### Frank Reidy Research Center for Bioelectrics

Dr. Richard Heller



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**Center for Bioelectrics** 



## Reidy Center for Bioelectrics

### University Level Research Center

- Founded in 2002
- 45 Researchers: Faculty, Post-docs, students, technicians and staff
- Leader in an International Consortium on Bioelectrics
  - Japan, Germany, France, Czech Republic, Slovenia and United States
- Over \$3 Million in Annual Research Expenditures
  - Research sponsors include National Institutes of Health, US Department of Defense, National Science Foundation, Commonwealth of Virginia and Private Industry



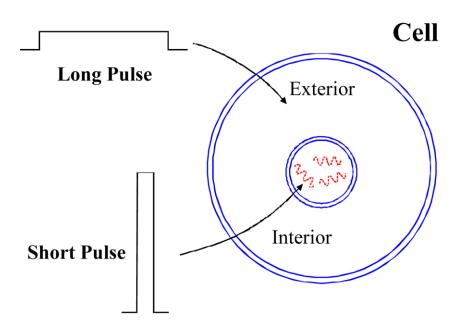
### Reidy Center for Bioelectrics

Report by "Nature.com" - January, 2011.

"The integration of physical sciences and engineering with biology is considered to be revolutionary, and will transform everything from health care to energy production to food."

The Center for Bioelectrics is an example of such a revolutionary approach. By bringing together scientists with diverse backgrounds, but complementing expertise, the bioelectrics team has positioned itself as an internationally recognized leader in this rapidly growing field.

### Pulsed Power: Affect Cell Functions [from delivery of molecules to release of calcium and induction of apoptosis]









# **Bioelectric Applications**

- Cellular interactions
- Wound healing
- Plasma medicine
- Cancer treatment
- Detection
- Cardiovascular
- DNA vaccines
- Neurological Applications



## **Bioelectric Applications**

### Wound Healing

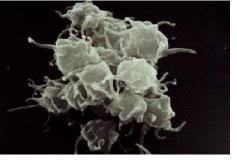
- Platelet gels nanosecond pulse electric fields to activate platelets
- Gene therapy delivery of plasmids encoding angiogenic factors to accelerate wound healing

### Using Pulsed Power to Activate Platelets for Wound Healing

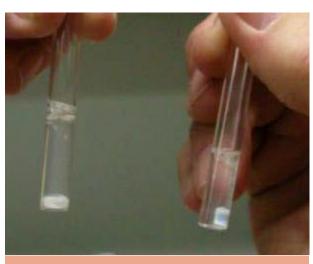
Platelets



**Before Activation** 



After Activation



### Bovine Thrombin is used in current clinical trial

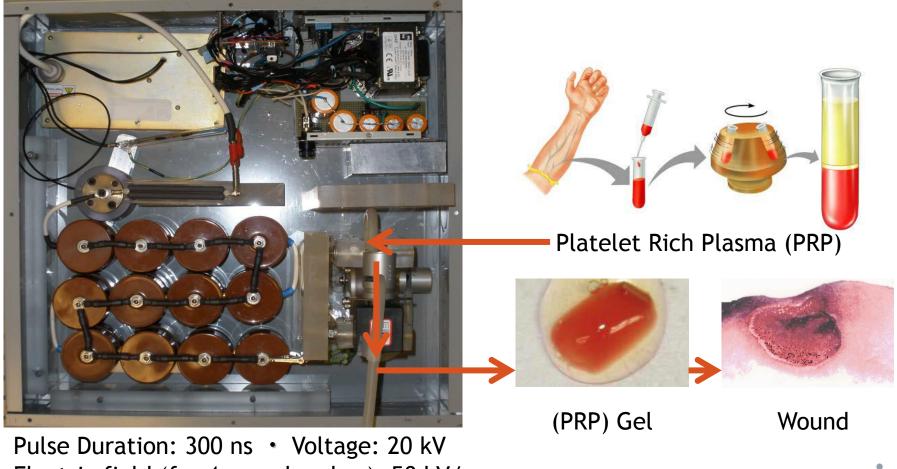
**Problems of Using Bovine Thrombin** 

- possible allergies and complications
- potential risk of contamination of Creutzfeldt-Jakob's disease (Mad-cow disease)

