

# Cautionary Note Regarding Forward Looking Statements

This presentation of PolyPid Ltd. (the "Company") contains "forward-looking statements" within the meaning of the Private Securities Litigation Reform Act and other securities laws. Words such as "expects," "anticipates," "intends," "plans," "believes," "seeks," "estimates" and similar expressions or variations of such words are intended to identify forward-looking statements. For example, the Company is using forward-looking statements when it discusses statements relating to our objectives, plans. and strategies, the expected timing of trials, the research, development, and use of our platform technologies, technologies, products and product candidates, potential benefits and advantages of our products and product candidates, and all statements (other than statements of historical facts) that address activities, events, or developments that the Company intends, expects, projects, believes, or anticipates will or may occur in the future, expected timing of completion of patient recruitment and top-line results of the SHIELD II study and US addressable market. Forward-looking statements are not historical facts, and are based upon management's current expectations, beliefs and projections, many of which, by their nature, are inherently uncertain. Such expectations, beliefs and projections are expressed in good faith. However, there can be no assurance that

management's expectations, beliefs and projections will be achieved and actual results may differ materially from what is expressed in or indicated by the forward-looking statements. Forward-looking statements are subject to risks and uncertainties that could cause actual performance or results to differ materially from those expressed in the forward-looking statements. For a more detailed description of the risks and uncertainties affecting the Company, reference is made to the Company's reports filed from time to time with the Securities and Exchange Commission ("SEC"), including, but not limited to, the risks detailed in the Company's Annual Report on Form 20-F, filed with the SEC on March 6, 2024. Forward-looking statements speak only as of the date the statements are made. The Company assumes no obligation to update forward-looking statements to reflect actual results, subsequent events or circumstances, changes in assumptions or changes in other factors affecting forward-looking information except to the extent required by applicable securities laws. If the Company does update one or more forward-looking statements, no inference should be drawn that the Company will make additional updates with respect thereto or with respect to other forward-looking statements.



# PolyPid Overview

Late clinical stage biopharma company

Unique LOCAL Prolonged Delivery of APIs Polymer-Lipid Encapsulation matriX (PLEX) Platform

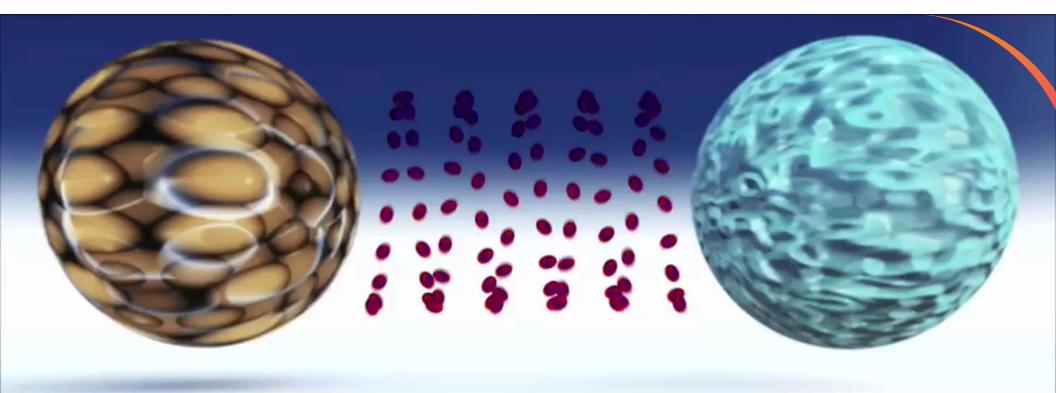
Lead Product D-PLEX<sub>100</sub> in Phase 3 trial

OncoPLEX Next Big Opportunity for Solid Tumors 176
granted and pending patents(1)

65 employees(1)

HQS
Global: Petach Tikva, Israel
US: New Jersey

**NASDAQ: PYPD** 



**LIPIDS** 

**POLYMERS** 



# Robust Pipeline with Multiple Near- and Longer-Term Inflection Points

Product candidate and indication	Preclinical	Phase 1	Phase 2	Phase 3	Key milestones
D-PLEX <sub>100</sub> Prevention of Surgical Complications  Prevention of SSI in Colorectal Abdominal  Prevention of SSI in Abdominal Surgery  Prevention of SSI in Orthopedic Surgery		abdominal colorectal surg		PK + Safety Study  Post-Approval Efficacy Study	SHIELD I study completed  SHIELD II Topline expected by 2Q 2025  2026  2027/2028
OncoPLEX  Post Surgical Tumor Resection (Adjuvant)  Intratumoral Solid Tumors (Neoadjuvant)					Pre-IND meeting completed (FDA) for GBM Pre-clinical stage



