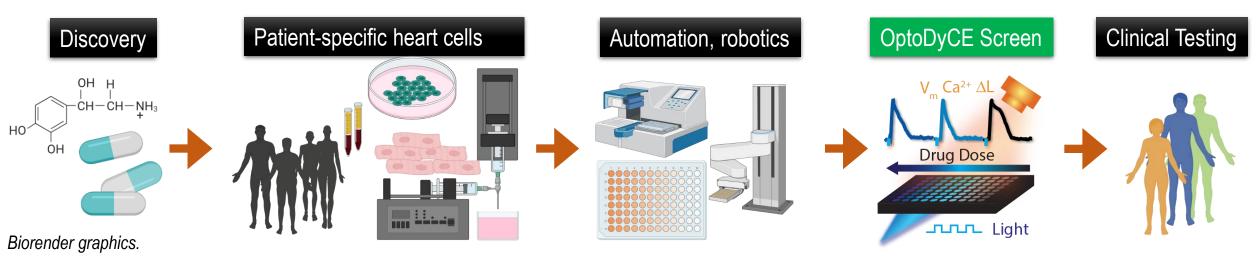
# **SPARKBio**

high-throughput optical cardiotoxicity screening

**Developed by The George Washington University** 

# **Technology Summary**

- First scalable automated all-optical system for high-throughput cardiac toxicity screening (OptoDyCE).
- Optically measures electrophysiology of a variety of primary and stem-cell-derived cells and tissues.
- Compatible with proprietary "spark-cell" spheroids: easy to manipulate, store and deploy for immediate use.
- No genetic modification of the studied cells of interest needed; no viral delivery needed; fast deployment.



Klimas, A. et al. Nat. Commun. 7, 11542 (2016) Klimas A et al. Prog Biophys Mol Biol, 154:62-70 (2020)

OptoDyCE is applicable to various experimental models for drug discovery and cardiotoxicity screening.

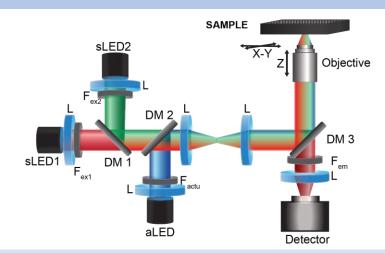
## **Problems with Drug Development Pipeline**

# Cardiotoxicity is one of the leading causes for discontinuation at different phases of drug development.

- Current methods of pre-clinical cardiotoxicity testing are slow, expensive, and lack the complexity of in vivo cardiomyocytes.
- There are no scalable and high-throughput compatible cellular cardiac electrophysiology assays in the market.
- Current automated systems can't characterize tissue-level/multicellular effects.

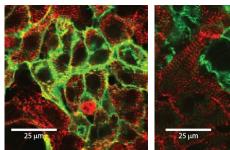
#### **Our Solution**

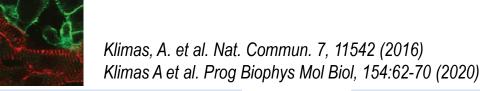
#### A Device and Method Providing Multi-Parameter, High-Content Information



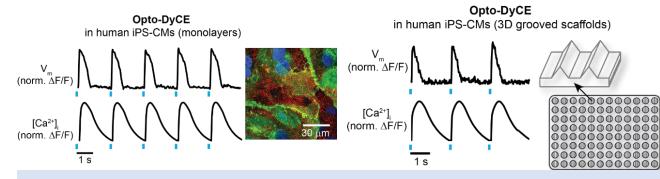
Light-based device provides simultaneous multiparameter control and measurement of hiPSC-CMs.

#### Gene Delivery Modular Delivery

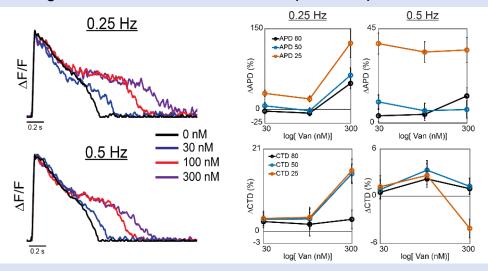




Light sensitivity (green) imparted on CMs (red) using gene delivery or "spark" cells.



Physiologically relevant, multi-parameter data can be obtained in different tissue configurations within 10 seconds per sample.

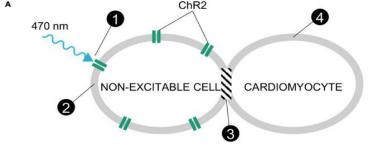


Pro-arrhythmic effects of a drug (Vanoxerine) recently removed from Phase III clinical trials were easily revealed using the OptoDyCE Platform.

