

# Bidirectional Targeted Killing approach of Genetically Engineered Escherichia coli surface antigen (fimbriae) BNT (Bionanotube) to tumor cell : A Molecular and Engineering Design Concept of Cancer Therapy

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## ABSTRACT

A bidirectional immune response and chemotherapeutic approach is designed on microbial surface antigen, a genetically Engineered Escherichia coli K-12, in which A tumor cell associated immune response will be triggered through GE Escherichia coli K-12 Surface antigen (fimbriae). This fimbriae (pili) is derived through cloning of the said tumor gene into bacteria and BNT derived by the isolated GE fimbriae can be isolated and purified. This BNT (Fimbriae), their nanotube will carry tumor cell killing therapeutic anticancer drugs, cytotoxicity will kill the tumor cells growth. In the recent years we have seen that the trends of microbial engineering emphasizing more towards immune response. We have observed the importance of biological evolution has been shifted to immune responses in ensuring the generations of cancer diversified by various organ specific activities and finally we have observed the functions of stem cells in curing cancer as regenerative medicines, where the questions of regenerations and repairing are still in debate. These repairing mechanisms are confined and are being transferred to microbial engineering. Ten time more bacterial population (10: 1 trillion) cells are existing in a healthy body and same would have been changed due to cancer, where the specific bacterial population will predominate the others to support tumor growth. The author will attempt to change the mode and motivations of bacterial population to a cancer patient, by application of GE E.coli K-12 surface antigen as BNT, immobilized with cancer drugs. In this process a bidirectional approach is proposed to be involved, one is to trigger the immune response (i.e. Lymphocytes T,-B- and Macrophage activities) through BNT and will be specific to kill tumor cells, since the GE bacteria is made by cloning of the specific tumor cells), where immune response is positive but pathogenesis of tumor cells are nil. Second is, since it is genetically engineered, the BNT of the specific E,coli will be more specific to identify the target tumor cell and will deliver the drug. BNT will act both as nanotube to immobilize cancer drug as drug carrier and immune response. The author proposes that this BNT might be more biocompatible compared to CNT (Carbon Nanotube) to prevent side effects Immune response of BNT is identified. However many more work is need to be completed to validate the concepts.

## I. INTRODUCTION

A bidirectional immune response and chemotherapeutic approach is designed on microbial surface antigen, a genetically Engineered **Escherichia coli K-12**. In this process A tumor cell associated immune response will be triggered through GE **Escherichia coli K-12** Surface antigen (fimbriae). [1-3] This fimbriae (pili) is derived through cloning of the said tumor gene into bacteria and BNT derived by the isolated GE fimbriae can be isolated and purified. This BNT (Fimbriae), their nanotube will carry tumor cell killing therapeutic anticancer drugs, cytotoxicity will kill the tumor cells growth. In the recent years we have seen that the trends of microbial engineering emphasizing more towards immune response. The importance of biological evolution has been shifted to immune responses in ensuring the generations of cancer diversified by various organ specific activities. Finally we have observed the functions of stem cells in curing cancer as a part of regenerative medicines. However the questions of regenerations and repairing are still in debate, [4-10]. Nano is 10<sup>-9</sup> m, BNT is Bionanotube and has been conceptualized on the basis of extracted surface antigen of bacteria and is reproduced in figures of SEM (Scanning Electron Microscopy). It is an extract of the GE bacteria of **Escherichia coli K-12** Bacterial protein could be infectious and regenerative. BNT is conceptualized for repairing and bidirectional application in cancer therapy. The mechanisms are confined and are being transferred to microbial engineering. Ten times more bacterial population (10: 1 trillion) cells are exiting in a healthy body of human cells in trillions. 10<sup>-6</sup> cells could be changed due to mutation into a benign and or in a malignant tumor. In cancer the specific bacterial population will predominate the others to support tumor growth and to manage/intensify the opportunistic infections.

Through GE-Fimbriae, and by triggering the immune response the metabolic activities of the major good bacterial population will help to protect the opportunistic infective bacterial population to a cancer patient. GE **E.coli K-12** surface antigen as BNT, immobilized with cancer drugs is proposed for specific bidirectional killing of cancer cell instead of multidirectional random cytotoxic effects of gold, platin, silver nanoparticles based Chemotherapeutic drugs, [11-20].

**II. BIDIRECTIONAL APPROACH**

A bidirectional approach is proposed to be involved in triggering immune cells (i.e. Lymphocytes T, B- and Macrophage) Since a BNT was isolated from GE microbes, which was engineered by cloning of tumor cell DNA and therefore it would be automatically specific to move surrounding tumor cells It will trigger the immune response against the growth of tumor cells. Immobilized drugs inside BNT will release the drug as specific application to tumor and the malignant nature of tumor. Bidirectional ( i.e. it will target the surface antigen of the tumor to bring the drug on the surface of the tumor and at the same time it will generate immune response of the body against said cancer/ tumor. The reason is that the fimbriae (pili) GE- **E.coli K-12** is made through cloning of tumor gene into **E.coli K-12**. According to immunological reaction, this bidirectional approach is unique, compare to any nanoparticle based chemotherapeutic drugs, since they attack not only the tumor cell but also the body surrounding cells.