# D-PLEX<sub>100</sub> is a Potential First-in-class for the prevention of SSIs

Indication:

Prevention of abdominal incisional SSI

Doxycycline (broad spectrum antibiotic)

FDA 505(b)(2) regulatory pathway

Administered directly into the surgical site for prolonged 30 days release

~12M Surgeries addressable market in US

Breakthrough Therapy, Fast Track, and QIDP designations

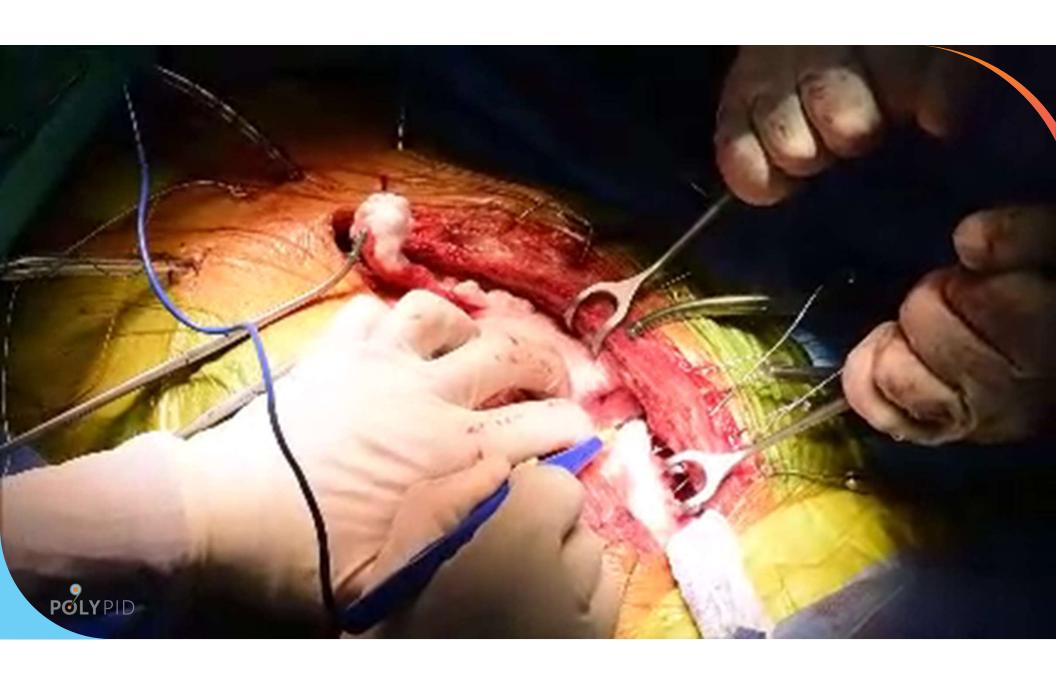
~12M procedures
Orthopedic 5.0M
Abdominal 4.5M

Gyn & Uro **2.5M** 

Sternum **0.5M** 

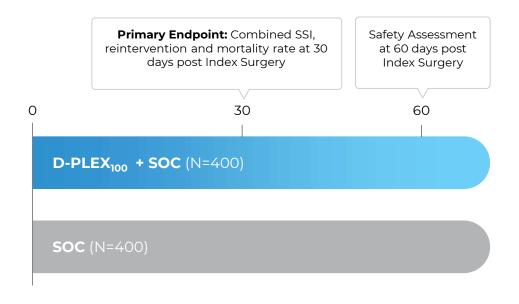






# Study Design and Timeline





#### Surgeries with large surgical incision

Total of 800 patient trial. Sample size is based on Interim Analysis review of the data by the Data Safety Monitoring Committee (DSMB)

#### **Current timing assumptions**

- Last patient in: Q1 2025
- Topline results: Q2 2025

#### Actions taken to de-risk SHIELD II

- Focused on population where SHIELD I was successful
- Conservative statistical assumptions on SSI rates
- Implemented lessons-learned: performed detailed debriefing with the site PIs, kept only high-performing sites
- Strengthened clinical ops team