System can dynamically interrogate 10,000 samples per day (20 years of patch clamp man-hours) and provide a more complete profile of potential cardiotoxic effects compared to traditional methods.

#### **Our Solution**

#### OptoDyCE-compatible "Spark-Cell" Spheroids for Optical Pacing of Cardiac Tissue



- ChR2 SPHEROID hiPS-CM SYNCYTIA ELECTROPHYSIOLOGY

- "Spark-cell" spheroids are scalable and cost-effective for contactless optical stimulation of cardiac cell constructs for high throughput drug screening.
- Confer optical responsiveness of cardiac tissue earlier than direct viral or liposomal genetic modification of the cardiomyocytes.
- Offer multimodal responsiveness optical, magnetic, ultrasound for navigation and actuation.



Chua C et al. Front Bioeng Biotechnol; 9:658594 (2022).

"Spark-cell" spheroids can be manufactured in a high-throughput (HT) format, and can be positioned onto hiPSC-cardiomyocytes, also grown in HT format

Chua C et al. Front Bioeng Biotechnol; 9:658594 (2022).

"Spark-Cell" spheroids can be frozen, transported, and quickly deployed on-site to confer optical pacing of cardiac cells

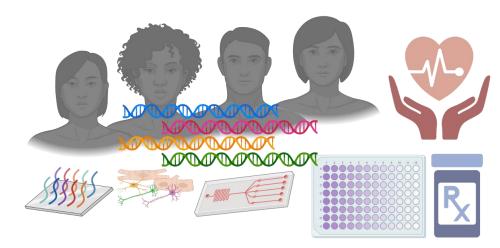
### **Target Customers**

### Industry

- High-throughput screening developers (Nanion, Axion Biosystems, Q-State Biosciences)
- Pre-clinical CROs doing cardio toxicity screening (Eurofins, Cardiomex, Cyprotex, etc.)
- Pharma companies (Bayer, Novartis, Bristol-Myers Squibb, Johnson & Johnson etc.)
- Biotech, gene therapy developers
- AI, machine learning companies for personalized medicine (In Silico Medicine)

### Research Institutions

- Academic labs
- University core facilities



## What makes SPARKBio unique?



- Allows precise optogenetic stimulation without genetic modification of cells
- Allows all-optical electrophysiology-simultaneous voltage, calcium optical measurements
- Safer for cells due to low intensity and long wavelength light used
- Spheroids respond to electrical, optical, magnetic, and ultrasound signals
- Compatible with standard high-throughput plate format (96- or 384-well plates)

# **Competitive advantages**

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		Patch Clamp	MEAs	Sciences	Q-State	OptoDyCE
General	Low Cost					✓
	High Throughput		✓	✓	✓	✓
	High Sensitivity	✓	✓	✓	✓	✓
	Modular					✓
	Multi-cell recording		✓	✓	✓	✓
	Contactless Actuation				✓	✓
	Contactless Recording			✓	✓	✓
Assay	Voltage	✓			✓	✓
	Calcium			✓	✓	✓
	Contraction					✓
Sample Type	Monolayer		✓	✓	✓	✓
	3D Structure					✓
	Single Cells	✓	✓	✓	✓	✓
	Human Cells	✓	✓	✓	✓	✓
Study Type	Cardiotoxicity	✓	✓	✓	✓	✓
	Chronic Study		✓	✓	✓	✓
	EC-Coupling					✓
	Mechanistic Screen					✓
	Lead Screening		✓	✓	✓	✓

### **Innovator**

### **Emilia Entcheva, PhD**

- Director of the Cardiac Optogenetics and Optical Imaging lab at the George Washington University
- PhD in Biomedical Engineering
- Pioneer of cardiac optogenetics and optical methods
- Extensive knowledge of cardiac electrophysiology experiments and computer modeling



### What does SPARKBio need?

- What we are looking for?
  - Serial entrepreneur with potential for CEO role
  - CSO/CTO
  - Investment / Financing
- Is there a market for SPARKBio's technology?
  - Yes. <u>Cardiotoxicity alone accounts for >30% of drugs withdrawn</u>. To avoid latestage drug attrition, pharmaceutical industries needs high-throughput platforms to screen drug-induced cardiac toxicity in vitro.
- What is the current development state of SPARKBio's technology?
  - Patents filed
  - Have fully-functional prototype