  |

| Primary Therapy for                                                                                                                                                                         | Stage II–IV Disease (Principles of Maintenance PARPi Therapy on OV-C, 3 of 12)                                                                                                                                                                                                                                                                                                                                                                        | Other Recommended Regimens                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    | Useful in Certain Circumstances                                                                                                                                                                                                                                                                                                                                                                                                                                                   |
|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <ul style="list-style-type: none"> <li>• High-grade serous</li> <li>• Endometrioid (grade 2/3)</li> <li>• Clear cell carcinoma<sup>d</sup></li> <li>• Carcinosarcoma<sup>d</sup></li> </ul> | <b>Preferred Regimens</b> <ul style="list-style-type: none"> <li>• Paclitaxel/carboplatin every 3 weeks<sup>g,h</sup></li> <li>• Paclitaxel/carboplatin/bevacizumab + maintenance bevacizumab<sup>g</sup> (ICON-7 &amp; GOG-218)</li> </ul>                                                                                                                                                                                                           | <ul style="list-style-type: none"> <li>• Paclitaxel weekly/carboplatin weekly<sup>g,h,k</sup></li> <li>• Docetaxel/carboplatin</li> <li>• Carboplatin/liposomal doxorubicin</li> <li>• Paclitaxel weekly/carboplatin every 3 weeks<sup>g</sup></li> <li>• Docetaxel/carboplatin/bevacizumab + maintenance bevacizumab (GOG-218)</li> </ul>                                                                                                                                                                                                                                                                                                                                                                    | <ul style="list-style-type: none"> <li>• Paclitaxel/cisplatin</li> <li>• Docetaxel/oxaliplatin/bevacizumab + maintenance bevacizumab</li> <li>• IV/IP paclitaxel/carboplatin</li> <li>• IV/IP paclitaxel/cisplatin (for optimally debulked stage II–III disease)</li> <li>• For carcinosarcoma: <ul style="list-style-type: none"> <li>▶ Carboplatin/ifosfamide</li> <li>▶ Cisplatin/ifosfamide</li> <li>▶ Paclitaxel/ifosfamide (category 2B)<sup>g</sup></li> </ul> </li> </ul> |
| Mucinous carcinoma <sup>d</sup>                                                                                                                                                             | <b>Preferred Regimens</b> <ul style="list-style-type: none"> <li>• 5-FU/leucovorin/oxaliplatin ± bevacizumab (category 2B for bevacizumab)</li> <li>• Capecitabine/oxaliplatin ± bevacizumab (category 2B for bevacizumab)</li> <li>• Paclitaxel/carboplatin every 3 weeks<sup>g,h</sup></li> <li>• Paclitaxel/carboplatin/bevacizumab + maintenance bevacizumab<sup>g</sup> (ICON-7 &amp; GOG-218)</li> </ul>                                        | <b>Other Recommended Regimens</b> <ul style="list-style-type: none"> <li>• Paclitaxel weekly/carboplatin weekly<sup>g,h,k</sup></li> <li>• Docetaxel/carboplatin</li> <li>• Carboplatin/liposomal doxorubicin</li> <li>• Paclitaxel weekly/carboplatin every 3 weeks<sup>g</sup></li> <li>• Docetaxel/carboplatin/bevacizumab + maintenance bevacizumab (GOG-218)</li> </ul>                                                                                                                                                                                                                                                                                                                                  | <b>Useful in Certain Circumstances</b> <ul style="list-style-type: none"> <li>• Paclitaxel/cisplatin</li> <li>• Docetaxel/oxaliplatin/bevacizumab + maintenance bevacizumab</li> </ul>                                                                                                                                                                                                                                                                                            |
| Low-grade serous/Grade I endometrioid <sup>d,e,f</sup>                                                                                                                                      | <b>Preferred Regimens</b> <ul style="list-style-type: none"> <li>• Paclitaxel/carboplatin every 3 weeks<sup>g,h</sup> ± maintenance letrozole (category 2B<sup>f</sup>) or other hormonal therapy (category 2B)<sup>i</sup></li> <li>• Paclitaxel/carboplatin/bevacizumab + maintenance bevacizumab<sup>g</sup> (ICON-7 &amp; GOG-218)</li> <li>• Hormone therapy (aromatase inhibitors: anastrozole, letrozole, exemestane) (category 2B)</li> </ul> | <b>Other Recommended Regimens</b> <ul style="list-style-type: none"> <li>• Paclitaxel weekly/carboplatin weekly<sup>g,h,k</sup></li> <li>• Docetaxel/carboplatin ± maintenance letrozole (category 2B<sup>f</sup>) or other hormonal therapy (category 2B)<sup>i</sup></li> <li>• Carboplatin/liposomal doxorubicin ± maintenance letrozole (category 2B<sup>f</sup>) or other hormonal therapy (category 2B)<sup>i</sup></li> <li>• Paclitaxel weekly/carboplatin every 3 weeks<sup>g</sup></li> <li>• Docetaxel/carboplatin/bevacizumab + maintenance bevacizumab (GOG-218)</li> <li>• Hormone therapy (leuprolide acetate, goserelin acetate, tamoxifen,<sup>j</sup> fulvestrant) (category 2B)</li> </ul> | <b>Useful in Certain Circumstances</b> <ul style="list-style-type: none"> <li>• Paclitaxel/cisplatin</li> <li>• Docetaxel/oxaliplatin/bevacizumab + maintenance bevacizumab (category 2B)</li> </ul>                                                                                                                                                                                                                                                                              |