The Interim Analysis is an Opportunity to Re-size the Trial and Ensure it has Sufficient Power to Confirm Treatment Benefit

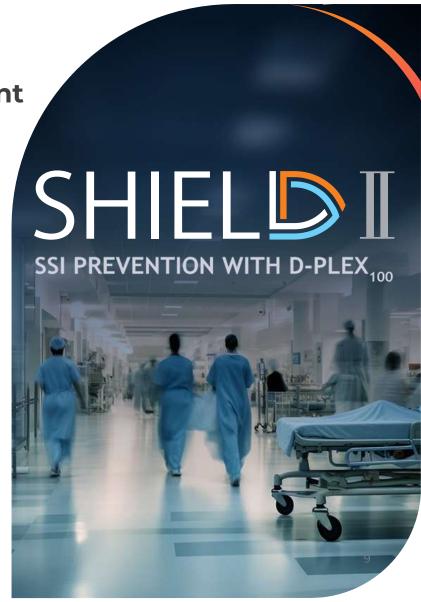
Unblinded interim efficacy analysis by DSMB at 430 patients recommended expanding the sample size to 800 patients - **the first stop after the minimum planned number of patients** 

We view this DSMB's recommendation as a **very favorable outcome**, as it is suggestive of **positive efficacy signals** from D-PLEX $_{100}$ 

The study is now **significantly de-risked** - futility option is now removed

**No safety issues** related to D-PLEX $_{100}$  have been observed in SHIELD II to date



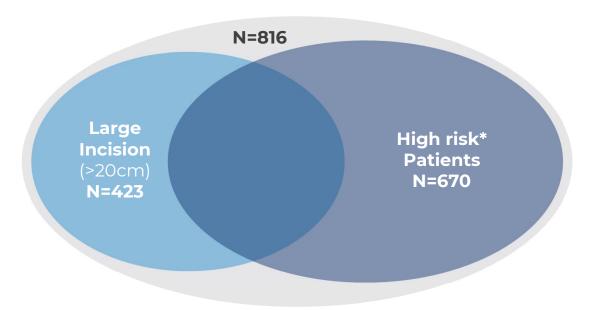


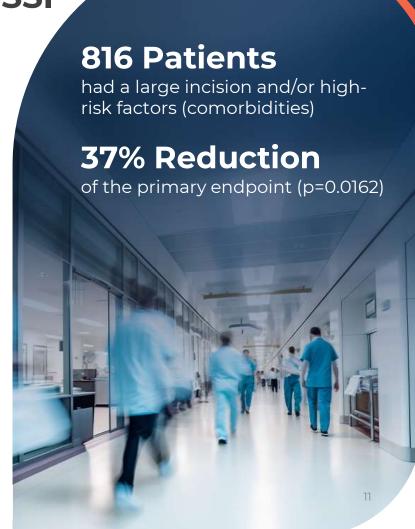
# SHIELD I: Deep Dive into the Large-Incision Subgroup

Parameter	<b>D-PLEX</b> (N=212)	Control (N=211)	Effect
Primary endpoint	17 (8%)	37 (17.5%)	<b>54%</b> (p=0.0032)
Key Secondary Efficacy Endpoints			
Infection rate during 30 days post abdominal surgery	9 (4.4%)	19 (9.7%)	55%
Number of subjects with at least 1 score of ASEPSIS >20	2 (1.0%)	5 (2.6%)	62%
Additional Efficacy Endpoints			
Incidence of SSSI rate during 30 days post surgery	9 (4.4%)	17 (8.7%)	49%
Incidence of DSSI rate during 30 days post surgery	0	2 (1.0%)	100%
Mortality rate within 30 days post abdominal surgery	6 (2.8%)	10 (4.7%)	40%
Time to adjudicated SSI during 30 days post index surgery (days)	8.0 (4, 28)	5.0 (1, 13)	NA
Number of subjects treated with IV Antibiotic as treatment for adjudicated SSI	1 (11.1%)	9 (47.4%)	77%
Number of subject with any surgical re-interventions	9 (4.4%)	19 (9.7%)	55%



D-PLEX<sub>100</sub> Effect on in Patients w/ SSI Risk Factors\*







\* Post-hoc analysis; patient related risk factors include BMI >30, smoking/COPD, diabetes, hypertension



