



INTRODUCTORY OVERVIEW AND PRESENTATION

SIGNPATH PHARMA INC.

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OVERVIEW

- *SignPath Pharma is a clinical stage biotechnology company. It focuses on creating heart safe versions of drugs which cause cardiotoxicity.*
- *SignPath's proprietary technology provides a solution to the most serious drug safety issue in the world (cardiac safety.)*
 - Funding to date:
 - Raised upwards of \$17 million through several “friends and family” rounds
 - No outstanding debt
 - Broad IP portfolio with many issued patents prevents competition
 - Delaware C Corp
 - Operates as an efficient “Virtual” company using outsourcing and contracted labs



PROBLEM: DRUG INDUCED CARDIAC SIDE EFFECTS

➤ *Arrhythmia caused by drugs:*

- **Leading cause of failure at every stage of drug development**
- Over 150 FDA approved drugs with this problem, including anti-cancer agents, antibiotics, antipsychotics, antihistamines and other drugs. Many of these drugs are dose limited by cardiac safety.
- Numerous pipeline drugs are stalled due to cardiac toxicities. Estimated that over 1/3 of drugs in development fail due to cardiac safety.

➤ *Cardiomyopathy caused by cancer chemotherapy:*

- Cardiac toxicity that can permanently damage the heart muscle and damage to other organs
- Prevents proven effective chemotherapy treatments from continued use
- Cardiomyopathy leads to reduced cumulative dose or temporary cessation of chemotherapy for many patients.



CARDIAC ANSWER: SPP4040

- *SPP4040 has prevented arrhythmia more than 35 well known problem drugs*
- *There is a 90% correlation between the animal models and human results for drugs impacting QT interval. Scientific risk is low.*
- *SPP4040 has shown protection against cardiotoxicity with chemotherapy regimens that otherwise result in significant cardiomyopathy.*
- *SPP4040 has been through several development iterations to achieve:*
 - Oral administration
 - Non-toxic even at high dosage (chronic dosing at 800x clinical dose)
 - Does not reduce or impact efficacy of target drug
 - Effective within minutes
 - 6 to 8 hour effectiveness after single dose
 - 100mg dosage for adult male. This will allow co-formulation with target drugs



SPP4040 MARKET POTENTIAL

- *Very large market of drugs which are limited by cardiac arrhythmia*
 - More than 150 FDA approved drugs with arrhythmia problems
 - Hundreds more drugs stalled in development because of arrhythmia
- *The potential addressable market is tens of billions of dollars. Examples include:*
 - Risperidone: Peak Sales \$4.7 billion
 - Citalopram: Peak Sales \$1.7 billion
 - Escitalopram: Peak Sales \$2 billion
 - Ondansetron: Peak Sales \$1.1 billion.
- *The chemotherapy cardiomyopathy market is tremendous. Many chemotherapy regimens are cardiotoxic. Chemotherapy drugs which cause cardiomyopathy include:*
 - Sutent
 - Herceptin



SPP 4040 MARKET POTENTIAL continued

- *SignPath's anti-arrhythmia strategy is to create second generation "heart safe" versions of drugs with proven efficacy.*
- *Plans are to co-formulate and license SPP4040 with problem drugs. This maximizes potential revenue by:*
 - Saving drugs with markets limited by cardiotoxicity
 - Extending patent life of marketed drugs
 - Entering license agreements with drug companies developing drugs which are experiencing cardiac safety concerns.
- *SignPath can also license its proprietary CorreQT technology to companies who are facing obstacles in developing pipeline drugs due to cardiac safety issues.*
- *SPP4040 clinical trials to begin in 2020.*



LIPOCURC™ (LIPOSOMAL CURCUMIN)

- *LipoCurc is Signpath's first clinical application of its technology to address cardiotoxicity.*
 - Curcumin pro-arrhythmic side effects eliminated.
 - No cardiotoxicity observed in 70 human patients.
- *LipoCurc has unique mechanisms of action, which allow for single agent activity, and synergy with other anti-cancer drugs.*
 - Clinical activity in human cancer
 - Kills cancer stem cells
 - Counteracts cancer related immunosuppression
 - Crosses the blood/brain barrier which allows for treatment of brain tumors
 - Observed efficacy in end-stage cancer patients who failed 6-7 prior treatments.
 - Accumulates preferentially (500x) in cancerous blood cells, which allows it to target blood cancers.
- *LipoCurc has minimal toxicity.*



LipoCurc—PHASE II CLINICAL TRIAL TARGETS

- **Glioblastoma** This will be done at Johns Hopkins. Market size is over \$3 billion despite the fact that there is currently no effective treatment
- **Mesothelioma** The Australian government is funding 100% of costs for Phase II testing. (Flinders University in Adelaide.) Market size is currently \$340 million (no effective treatment)
- **Multiple Myeloma** Phase II site not yet finalized. Market size is expected to be \$38 billion in five years
- **Soft Tissue Sarcoma** Phase II trials to be held at University of Nebraska. The global market size is expected to be \$1.2 billion by 2023.



INTELLECTUAL PROPERTY PROTECTION

- *Extremely Broad Patent Protection:*
- Numerous issued and pending patents protecting cardiac arrhythmia and cardiomyopathy protection.
 - Patent position prevents competition in the field of mitigating arrhythmia using lipids and empty liposomes
 - Broad use patents, and specific composition of matter patents
- Patent protection on LipoCurc, with both composition of matter and method of use patents issued and pending.



CURRENT STAGE AND PRODUCT DEVELOPMENT

➤ *SPP4040 Progress To Date:*

- Proven successful with more than 35 drugs in animal testing
- Animal studies show no toxicity
- Planned trials will be relatively simple, quick, and inexpensive

➤ *LipoCurc Progress To Date:*

- Completed phase I trials in 70 humans.
- End stage cancer patients who had failed 6-7 prior treatments showed dramatic decreases in their cancer markers and clinical benefit.



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