**III. MATERIALS AND METHOD**

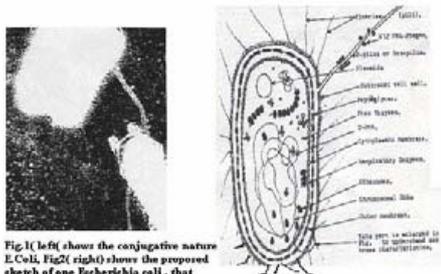


Fig.1 (left) shows the conjugative nature E.Coli. Fig.2 (right) shows the proposed sketch of one Escherichia coli , that carries fimbriae and pili( sex).

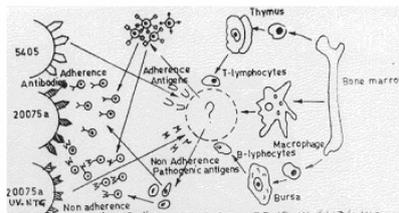
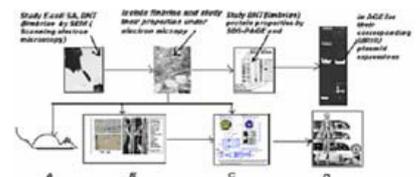


Fig.6. The proposed immune mechanism, which was observed in Bab/C mice by peritoneal inoculum of donor, mutant and hybrid GE fimbriae, carried donor adherence gene in vector plasm id cloning and isolation.



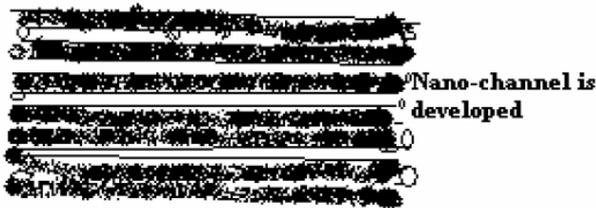
A: Mortality of mice were tested primarily by isolated BNT, by its immune responses against 024:EP/EC donor  
 B: The concept of slow vaccination was conceptualized and therefore scaffold ( matrix based immobilized fimbriae(BNT) applications was thought.  
 C: BNT as nanochannel was pararely conceptualised to validate, that the BNT may be used for slow vaccination, against specific infections. So the nanochanneling properties were observed by BNT enclosed araldite membrane attached with the help of small membrane cassette.....  
 D: Finally the said concept has been perpetuated to design vaccine against Cancer or Oncogenes in case of human trial as future objective.

Figure 1

Figure 2

Figure 3

Steps involved in approaching the model of Bidirectional. Fig.1 represents the SEM view of Escherichia coli and its gene transfer in natural conjugation sex pili, fimbriae (pili) is reproduced, as shown in right, Fig.1. Fig.2, shows the immune response model and Fig.3 describes the steps involved Nanochanneling Design in given below Fig 4.



**Clusture of Fimbriae at the length of 2-5 micron. are used to immobilized cancer drug specific to organ to organs. Fimbriae will do the immune response, since they are genrated in hybrid Escherichia coli K-12 gene cloning. The gene was isolated from the cancer cell.**

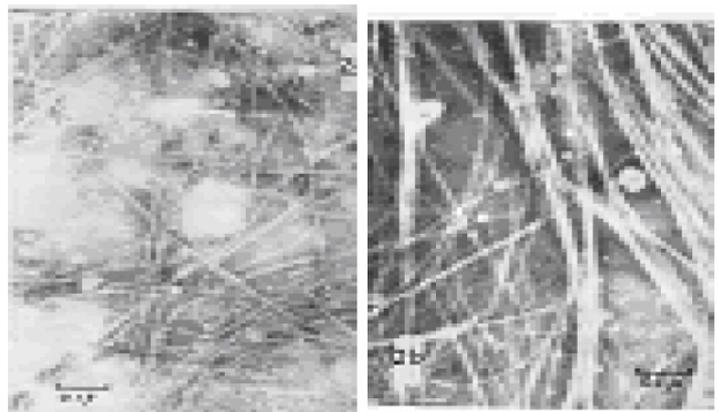


Figure 4

**IV. CONCLUSION**

The Cancer treatments should be carried out through vaccination process, through genetically engineered microbes, carrying the Genes of that particular cancer cells in the form of BNT. BNT would be more compatible to the human body compared to CNT (SWNT and or DWCNT).

