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(54) Title: COMPOSITIONS AND METHODS TO TREAT CANCER WITH CUPREDOXINS AND CPG RICH DNA

(57) Abstract: The present invention relates to compositions comprising CpG rich DNA from Pseudomonas aeruginosa. The com-
positions optionally comprise a cupredoxin. The present invention includes specific CpG DNAs from Pseudomonas aeruginosa that
are useful for treating cancer and other conditions in patients. These compositions are optionally in a pharmaceutically acceptable
carrier and also optionally comprise a cupredoxin. The present invention further relates to methods to express proteins near cancer
cells. These methods may be used to express therapeutic or diagnostic proteins near cancer cells in a patient suffering from cancer
or other conditions, and can also be used for diagnosing cancer in a patient. This method uses the gene for azurin from P. aeruginosa
as an expression system for azurin or heterologous proteins in P. aeruginosa or heterologous cells.

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INTERNATIONAL SEARCH REPORT

International application No.

PCT/US07/86393

A. CLASSIFICATION OF SUBJECT MATTER
 IPC: Please See Continuation Sheet

USPC: Please See Continuation Sheet
 According to International Patent Classification (IPC) or to both national classification and IPC

B. FIELDS SEARCHED

Minimum documentation searched (classification system followed by classification symbols)
 U.S. : Please See Continuation Sheet

Documentation searched other than minimum documentation to the extent that such documents are included in the fields searched

Electronic data base consulted during the international search (name of data base and, where practicable, search terms used)
 Please See Continuation Sheet

C. DOCUMENTS CONSIDERED TO BE RELEVANT

Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WO 2005/018662 A1 (BOARD OF TRUSTEES OF THE UNIVERSITY OF ILLINOIS) 03 March 2005 (03.03.2005), see entire document.	1-32
A, P	WO 2007/024368 A2 (THE BOARD OF TRUSTEES OF THE UNIVERSITY OF ILLINOIS) 01 March 2007 (01.03.2007), see entire document.	1-32
A	US 7084105 B2 (CHAKRABARTY et al) 01 August 2006 (01.08.2006), see entire document.	1-32
A	US 2006/0149037 A1 (CHAKRABARTY et al) 06 July 2006 (06.07.2006), see entire document.	1-32
A, T	US 7381701 B2 (CHAKRABARTY et al) 03 June 2008 (03.06.2008), see entire document.	1-32
A, P	US 2006/0292136 A1 (CHAKRABARTY et al) 28 December 2006 (28.12.2006), see entire document.	1-32

Further documents are listed in the continuation of Box C. See patent family annex.

* Special categories of cited documents:	"T"	later document published after the international filing date or priority date and not in conflict with the application but cited to understand the principle or theory underlying the invention
"A" document defining the general state of the art which is not considered to be of particular relevance	"X"	document of particular relevance; the claimed invention cannot be considered novel or cannot be considered to involve an inventive step when the document is taken alone
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C (Continuation). DOCUMENTS CONSIDERED TO BE RELEVANT		
Category *	Citation of document, with indication, where appropriate, of the relevant passages	Relevant to claim No.
A	WU. H.J. Azurin of pathogenic <i>Neisseria</i> ssp. is involved in defense against hydrogen peroxide and survival within cervical epithelial cells. <i>Infection and Immunity</i> . December 2005, Vol 73. No. 12, pages 8444-8448, see entire document.	1-32
A	PUNJ. V. Microbial-based therapy of cancer. <i>Cancer Biology and Therapy</i> . August 2004, Vol 3. Issue 8, pages 708-714, see entire document.	1-32
A	PUNJ. V. Bacterial cupredoxin azurin and its interactions with the tumor suppressor protein p53. <i>Biochemical and Biophysical Research Communications</i> . 2003, Vol 312, pages 109-114, see entire document.	1-32
Y, P	MAHFOUZ. M. Bacterial proteins and CpG-rich extrachromosomal DNA in potential cancer therapy. <i>Plasmid</i> . 2007, Vol 57, pages 4-17, see entire document.	1-32
A	PUNJ. V. Bacterial cupredoxin azurin as a inducer of apoptosis and regression in huamn breast cancer. <i>Oncogene</i> . 2004, Vol. 23, pages 2367-2378, see entire document.	1-32
A	YAMADA. T. Bacterial redox protein azurin, tumor suppressor protein p53, and regression of cancer. <i>PNAS</i> . October 29, 2002, Vol 99. No. 22, pages 14098-14103, see entire document.	1-32
Y	YAMADA. T. Internalization of bacterial redox protein azurin in mammalian cells: entry domain and specificity. <i>Cellular Microbiology</i> . 2005, Vol 7. No. 10, pages 1418-1431, see entire document.	1-32

INTERNATIONAL SEARCH REPORT

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Continuation of B. FIELDS SEARCHED Item 1:

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Continuation of B. FIELDS SEARCHED Item 3:

WEST, PALM, SCIENCEDIRECT, PUBMED, BIOSIS, MEDLINE, BIOTECHDS, CA, CABA, CAPLUS, DISSABS, AGRICOLA, CONFSCI, LIFESCI, SCISEARCH

search terms: inventor names pseudomonas aeruginosa, nucleic acid, dna, peptide, cupredoxin, cancer, melanoma, azurin, treatment, CpG, bacterial dna, copper redox protein, neisseria, vibrio bordetella, laz, plastocyanin, stellacyanin, detect, diagnosis, fusion protein, heterologous, breast cancer, lung, colorectal, prostate, pancreas, pharmaceutical

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