



PledPharma

# COMPANY PRESENTATION

November 2016

# Decision to develop PledOx<sup>®</sup> to market registration

## Developing PledOx<sup>®</sup> to market registration

- Confidence in the PledOx<sup>®</sup> project
  - Unique Phase IIb data (PLIANT trial)
  - PFS data in PLIANT follow-up in May 2016
  - Positive feedback from EMA and FDA regarding commencement into Phase III studies
- Development of PledOx<sup>®</sup> to market registration expected to cost approx. SEK 750 million
  - SEK 400m to finance 2 pivotal Phase III trials until “top-line” data is available in 2020 – **adjuvant** and **metastatic** setting
  - Follow-up and market registration process expected to be financed through equity raise or strategic partnership at later stage
- Phase III trials will unlock the value potential of PledOx<sup>®</sup>
  - A significantly higher value for completed Phase III studies
  - A larger proportion of the value will be attributed to PledPharma

# PledOx<sup>®</sup>

(calmangafodipir)







**Chemotherapy can cure cancer...**

...but is associated with dose limiting and debilitating toxicity

# Neuropathy is one of the most feared side effects of chemotherapy

## Most common dose limiting side-effects of oxaliplatin (used in FOLFOX treatment)

Side-effect	Incidence	Unmet need	Treatment options
<b>Neuropathy</b> (Nerve damage)	40-60% <sup>1</sup>	 High	<ul style="list-style-type: none"> <li>Often have to reduce/discontinue treatment</li> <li>Can have long-term effects</li> <li><b><u>No treatment currently available</u></b></li> </ul>
<b>Thrombocytopenia</b> (Low platelet levels)	76% <sup>2</sup>	 Medium/High	<ul style="list-style-type: none"> <li><b><u>Limited treatment options</u></b>, dose-limiting</li> </ul>
<b>Febrile Neutropenia</b> (Fever with low white blood cells)	4% <sup>2</sup>	 Medium	<ul style="list-style-type: none"> <li>Treatable with antibiotics, G-CSFs</li> <li>Severity and high cost of complications keep it top-of-mind with payers and prescribers</li> </ul>
<b>Neutropenia</b> (Low white blood cells)	53% <sup>2</sup>	 Medium	<ul style="list-style-type: none"> <li>G-CSFs can manage, but pre-treatment is restricted to certain patient populations</li> <li>Dose-limiting</li> </ul>

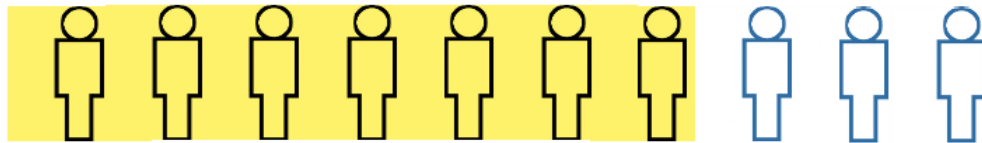
Source:

1. Loprinzi et al., 2013 JCI; Ventzel et al. 2015 Pain
2. Eloxatin Label.

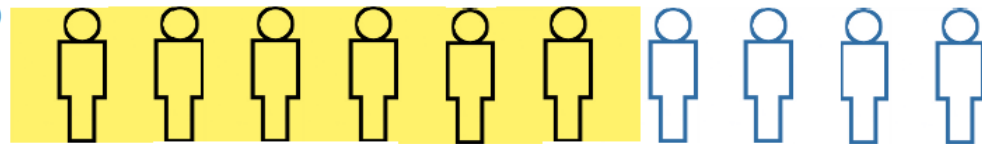
# Incidence of CIPN

## How common is it?

1 month after chemo



3 months after chemo



6 months or longer



(Seretny *et al.* 2014)

# Some important quotes on the impact of CIPN

Symptoms have substantial impact on an individual's quality of life (Tanay *et al.* 2016)

**"It is now 14 years later..."**

"My feet don't feel like they're 100% part of me. They feel slightly disconnected."