**12M 8M** Europe total addressable US total addressable market in-patient market in-patient surgical procedure surgical procedures **4.4M Abdominal Surgeries** Abdominal Surgeries •7,400K\*\*← Core Target Surg. Procedures

<sup>\*</sup> Assuming additional safety and PK Study for US ; Expected Abdominal Indication in Europe based on SHIELD 2 phase 3 trial

<sup>\*\*</sup>Source IQVIA PM&I Global FlexView. Internal analysis

# Demonstrated Economic Benefits will be Essential for Market Access and Sales Uptake

#### **Direct cost**

SSI costs ~\$25K/patient<sup>1</sup> on average

- Prolonged length of stay and higher readmission rates
- Re-operation in some cases (to debride and remove infected / necrotic tissues)

#### Indirect cost

CMS 1-3% penalty on all the yearly Hospital Medicare reimbursement

### **Reputational cost**

Hospital SSI rates are public information and have direct influence on hospital ranking by CMS and U.S. News best hospitals ranking



1. Stone PW. Economic burden of healthcare-associated infections: an American perspective. Expert Rev Pharmacoecon Outcomes Res. 2009 Oct;9(5):417-22.



# **Global Go-to-market Strategy**

Partnerships with leading pharma companies with established hospital-focused commercial capabilities and resources

### Agreement highlights



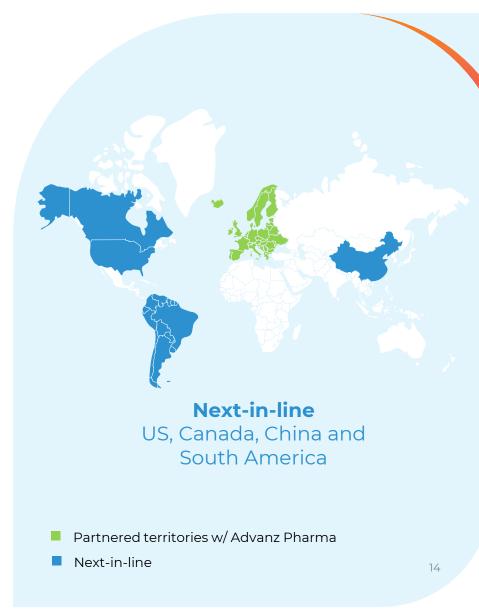
Includes European Economic Area and UK

Potentially receive over \$115 million in upfront and milestone payments as well as royalties on net sales

\$2.7 million upfront payment paid upon signature of licensing agreement Focused on abdominal and cardiac indications

Signed licensing agreement includes transfer price, development and sales-related milestone payments and royalties

Development-related milestones for a total of up to \$25 million





# R&D Collaboration Agreement with Immunogenesis for Controlled and Prolonged Intratumoral Delivery

- Houston-based company founded by Dr. Michael Curran, Associate Professor of Immunology at the MD Anderson Cancer Center
- Combined Immunogenesis potent anti-tumor STING agonist with PolyPid's technology to enhance treatment for solid tumors
- Aim to overcome shortcoming of STING agonists molecules, which are susceptible to rapid clearance





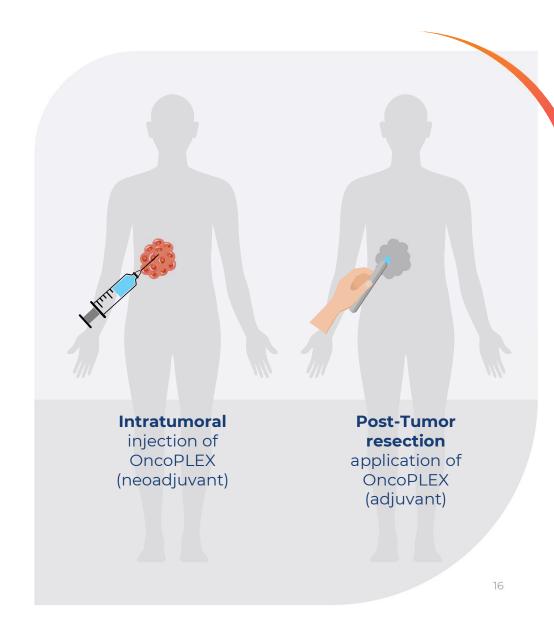
# OncoPLEX - The Next Big Opportunity @ PolyPid

