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(54) **Title:** COMBINATION OF A VEGF RECEPTOR INHIBITOR WITH A CHEMOTHERAPEUTIC AGENT

(57) **Abstract:** The present invention relates to a combination therapy for treating patients suffering from proliferative diseases or diseases associated with persistent angiogenesis. The patient is treated with: (a) a VEGF inhibitor compound; and (b) one or more chemotherapeutic agents selected from the group consisting of: i. an aromatase inhibitor; ii. an anti-estrogen, an anti-androgen (especially in the case of prostate cancer) or a gonadorelin agonist; iii. a topoisomerase I inhibitor or a topoisomerase II inhibitor; iv. a microtubule active agent, an alkylating agent, an anti-neoplastic antimetabolite or a platin compound; v. a compound targeting/decreasing a protein or lipid kinase activity or a protein or lipid phosphatase activity, a further anti-angiogenic compound or a compound which induces cell differentiation processes; vi. a bradykinin 1 receptor or an angiotensin II antagonist; vii. a cyclooxygenase inhibitor, a bisphosphonate, a heparanase inhibitor (prevents heparan sulphate degradation), e.g., PI-88, a biological response modifier, preferably a lymphokine or interferons, e.g., interferon γ, an ubiquitination inhibitor, or an inhibitor which blocks anti-apoptotic pathways; viii. an inhibitor of Ras oncogenic isoforms or a farnesyl transferase inhibitor; ix. a telomerase inhibitor, e.g., telomestatin; x. a protease inhibitor, a matrix metalloproteinase inhibitor, a methionine aminopeptidase inhibitor, e.g., bengamide or a derivative thereof, or a proteasome inhibitor, e.g., PS-341; xi. agents used in the treatment of hematologic malignancies or FMS-like tyrosine kinase inhibitors; xii. an HSP90 inhibitors; xiii. HDAC inhibitors; xiv. mTOR inhibitors; xv. somatostatin receptor antagonists; xvi. integrin antagonists; xvii. anti-leukemic compounds; xviii. tumor cell damaging approaches such as ionizing radiation; xix. EDG binders; xx. anthranilic acid amide class of kinase inhibitors; xxi. ribonucleotide reductase inhibitors; xxii. S-adenosylmethionine decarboxylase inhibitors; xxiii. antibodies against VEGF or VEGFR; xxiv. photodynamic therapy; xxv. angiostatic steroids; xxvi. implants containing corticosteroids; xxvii. AT1 receptor antagonists; and xxviii. ACE inhibitors.

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