**"I've fallen down the stairs a few times."**

"I started to drop things. My hands are numb."

"Tingling in the hands and feet. They burn. It's like an anthill and it runs in the feet."

**"People don't understand about it all."**

"It has affected my mobility more than anything else. I can't go very far. Sometimes, I can't go out at all."

**"It was like my nails were coming away from my nail bed"**

**"I used to work with computer and I can't feel the keys."**

"I felt like I had pebbles or marbles in my shoes."

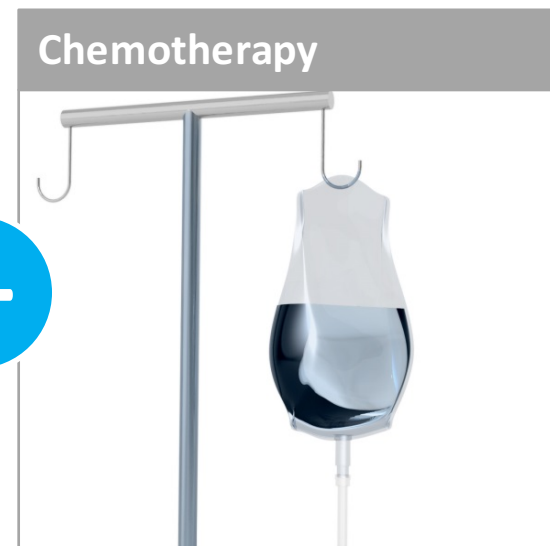
**"I spilled a cup of hot tea on someone as I lost grip."**

"I had to really kind of hang on just so I didn't fall over."

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# PledOx<sup>®</sup> aims to become a standard pre-treatment to chemotherapy to help prevent neuropathy

PledOx<sup>®</sup> infusion 5 minutes  
before chemotherapy



**FOLFOX – 12 cycles  
over 6 months  
in the adjuvant  
setting**

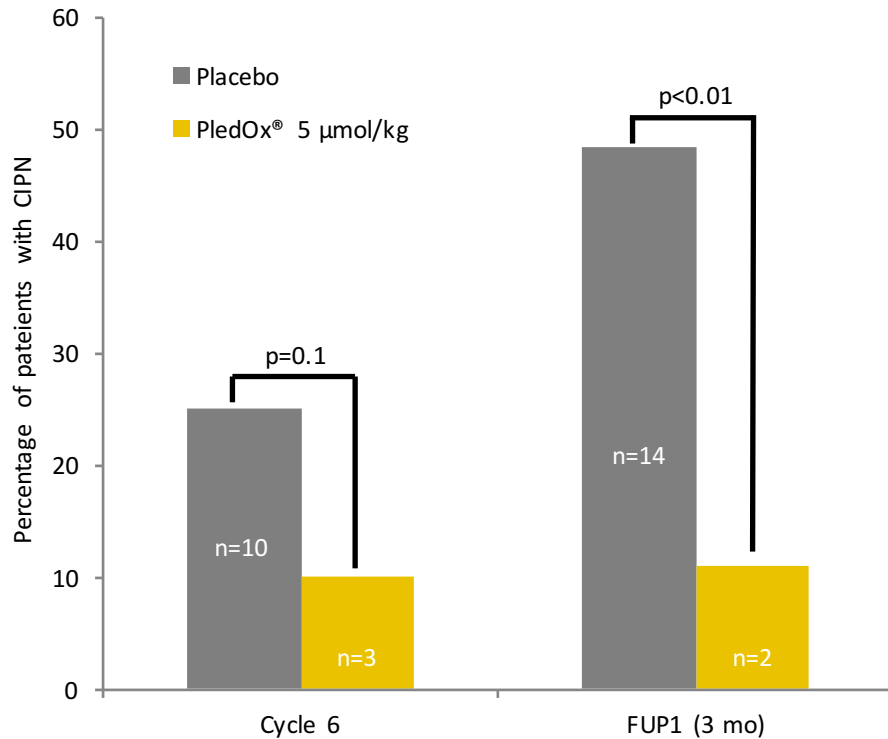
**CAPOX – 8 cycles  
over 6 months  
in the adjuvant  
setting**

