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(54) Title **COMBINATION THERAPY FOR THE TREATMENT OF OVARIAN CANCER**

(56) References Cited: WO-A1-2008/075330, US-A1- 2007 166 388, US-A1- 2006 013 819  
COHEN MARTIN H ET AL: "FDA drug approval summary: bevacizumab plus FOLFOX4 as second-line treatment of colorectal cancer.", THE ONCOLOGIST MAR 2007, vol. 12, no. 3, March 2007 (2007-03), pages 356-361, XP002712135, ISSN: 1083-7159  
BAMIAS A ET AL: "Angiogenesis: a promising therapeutic target for ovarian cancer.", CRITICAL REVIEWS IN ONCOLOGY/HEMATOLOGY DEC 2012, vol. 84, no. 3, December 2012 (2012-12), pages 314-326, XP002698499, ISSN: 1879-0461  
LORTHOLARY A ET AL: "Weekly paclitaxel as a single agent or in combination with carboplatin or weekly topotecan in patients with resistant ovarian cancer: the CARTAXHY randomized phase II trial from GINECO", ANNALS OF ONCOLOGY, vol. 23, 2012, pages 346-352, Clinical Trials Identifier : NCT00976911: "Aurelia: A study of Avastin (Bevacizumab) added to chemotherapy in patients with platinum-resistant ovarian cancer", The National library of Medicine , 15 March 2011 (2011-03-15), Retrieved from the Internet: URL:clinicaltrials.gov [retrieved on 2017-06-12]  
RICHARDSON D L ET AL: "Combination gemcitabine, platinum, and bevacizumab for the

treatment of recurrent ovarian cancer", GYNECOLOGIC ONCOLOGY, ACADEMIC PRESS, LONDON, GB, vol. 111, no. 3, 1 December 2008 (2008-12-01), pages 461-466, XP025691004, ISSN: 0090-8258, DOI: 10.1016/J.YGYNO.2008.08.011 [retrieved on 2008-09-30]

MCGONIGLE KATHRYN F ET AL: "Combined weekly topotecan and biweekly bevacizumab in women with platinum-resistant ovarian, peritoneal, or fallopian tube cancer: results of a phase 2 study.", CANCER 15 AUG 2011, vol. 117, no. 16, 15 August 2011 (2011-08-15), pages 3731-3740, XP002712134, ISSN: 1097-0142

COHN D E ET AL: "Bevacizumab and weekly taxane chemotherapy demonstrates activity in refractory ovarian cancer", GYNECOLOGIC ONCOLOGY, ACADEMIC PRESS, LONDON, GB, vol. 102, no. 2, 1 August 2006 (2006-08-01), pages 134-139, XP024921475, ISSN: 0090-8258, DOI: 10.1016/J.YGYNO.2006.01.030 [retrieved on 2006-08-01]

D. M. O'MALLEY & AL: "Addition of bevacizumab to weekly paclitaxel significantly improves progression-free survival in heavily pretreated recurrent epithelial ovarian cancer", GYNECOLOGIC ONCOLOGY, vol. 121, 1 January 2011 (2011-01-01), pages 269-272,

J. A. LEDERMANN, F. A. RAJA: "Clinical trials and decision-making strategies for optimal treatment of relapsed ovarian cancer", EUROPEAN JOURNAL OF CANCER, vol. 47, no. 3, 1 September 2011 (2011-09-01), pages s104-s115,

GERBER HANS-PETER ET AL: "Pharmacology and pharmacodynamics of bevacizumab as monotherapy or in combination with cytotoxic therapy in preclinical studies", CANCER RESEARCH, AMERICAN ASSOCIATION FOR CANCER RESEARCH, US, vol. 65, no. 3, 1 February 2005 (2005-02-01), pages 671-680, XP002481174, ISSN: 0008-5472

MONK B J ET AL: "Salvage bevacizumab (rhuMAB VEGF)-based therapy after multiple prior cytotoxic regimens in advanced refractory epithelial ovarian cancer", GYNECOLOGIC ONCOLOGY, ACADEMIC PRESS, LONDON, GB, vol. 102, no. 2, 1 August 2006 (2006-08-01), pages 140-144, XP024921476, ISSN: 0090-8258, DOI: 10.1016/J.YGYNO.2006.05.006 [retrieved on 2006-08-01]