# Neoadjuvant CTx for advanced EOC

- Not good candidates for PDS d/t advanced age, frailty, poor performance status, comorbidities
  - Unlikely to be optimally cytoreduced
- **Neoadjuvant CTx (NAC)** to reduce tumor burden before surgery
- increase likelihood of optimal cytoreduction
  - may improve condition and reduce perioperative risks

# Neoadjuvant CTx for advanced EOC

- NAC x 3-4 cycles → IDS → postop CTx x 3-4 cycles (total 6-8 cycles)  
(same chemotherapy regimen for both NAC and postop CTx)
- Paclitaxel/Carboplatin q 3wks x 3~4 cycles
- Biopsy for histologic confirmation before NAC !  
(core biopsy preferred than cytologic diagnosis)

## 1. 선행화학요법 (neoadjuvant)

| 연번 | 항암요법                                                                                                      | 투여대상                                 |
|----|-----------------------------------------------------------------------------------------------------------|--------------------------------------|
| 1  | paclitaxel + carboplatin<br>(제2007-7호: 2007.11.20, 개정 제2015-291호: 2015.12.1,<br>개정 제2022-190호: 2022.8.1.) | 수술 불가능한 stage IIIc~IV<br>(투여기간: 3주기) |

# Maintenance Tx

- Bevacizumab
  - Paclitaxel/Carboplatin/Bevacizumab → Bevacizumab
- PARPi
  - Olaparib / Olaparib+bevacizumab
  - Niraparib

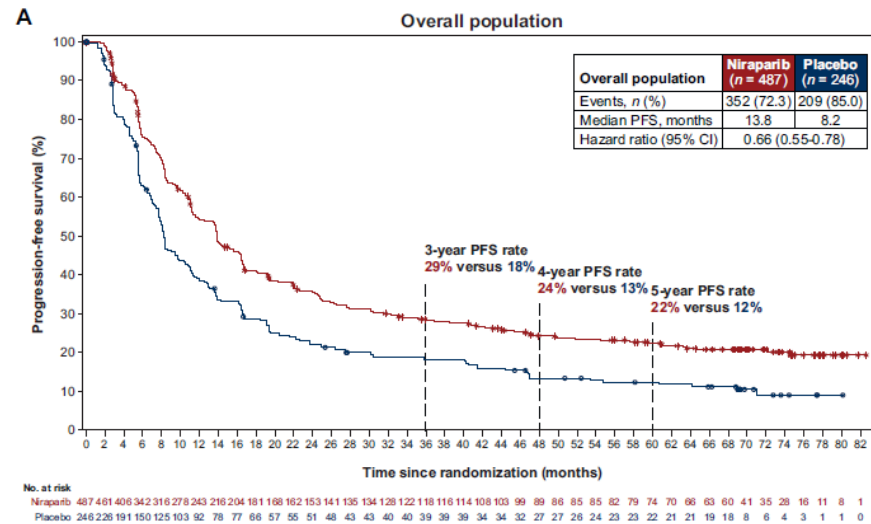
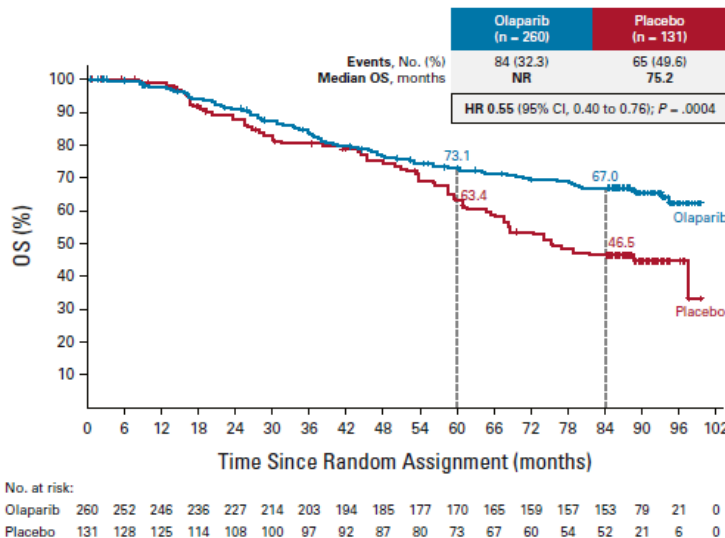
Individualized maintenance therapy based on genetic and molecular testing