# PledOx<sup>®</sup> – Significant reduction of number of neuropathies while having no apparent negative interaction on chemo treatment



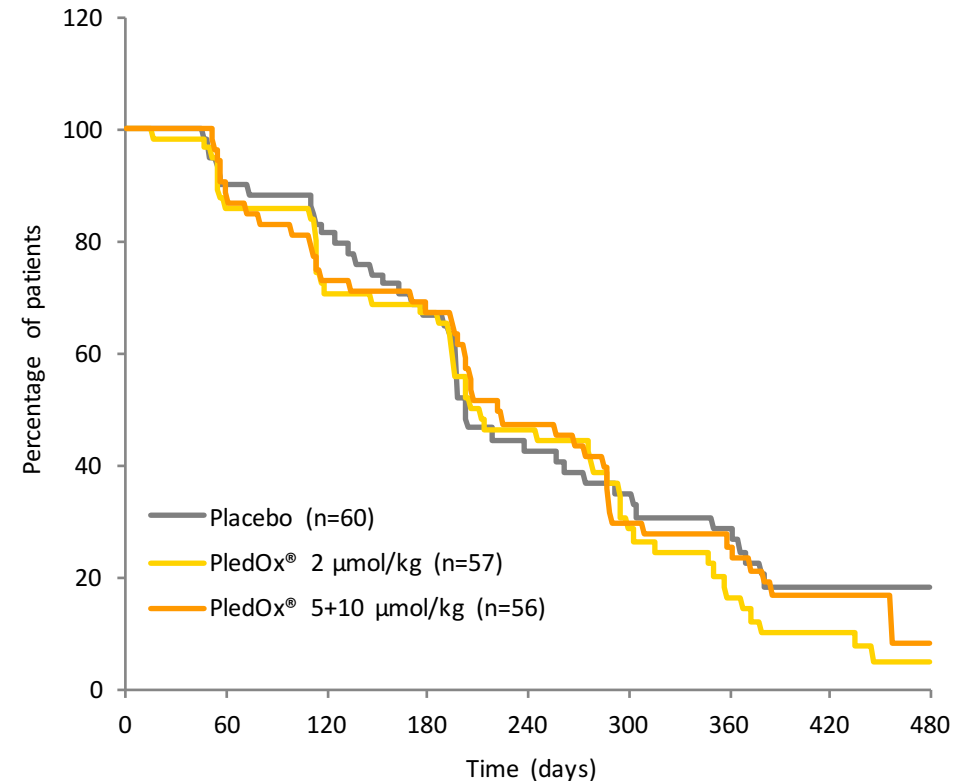
## Responder analysis – PLIANT trial

Responder analysis on PLIANT patients scoring any grade 3 or worse numbness or tingling at cycle 6 and 3 months after EOT



## Progression Free Survival – PLIANT trial

Time to progression



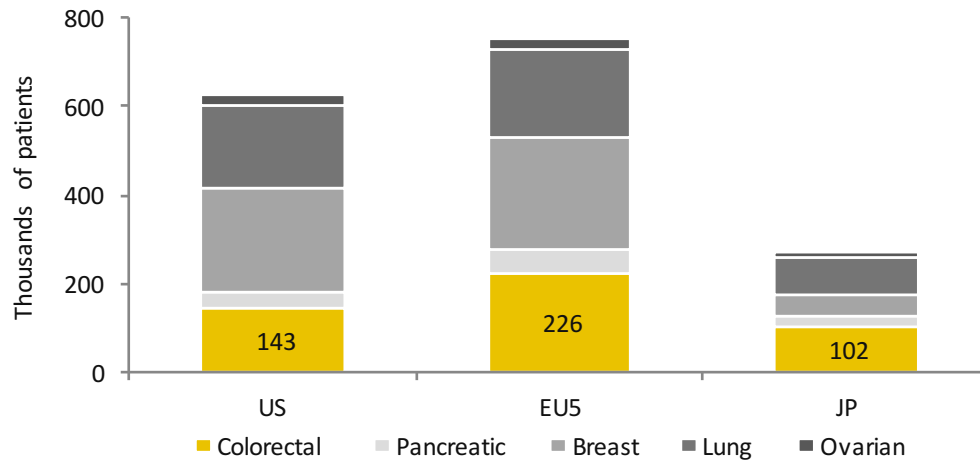
**Significant reduction of number of patients with both acute and chronic CIPN**