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**V. REFERENCES**

- [1] Brahma NK (2012): (a) Design and Control of one Photo-bioreactor; (b) Up and Down Stream in a Chemical and Bioprocess Engineering.- RACET( Recent Advancement of Chemical Engineering and Technology) – Kerala NIT (National Institute of Technology) as paper of AIS- IEI.; and also in Chemcon-2012, Bangalore, AP.
- [2] Brahma NK (2012) Book entitled Introduction to Chemical Science and Engineering, Lap-Lambert- Germany.
- [3] Brahma NK (2013) Bionanotube: A breakthrough Approach in Chemotherapy, Chemcon-2013, Bangalore, AP.
- [4] Brahma NK (2014) Microbial Fuel Cell for safe and clean fuel. Chemcon-2014, Chandigarh, PU.
- [5] Brahma NK (2015) Biosafety and its Importance in Managements of Biological and Chemical Disaster and Terrorism, One Day Seminar (ODS) on Disaster Management against Biological and Chemical Terrorism, IEI/ WBSC).
- [6] Bhar, Basu, Chakrabarty and Brahma (2015), To Study the Activity of Green Sand for Removal of Arsenic and Bacteria from Water, All India Seminar on Recent Developments in Chemical reactors & Reaction Engineering for Security in Energy, Environment, Food and Healthcare. IEI/ EBSC.
- [7] Brahma NK et.al, (2015) Organized ODS, entitled “Chemical Engineering and Biotechnology Process Intensification through Innovative Computation.
- [8] Brahma NK (2016) Organized ODS on Application of Image Processing in Chemical Engineering, Biology and Medicine, IEI/ WBSC.
- [9] Brahma NK (2016) Algae Culture in a Photo bioreactor need Specific Growth, Media, Light Distribution and Operational Aspects of Chemical Engineering, SEED( Synthetic Biology, Engineering, Evolution and Design), Society of Biological Engineering, AIChE USA- Chicago.
- [10] Brahma NK (2016) Plasmid Gene Transfer in vitro into Auxotroph Pheno-Genotypes, F-, Escherichia coli K-12 Generate Hybrid Prototroph to Damage Clonal Selectivity, SEED,. Society of Biological Engineering, AIChE- USA- Chicago.
- [11] Brahma NK (2016) Bionanotube and Regenerative Medicine: A New look to Molecular and Engineering Design, 31 Indian Engineering Congress, Kolkata.
- [12] Brahma NK and Basu S, (2017) Understanding the Paradigm Shift of Diagnosis (Environment, Medical and Social Changes) Through Electronic Media (Bioinformatics, E-telemedicine) Digitization (Imaging Medical MI). Genomic and Regenerative Sciences and by Increasing Human Relations. IEI Technorama.
- [13] Brahma NK (2016) Haemoperfusive Membrane Encapsulated Activated Charcoal Packed Column Artificial Kidney in Replacing Haemodialysis, SEED, Society of Biological Engineering, AIChE (American Institute of Chemical Engineering), USA- Chicago.
- [14] Brahma NK (2016) Organized and Presented Paper, Learning LCA &EIA for Climate Change Through Process Intensification, Genetic Engineering and Nanotechnology, AIS (All India Seminar) entitled “Environmental Impact Assessment of Chemical and Allied Industries for Sustainable Development and Climatic Change”. IEI/ WBSC.
- [15] Brahma NK (2017) Molecular and Engineering Design Concept of Bacterial Surface Antigen (Fimbriae) involved in Cancer Therapy, SEED, Society of Biological Engineering, AIChE, Canada Vancouver.
- [16] Brahma NK (2017) Bidirectional Targeted Killing Approach of Genetically Engineered Escherichia coli Surface Antigen (fimbriae), BNT (Bionanotube) to tumor cell: A Molecular and Engineering Design concept of Cancer Therapy, Nanosmat-UK, Hong-Kong City University.
- [17] Brahma NK (2017) Book, Entitled “Molecular and Engineering Concepts of Microbiology, Lap-Lambert-Germany, USA.
- [18] Brahma NK (2017) Organized ODS, entitled “Advancement of Chemical Engineering and Biotechnology in Securing Health, Energy and Food, and paper entitled “Paradigm Shift of Biology” IEI/ WBSC.
- [19] Brahma, NK (1999) Genetically Engineered Hybrid 5405 E.coli K-12 fimbriae in vaccination With Balb/c mice against O26: serotype EPEC diarrhea. Ind. J.Chem. Engg. 39 (1) p: 17-, Cited in British Journal catalogue (BJC)-UK.
- [20] Grireshawr M, Sarkar D and Brahma NK (2004) Simulation of HIV Infection in Human Immune system as Diagnostic Tool, Ind. chem. Engg.46 (1):P; 13-20. Cited in BJC-UK.