CHURA ET AL: "Bevacizumab plus cyclophosphamide in heavily pretreated patients with recurrent ovarian cancer", GYNECOLOGIC ONCOLOGY, ACADEMIC PRESS, LONDON, GB, vol. 107, no. 2, 1 November 2007 (2007-11-01), pages 326-330, XP022350665, ISSN: 0090-8258, DOI: 10.1016/J.YGYNO.2007.07.017

TSUNETOH SATOSHI ET AL: "Topotecan as a molecular targeting agent which blocks the Akt and VEGF cascade in platinum-resistant ovarian cancers.", CANCER BIOLOGY & THERAPY 1 DEC 2010, vol. 10, no. 11, 1 December 2010 (2010-12-01), pages 1137-1146, XP002712136, ISSN: 1555-8576

WRIGHT JASON D ET AL: "Bevacizumab combination therapy in recurrent, platinum-refractory, epithelial ovarian carcinoma: A retrospective analysis.", CANCER 1 JUL 2006, vol. 107, no. 1, 1 July 2006 (2006-07-01), pages 83-89, XP002698498, ISSN: 0008-543X

Enclosed is a translation of the patent claims in Norwegian. Please note that as per the Norwegian Patents Acts, section 66i the patent will receive protection in Norway only as far as there is agreement between the translation and the language of the application/patent granted at the EPO. In matters concerning the validity of the patent, language of the application/patent granted at the EPO will be used as the basis for the decision. The patent documents published by the EPO are available through Espacenet (<http://worldwide.espacenet.com>) or via the search engine on our website here: <https://search.patentstyret.no/>

**PATENTKRAV**

1. Bevacizumab for anvendelse i en fremgangsmåte for behandling av en pasient som er diagnostisert med en platinaresistent epitelial eggstokkreft (EOC), et platinaresistent egglederkarzinom (FTC) eller et platinaresistent primært peritonealt karzinom (PPC), hvori fremgangsmåten omfatter å administrere til pasienten en effektiv mengde av bevacizumab og paklitaksel, hvori pasienten mottok to eller færre forutgående anti-kreft-behandlingsprogrammer, hvori behandlingen forlenger pasientens gjennomsnittlige progresjonsfrie overlevelsestid sammenlignet med en pasient med en platinaresistent epitelial eggstokkreft (EOC), et platinaresistent egglederkarzinom (FTC) eller et platinaresistent primært peritonealt karzinom (PPC) som kun mottar paklitaksel.
2. Bevacizumab for anvendelse ifølge krav 1, hvori pasienten ikke er refraktær overfor tidligere platinabehandling.
3. Bevacizumab for anvendelse ifølge krav 1, hvori pasienten har en målbar sykdom ifølge RECIST 1.0 eller en CA-125 påvisbar sykdom ifølge GCIG-kriteriene.
4. Bevacizumab for anvendelse ifølge krav 1, hvori pasienten har en ECOG-ytelsesstatus på 0-2 og en forventet gjenværende levetid på minst 12 uker.
5. Bevacizumab for anvendelse ifølge krav 1, hvori den effektive mengden av paklitakselet administreres ved 80 mg/m<sup>2</sup> som en 1-times intravenøs infusjon på dag 1, 8, 15 og 22 hver fjerde uke.
6. Bevacizumab for anvendelse ifølge krav 1, hvori den effektive mengden av bevacizumab er 10 mg/kg intravenøst hver annen uke eller 15 mg/kg intravenøst hver tredje uke.
7. Bevacizumab for anvendelse ifølge krav 6, hvori den effektive mengden av bevacizumab initialt administreres intravenøst over 90 minutter, med etterfølgende infusjoner over 60 minutter og deretter 30 minutter.
8. Bevacizumab for anvendelse ifølge krav 1, hvori pasienten har et platinafritt intervall (PFI) på mindre enn 3 måneder.
9. Bevacizumab for anvendelse ifølge krav 1, hvori pasienten har

abdominal ascites.

10. Bevacizumab for anvendelse ifølge krav 1, hvori behandlingen ytterligere forbedrer pasientens objektive responsrate (ORR) sammenlignet med en pasient med platinaresistent eggstokkreft som kun mottar paklitaxel.