New approach for solid tumors - every year, 1.6 million new cases of solid tumors in the U.S. alone

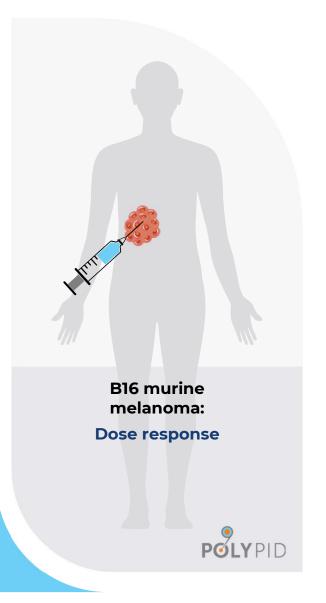
Prolonged 3wks release of docetaxel: intratumoral (neoadjuvant) and post surgical resection of the tumor (adjuvant)

Evaluated successfully in various animal models

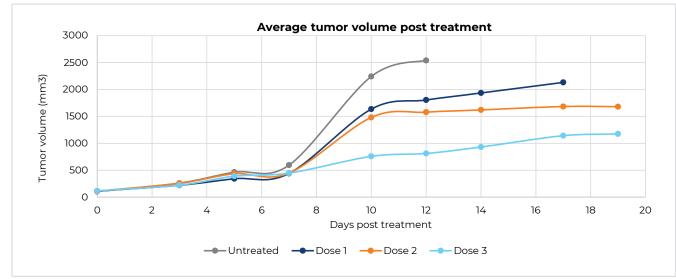
Pre-IND meeting (FDA) completed for GBM





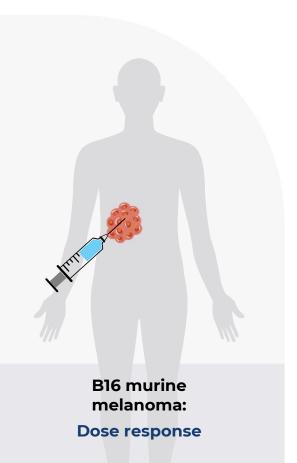


# Single Intratumoral Injection of OncoPLEX reduced tumor growth

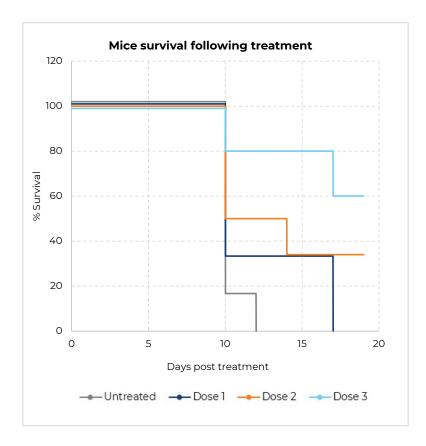


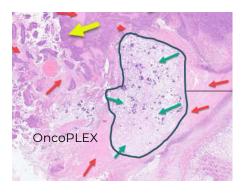
#### **Key Takeaways**

- · OncoPLEX spheres remain anchored to the injection site over the entire period
- The prolonged and constant release mechanism allows the released drug to generate an effective microenvironment far from the injection site

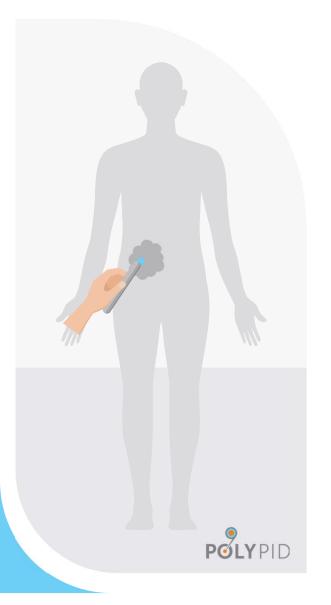


# 60% Survival at Day 19 for the Most Effective Dose



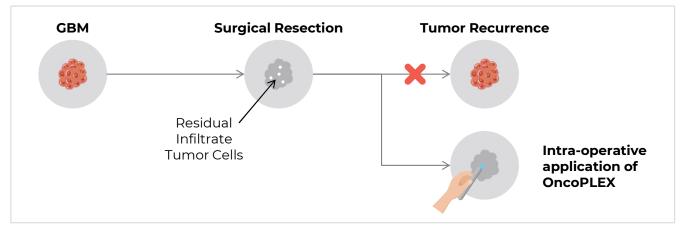


OncoPLEX Focal Deposits are mostly Surrounded by **Necrotic Tumor Tissues** and Inflammation