BRCA gene variant  
or  
Positive homologous  
recombination deficiency test

- PARP inhibitor alone (olaparib or niraparib)
- Olaparib + bevacizumab

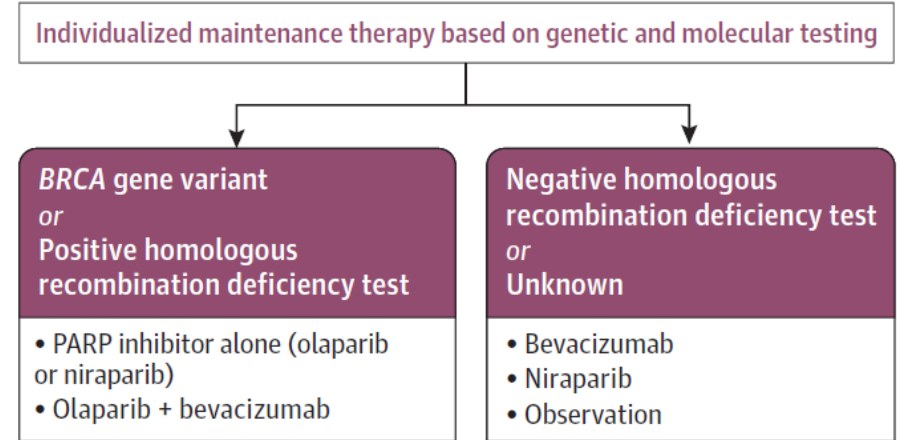
Negative homologous  
recombination deficiency test  
or  
Unknown

- Bevacizumab
- Niraparib
- Observation



# Maintenance Tx

- PARPi
  - Olaparib / Olaparib+bevacizumab
  - Niraparib



| Drugs                           | Results                                                                                                   | Dosage                           |
|---------------------------------|-----------------------------------------------------------------------------------------------------------|----------------------------------|
| Olaparib (SOLO1)                | Median PFS 56 vs. 13.8 mo (HR 0.33) in BRCAmut+                                                           | 300mg bid (2y)                   |
| Niraparib (PRIMA)               | Median PFS 21.9 vs. 10.4 mo (HR 0.43) in HRD+<br>Median PFS 13.8 vs. 8.2 mo (HR 0.62) in overall (HRD+/-) | 200mg qd                         |
| Olaparib +Bevacizumab (PAOLA-1) | Median PFS 37.2 vs. 17.7 mo (HR 0.33) in HRD+                                                             | 300mg bid (2y)<br>+15mg/kg (15m) |

# Maintenance Tx

## 4. 유지요법 (maintenance)

| 연번 | 항암요법                                                                                                                          | 투여대상                                                                                                                     | 투여단계 |
|----|-------------------------------------------------------------------------------------------------------------------------------|--------------------------------------------------------------------------------------------------------------------------|------|
| 1  | niraparib <sup>주1</sup>                                                                                                       | 가. 1차 백금기반요법에 반응(CR 또는 PR)한 진행성 상동재조합결핍 양성(BRCA 변이 또는 유전체 불안정성) 상피성 난소암, 난관암, 일차 복막암<br>※ 백금계 항암제 완료 후 12주 이내 투여         | -    |
|    | (제2019-331호: 2019.12.1, 개정 제2021-26호: 2021.2.1, 개정 제2021-242호: 2021.10.1, 개정 제2022-190호: 2022.8.1, 개정 제2024-219호: 2024.10.1.) | 나. 2차 이상의 백금기반요법에 반응(CR 또는 PR)한 백금민감성 재발성 BRCA 변이 고도 장액성 난소암, 난관암, 일차 복막암<br>※ 백금계 항암제 완료 후 8주 이내 투여                     |      |
| 2  | olaparib(tablet) <sup>주2</sup>                                                                                                | 가. 1차 백금기반요법에 반응(CR 또는 PR)한 진행성 BRCA 변이 고도 상피성 난소암, 난관암, 일차 복막암<br>※ 백금계 항암제 완료 후 8주 이내 투여<br>※ 투여기간: 최초 투여 후 2년까지 급여 인정 | -    |
|    | (제2021-242호: 2021.10.1, 개정 제2022-190호: 2022.8.1.)                                                                             | 나. 2차 이상의 백금기반요법에 반응(CR 또는 PR)한 백금민감성 재발성 BRCA 변이 고도 상피성 난소암, 난관암, 일차 복막암<br>※ 백금계 항암제 완료 후 8주 이내 투여                     |      |