# PledOx<sup>®</sup> is currently focused on colorectal cancer and oxaliplatin

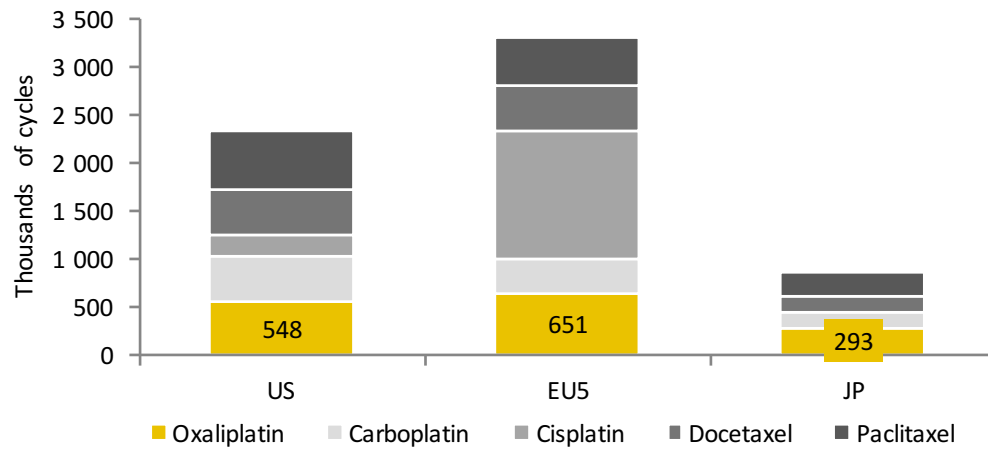


Annual applicable cancer patients<sup>1</sup>



PledOx<sup>®</sup> is initially targeting 470k annual colorectal cancer patients

Annual number of cycles used<sup>2</sup>



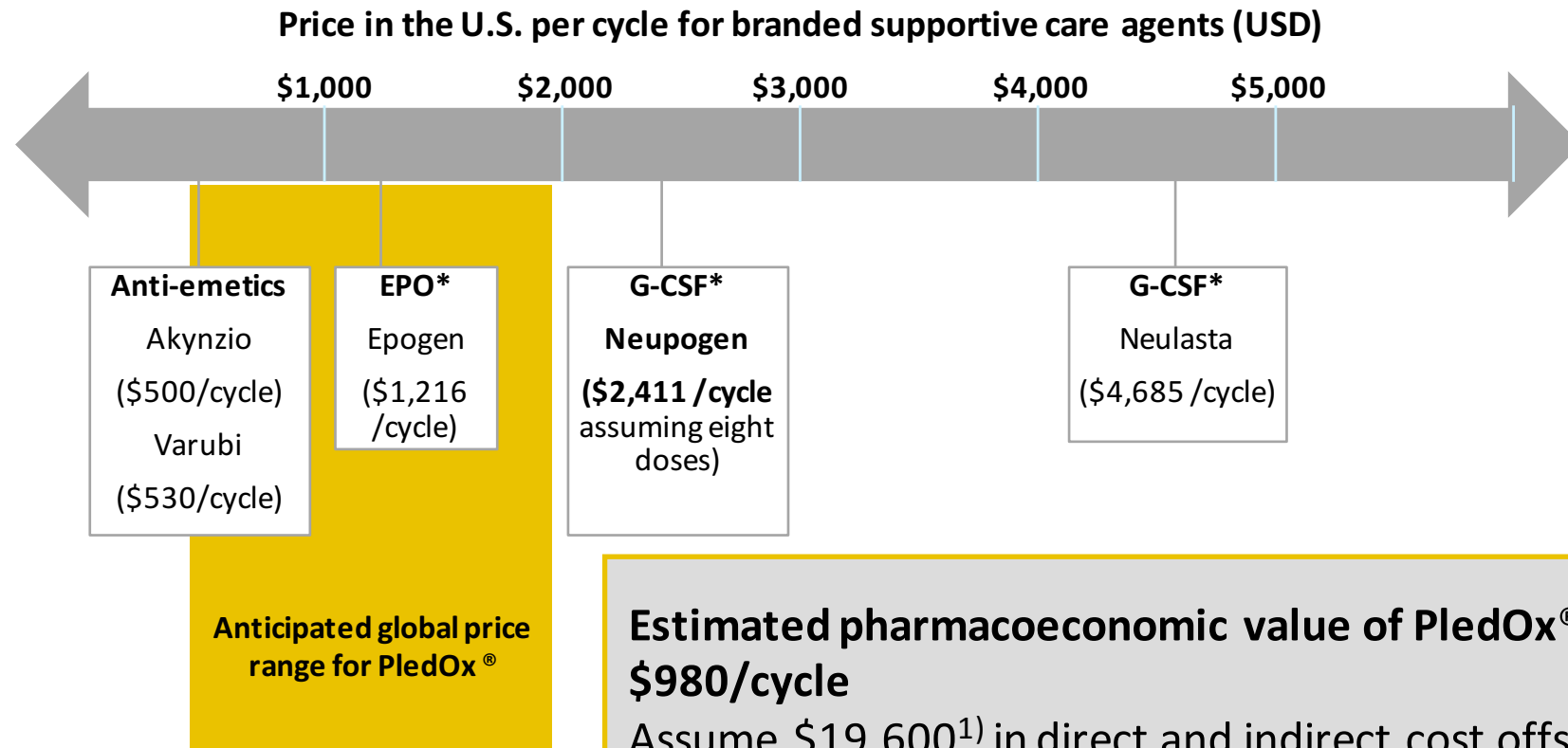
1.5m treatment cycles for colorectal cancer annually

Source:

1. SEER, EUCAN, GLOBOCAN.

2. IMS Data on unit sales of chemotherapy agents with regimen assumptions applied; no data for cisplatin in JP.

# Anticipated global peak sales USD 1.2 – 2.4 billion in CRC



## Estimated pharmacoeconomic value of PledOx<sup>®</sup> \$980/cycle

Assume \$19,600<sup>1)</sup> in direct and indirect cost offsets x 40% reduction / 8 cycles, does not take into account cost for chronic neuropathy, fall accidents<sup>2)</sup> and that 2/3 CIPN patients reportedly have to stop working<sup>3)</sup>

Source:

1. Pike et al. 2012
2. Kolb et al. 2016
3. Calhoun et al. 2001

# PledOx<sup>®</sup> is currently focused on colorectal cancer and oxaliplatin with further potential



PledOx<sup>®</sup> has applicability in the treatment of the most common types of cancer:

- Colorectal cancer
- Lung cancer
- Ovary cancer
- Breast cancer
- Pancreatic cancer

PledOx<sup>®</sup> can potentially be used in a broad range of platinum- and taxane-based chemotherapy regimens:

- Oxaliplatin
- Cisplatin
- Carboplatin
- Paclitaxel
- Docetaxel
- Immunotherapy

Initial focus

Future expansion

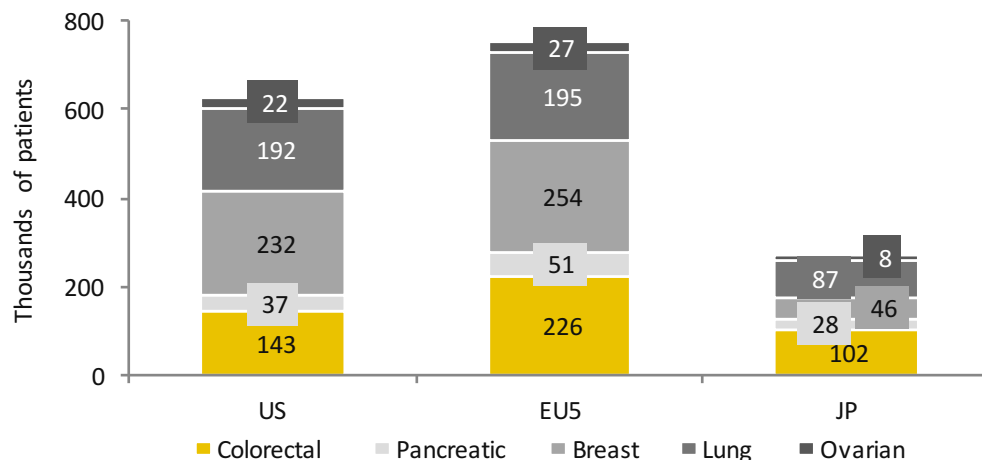
Further investigation

Opportunity to broaden market to taxanes, breast cancer and beyond

# PledOx<sup>®</sup> could be applicable to 1.7m cancer patients in US, EU5, JP

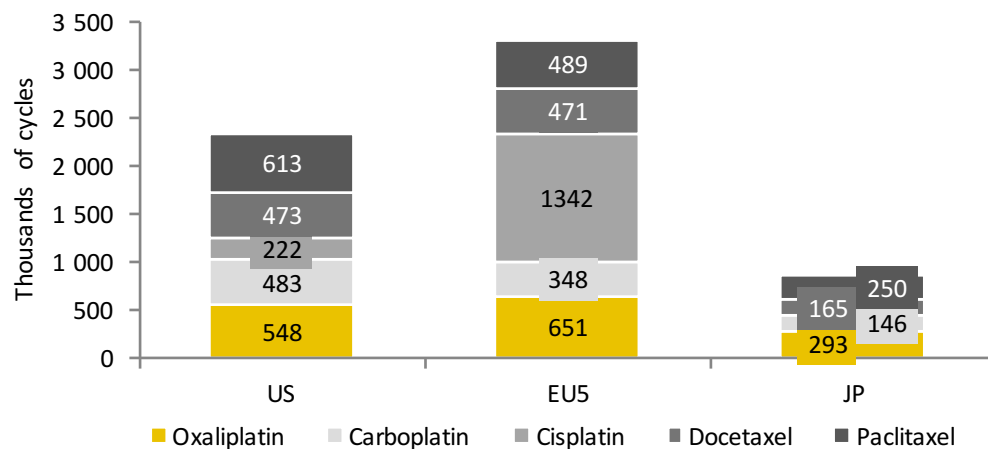


Annual applicable cancer patients<sup>1</sup>



PledOx<sup>®</sup> is expected to have efficacy in all these forms of cancer

Annual number of cycles used<sup>2</sup>



Source:

1. SEER, EUCAN, GLOBOCAN.

2. IMS Data on unit sales of chemotherapy agents with regimen assumptions applied; no data for cisplatin in JP.

# PledOx<sup>®</sup> – Summary of results from PLIANT and implied treatment benefits



## Summary of results from PLIANT

- Statistically significant reduction of chronic patient-reported neuropathies after end of treatment
- No apparent negative impact on anti-cancer effect of chemotherapy
- Reduced frequency, onset and duration of grade 2 or worse investigator-reported neuropathies
- A very benign safety profile
- No negative influence on other toxicities

## PledOx<sup>®</sup> treatment benefits

- **For oncologist:** Fewer dose-reductions and/or treatment discontinuations due to CIPN
  - Oxaliplatin can be used as an important tool in subsequent lines of therapy
  - No negative interaction with anti-cancer treatment
  - No extra “chair time”; no change to treatment paradigm
- **For cancer patients:** Fewer patients with persistent CIPN
  - Significantly improved quality of life
  - Lower morbidity with fewer falls, cuts, etc.
  - Convenient treatment paradigm