# Post-Surgical Resection in GBM & Other Solid Tumors

- Prolonged 3wks. local release of Docetaxel directly in the tumor resection pocket
- Evaluated successfully in various animal models
   75% overall tumor free survival in resected colon carcinoma tumor
- 98% tumor growth inhibition (day 41) in resected GBM tumor mouse model compared to the untreated control (p<0.001)</li>
- 60% survival (day 41) in resected GBM tumor mouse model vs 20% for the systemic treated mice (p=0.0165)
- 75% overall tumor free survival in resected colon carcinoma tumor mouse model compared to 25% for the systemic docetaxel arm
- Pre-IND meeting in GBM completed (FDA)



# State-of-the-Art Manufacturing Facility

PolyPid was granted Manufacturer Authorization and Good Manufacturing Practice – **GMP - certification** by Israel's MoH and EU qualified person for its state-of-the-art ~18,000 square feet (~1,700 m²) manufacturing facility.

### **Investment**

machinery, qualifications and validations

### **Supply capacity**

expected to meet commercial demand for the first 4-5 years from launch









Stock Information			
Listing	NASDAQ		
Ticker	PYPD		
52-week range <sup>1</sup>	\$2.95-\$9.20		
Market cap <sup>1</sup>	\$32 M		
Share Structure			
Shares outstanding	10.2 M		
Pre-funded warrants	1.6 M @ \$0.0001 exercise price		
Warrants	1.7 M @ \$3.61 exercise price		
Warrants	6.7 M @ \$4.00 exercise price		
Warrants	3.4 M @ \$5.50 exercise price		
Options	1.8 M @ \$6.73 WAEP		

# **Top Holders**









# **Analyst coverage**





Balaji Prasad







# **Key Accomplishments**

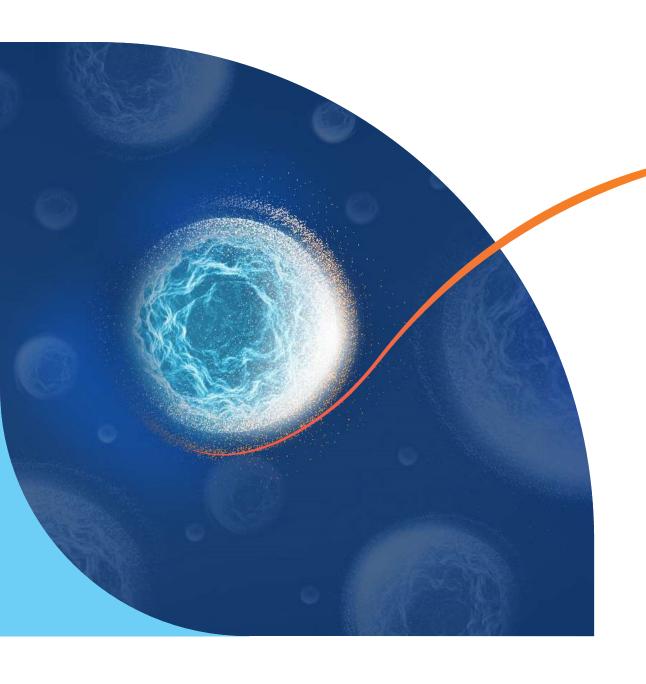
Raised over \$65 million from existing and new life science-focused investors Advanced the development of OncoPLEX including pre-IND studies

Signed a commercialization agreement for Europe - deal worth >\$110 million in milestone payments plus royalties and transfer price Initiated a second Phase 3 for prevention of SSI in abdominal colorectal resection with large incisions – top line expected by Q2 2025

Completed the largest Phase 3 trial in prevention of SSI in colorectal resection in over a decade Completed process validation for D-PLEX $_{100}$  and passed cGMP inspection – manufacturing facility ready for EU launch