Thrombin nsPEF activated gel activated gel



### **Compact Pulsed Power Generator for Platelet Gel Production in Doctor's Office**



Electric field (for 4 mm chamber): 50 kV/cm Repetition rate: 3 Hz



# Platelet Gel for Wound Healing (near term) & Ischemia (mid-long term)

- Dental surgery, orthopedics and plastic surgery
- Gel Advantages accelerated healing, less bleeding (enhanced hemostasis) and bruising, less pain, lower risk of infection, less chance of fluid collection (seroma)



Plasmid = Vascular Endothelial Growth Factor



Ferraro, et al Mol Ther. 17:651, 2009



### Gene Therapy for Wound Healing (mid-long term)

- Electroporation type pulses
- Dependent on factor delivered
- Utilizing gene transfer to the skin of an angiogenic factor
- Utilized for large defects flaps



## **Bioelectric Applications**

### ✤ Plasma

- Use of cold plasma to destroy bacteria
  - Prevention of wound infections
  - Decontamination of food
  - Decontamination of surfaces



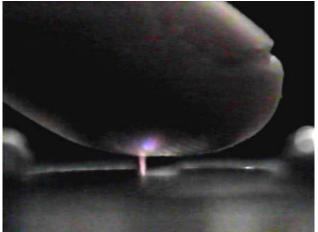
A. baumannii



P. aeruginosa



S. aureus

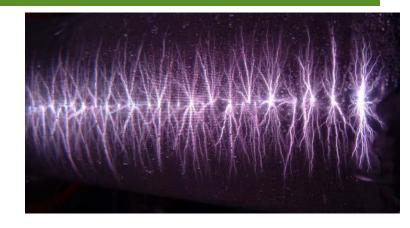


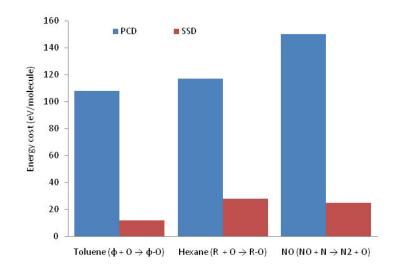
### <u>(</u>)

### **Bioelectric Applications**

### Plasma

- Other applications
  - Removal of nox
    from diesel exhaust
  - Air purification
  - Fuel reforming
  - Extraction of radioactive tritium







### Cold Plasma (near-mid term)

- Destroy microorganisms including pathogenic bacteria
- Decrease chance of infection in wound healing
- Hand sanitizing or other surfaces
- Disinfecting rooms hospitals, schools
- Dentistry applications
- Decontamination of equipment
- Decontamination of food



# Plasma Streamers with or without steam (near-mid term)

- Remove nox from diesel exhaust
- Air purification
- Fuel reforming
- Extraction of radioactive tritium

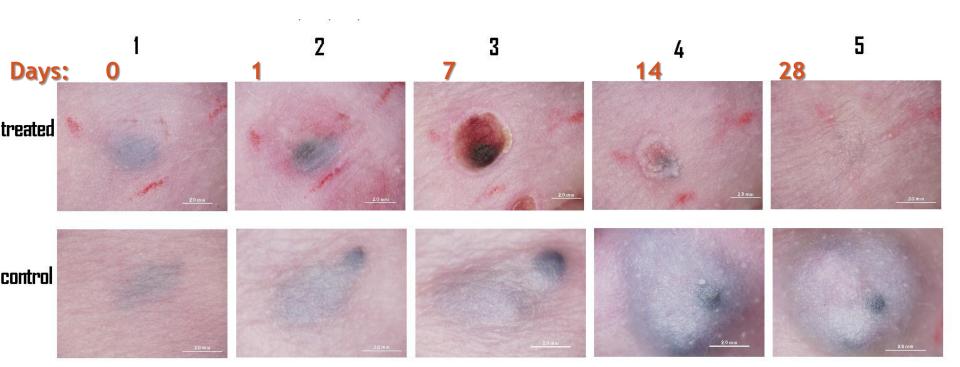


## **Bioelectric Applications**

### Treatment of Cancer

- Ablation therapy nanosecond pulse electric fields to destroy tumor cells - work performed in melanoma, squamous cell carcinoma, liver cancer, pancreatic cancer and breast cancer
- Gene therapy micro-millisecond pulse electric fields to deliver plasmid DNA to stimulate immune system. Phase I clinical trial successfully completed. Phase II trial initiated.

### NsPEFs Decrease B16F10 Tumor Size





### Electrogene Transfer of pIL-12 to Melanoma Patients

Phase I does escalation study (0.1 - 1.6 mg/ml) to evaluate safety and tolerability of approach. Secondary objective is to evaluate response.