주1. 유지요법 시행 직전 투여된 백금기반요법은 bevacizumab 포함 요법을 제외하며, 이전에 PARP 억제제를 투여받은 적이 없어야 함

주2. 유지요법 시행 직전 투여된 백금기반요법(bevacizumab 포함 요법 제외)은 최소 4주기 이상 투여해야 하며, 이전에 PARP 억제제를 투여받은 적이 없어야 함

# Paclitaxel

- Dose: 135(24h)-175(3h) mg/m<sup>2</sup> ; weekly 60-80 mg/m<sup>2</sup>
- 3-weekly regimen: infusion over 3 hours
- Weekly regimen: infusion over 1 hour
- Should be administered before platinum (carboplatin/cisplatin) !!
- Hypersensitivity reactions
- Neuropathy : ~60%, dose-dependent
- Myalgia, arthralgia

# Paclitaxel

## – Hypersensitivity reactions

: ~20-40%, usually within 2-5 minutes of infusion

/ urticaria, flush, chest tightness, dyspnea, hypotension

## - Premedications : three-drug prophylactic regimen

: Dexamethasone 20 mg IV 30 mins before paclitaxel

(alternative: 20 mg PO/IV at 12 and 6 hrs before paclitaxel)

+ Diphenhydramine 50 mg IV (Chlorpheniramine 1 amp), 30 mins before paclitaxel

+ Famotidine 20 mg IV, 30 mins before paclitaxel

## – Management

- Discontinue the infusion
- Antihistamine + hydrocortisone 100 mg IV
- If symptom subsides, restart with slower infusion rate
- prn) Desensitization protocol (알레르기내과 consult)



# Docetaxel

- Dose: 60-75 mg/m<sup>2</sup>
- Infusion over 1 hour
- Administered before platinum
- Fluid retention (edema, pleural effusion..; ~60%)
  - ↔ dexamethasone 8 mg bid 3-5 days, starting 1 day before docetaxel
- Hypersensitivity reaction

# Carboplatin

- Nephrotoxicity
  - ↔ Dosing with Calvert formula  
(desired AUC x (25 + GFR))
- Myelosuppression
- Monitoring parameters
  - : CBC, renal function test (CCr: Cockcroft formula), LFT

$$*CCr \text{ (female)} = ((140 - \text{age}) \times \text{Bwt (kg)} / (72 \times \text{SCr})) \times 0.85$$



## 2. Platinum

### 1) Desensitization protocols (Brigham and Women's Hospital, 2010)

- Platinum 제제는 rechallenge시 50%이상에서 severe reactions을 일으킬 수 있으므로, resuscitation drugs (epinephrine, antihistamines, bronchodilators, steroids) 준비 후 시작
- Allergy 내과에 consult 후 시행

#### • Solution preparation (carboplatin 500 mg 기준)

|            | Volume (NS) | Concentration | Total mg of drug |
|------------|-------------|---------------|------------------|
| Solution 1 | 250 mL      | 0.02 mg/mL    | 5                |
| Solution 2 | 250 mL      | 0.2 mg/mL     | 50               |
| Solution 3 | 250 mL      | 1.772 mg/mL   | 496.065          |

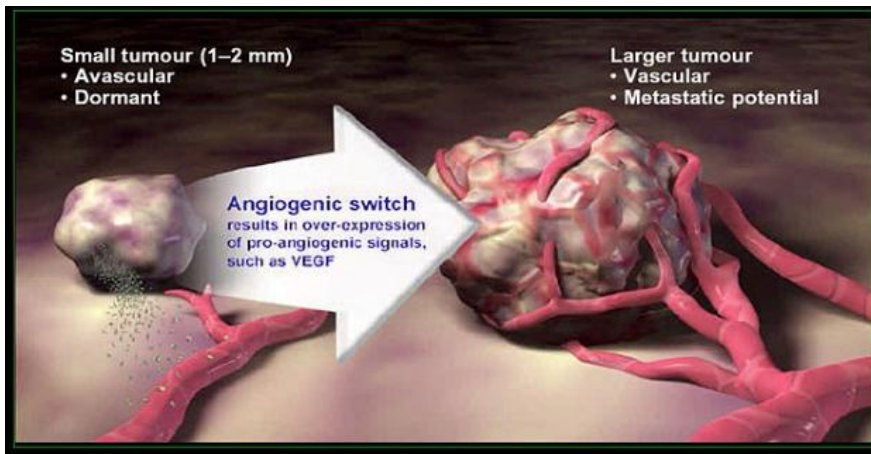
#### • Protocol for administration (carboplatin 500 mg 기준)