# **THANK YOU**

Polypid.com

# PLEX based product typical presentation POLYPID

### Solid spheres

Micron range in diameter

## Dry format (powder) and sterile

Ready to use

### **Each particle contains**

the PLEX formulation & the Pre-Encapsulated API/APIs

### The PLEX formulation

is predesigned to achieve the needed release characteristics

## The dry powder can be

prepared for administration by either:

**Hydration into a paste**, to be applied locally into the wound/tumor bed during the surgery

**Injected (≥21G) as a paste**, or as a dry powder – Once or multiple applications



# Recognizes the Potential Value of D-PLEX<sub>100</sub> in SSI



### 3 Fast Track Designations

More frequent meetings with the FDA to discuss the development plan

Eligible for accelerated approval and priority review, if relevant criteria are met

Rolling Review

## 3 Qualified Infectious Disease Product (QIDP) Designations

All the benefits of Fast Track

Additional 5-years of market exclusivity

Improved CMS add-on payment, increase of the NTAP from 50% to 75%

# Breakthrough Therapy Designation

All the benefits of Fast Track

Intensive guidance from FDA on an efficient drug development program

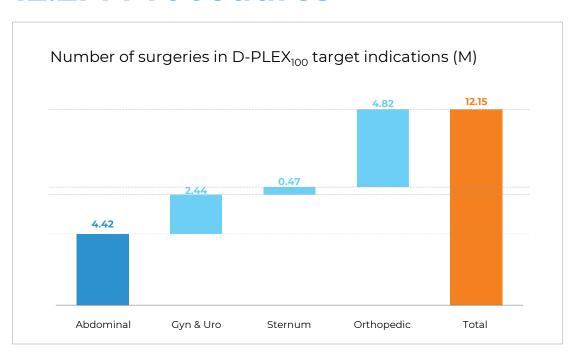
Organizational commitment from FDA involving senior managers



# **Total US Addressable Market**

US TAM for D-PLEX<sub>100</sub> is Over

# **12.2M Procedures**





# Main drivers of surgery volumes

### **Abdominal surgeries**

- Herniorrhaphies 2.1M / year
- · Cholecystectomies 616K/year
- Colorectal resection 544K / year

#### **Gynecology & Urology surgeries**

- Hysterectomies 660K /year
- Oophorectomies 1.1M / year

#### **Orthopedic surgeries**

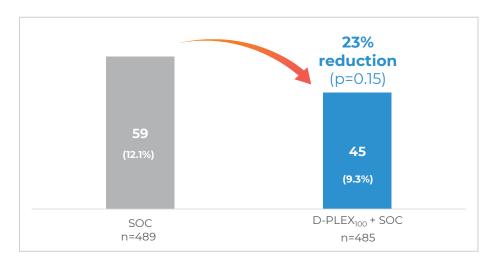
- Joint replacement 1.8M / year
- Long bone fraction 2M / year
- Spine procedures 1M /year





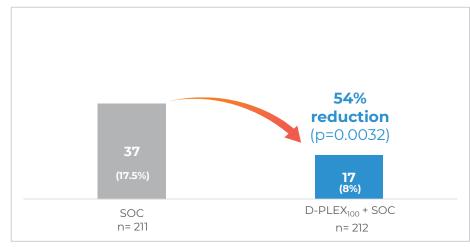
### All cohort

(primary endpoint\*, ITT)



# Large incisions complex surgeries – pre-specified subgroup analysis

(primary endpoint, incisions >20cm)



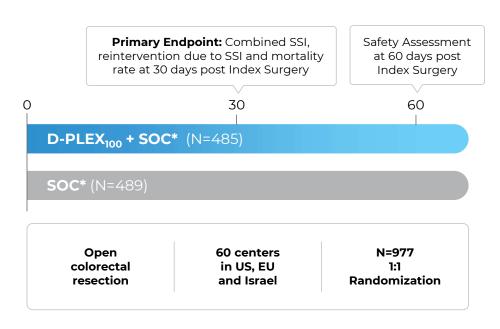


SHIELD I Study
was the Largest
Phase 3 Study of
Infection Prevention in
Colorectal Surgery in
Over a Decade



# Assess efficacy and safety of D-PLEX<sub>100</sub> for prevention of deep and superficial incisional SSI after elective abdominal colon surgery

(prospective, multicenter, randomized, controlled, two arm, double-blind study)



\*SOC - Standard of Care 28