Patients with Stage III or IV malignant melanoma with cutaneous lesions. To enroll a patient must have at least two lesions and up to 4 can be treated.

Patients receive 3 treatments (Days 1, 5 and 8). 1300 V/cm 100



### Results

Safety and tolerability

- No serious adverse events related to therapy
- Blood work all within normal ranges
- No unanticipated adverse device effects
- All subjects enrolled completed therapy

### Response

- Increased levels of IL-12 following treatment
- Histological evidence of tumor necrosis and lymphocytic infiltrate
- 19 patients with additional disease
  - 3 had a complete response
  - 2 have long-term stable disease
  - 3 had stable disease for 4-6 months



Daud, et al, J Clin Oncol, 2008





### Cancer Therapy

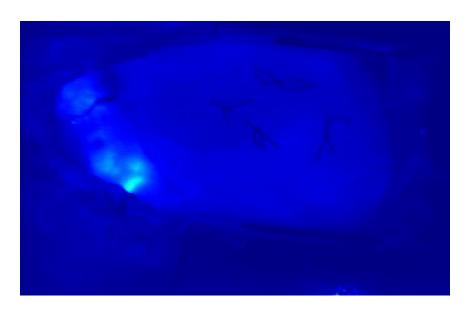
- Ablation (near-mid term)
  - Nanosecond pulse electric fields induce apoptosis
  - Any solid tumor
- Gene Therapy (mid term)
  - Deliver genes encoding immune modifiers
  - Induce a systemic response from local treatment
  - Potential for most metastatic cancers

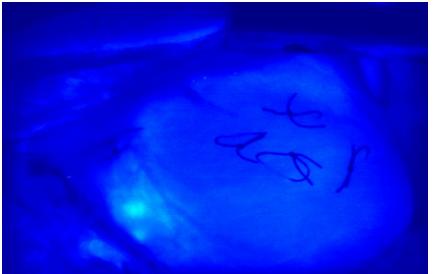


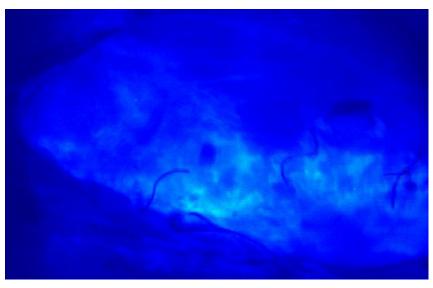
## **Bioelectric Applications**

### Cardiovascular

- Coronary artery disease gene therapy to assist revascularization
- Peripheral vascular disease gene therapy to assist revascularization



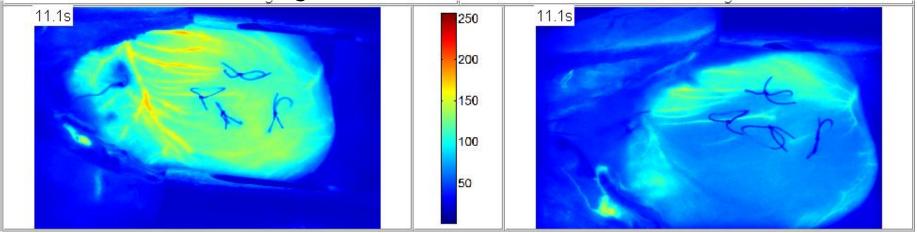




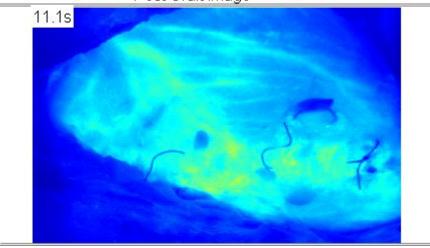


#### Pre Occlusion Image

Post Occlusion Image



### Post Treatment Image







### Gene Therapy for Wound Healing (mid-long term)

- Electroporation type pulses (long pulses)
- Dependent on factor delivered
- Peripheral vascular disease
- Coronary artery disease

### **Center for Bioelectrics - Future**

# Electric/electromagnetic fields are powerful tools

- Tremendous translational opportunities
- Multiple new applications

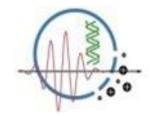
## Expansion

 Recruit new faculty - target plasma, cancer biology, immunology, cardiovascular and neuroscience expertise









Frank Reidy Research Center for Bioelectrics

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