| Step | Solution | Rate<br>(mL/h) | Time<br>(mins) | Volume infused<br>(mL) | Administered<br>dose (mg) |
|------|----------|----------------|----------------|------------------------|---------------------------|
| 1    | 1        | 2              | 15             | 0.5                    | 0.01                      |
| 2    | 1        | 5              | 15             | 1.25                   | 0.025                     |
| 3    | 1        | 10             | 15             | 2.5                    | 0.05                      |
| 4    | 1        | 20             | 15             | 5                      | 0.1                       |
| 5    | 2        | 5              | 15             | 1.25                   | 0.25                      |
| 6    | 2        | 10             | 15             | 2.5                    | 0.5                       |
| 7    | 2        | 20             | 15             | 5                      | 1                         |
| 8    | 2        | 40             | 15             | 10                     | 2                         |
| 9    | 3        | 10             | 15             | 2.5                    | 4.4292                    |
| 10   | 3        | 20             | 15             | 5                      | 8.8583                    |
| 11   | 3        | 40             | 15             | 10                     | 17.7166                   |
| 12   | 3        | 75             | 210            | 262.5                  | 465.0609                  |

**알레르기내과 consult!**

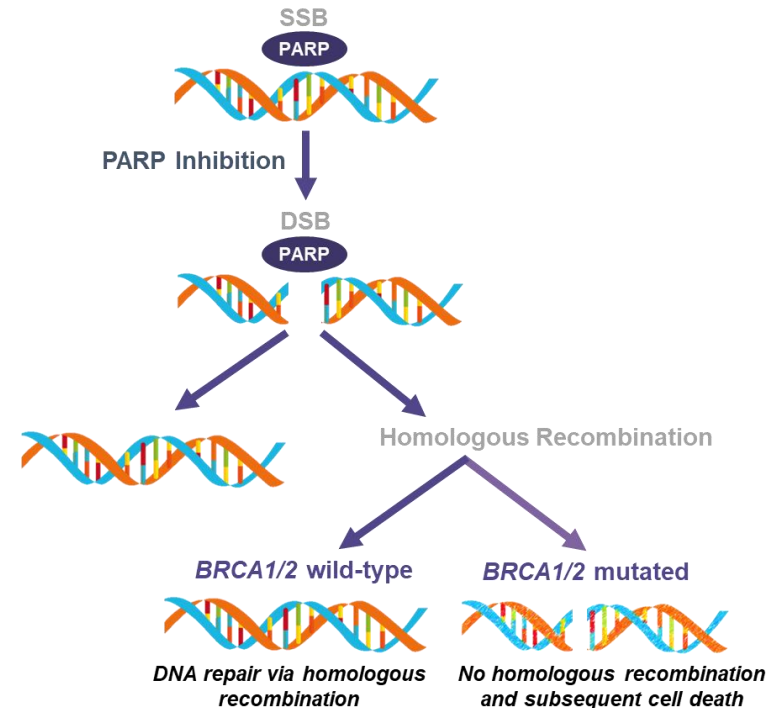
# Bevacizumab

- Mechanisms of action
  - : a recombinant humanized monoclonal Ab that blocks angiogenesis by inhibiting VEGF-A
- S/E: hypertension, proteinuria, hemorrhage, VTE, interruption of wound healing, GI fistula, ischemic/hemorrhagic strokes
- Monitoring: BP check, U/A (prn, 24hr urine collection in Prot 2+ pts)



# PARP inhibitors

- Mechanisms of action
  - : ‘Synthetic lethality’ in BRCA mutated cancers (germline/somatic BRCAm)
- S/E: anemia, neutropenia, thrombocytopenia, fatigue, N/V, diarrhea, MDS/AML (<1.5%)
- Monitoring: CBC, LFT, BUN/Cr (Chest PA if pneumonitis sx occurs)



# 2026년 대한부인종양학회 제7회 동계학술대회 with Chemo-TIP Review

일자 2026년 1월 17일 (토)

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## Thank you for your attention!



대한부인종양학회  
Korean Society of Gynecologic Oncology