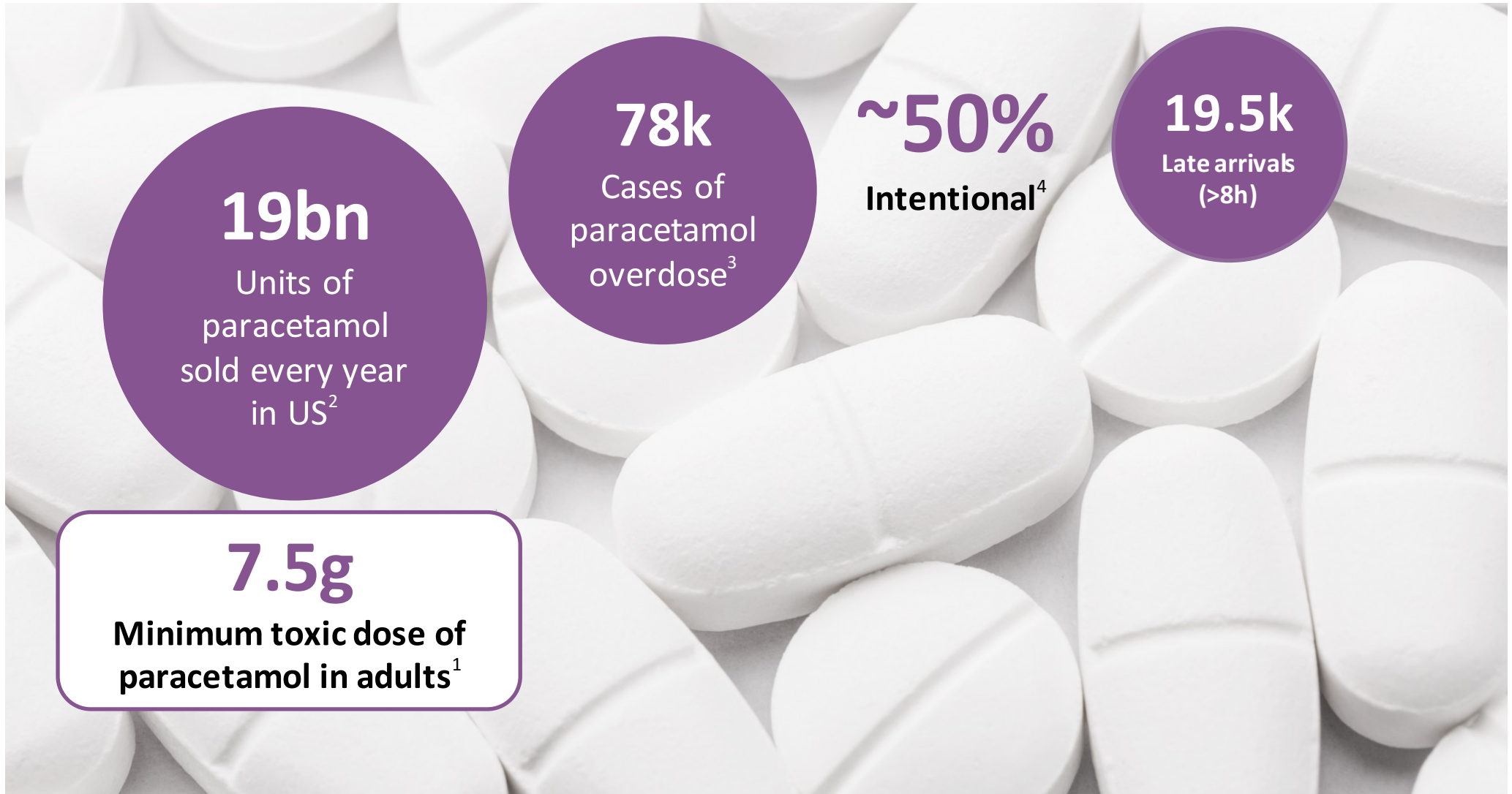
# Aladote<sup>®</sup>

(differentiated product from PledOx<sup>®</sup>)



**Paracetamol is one of the most largely used drugs...**  
**...but accidental or intentional overdose is a growing problem**

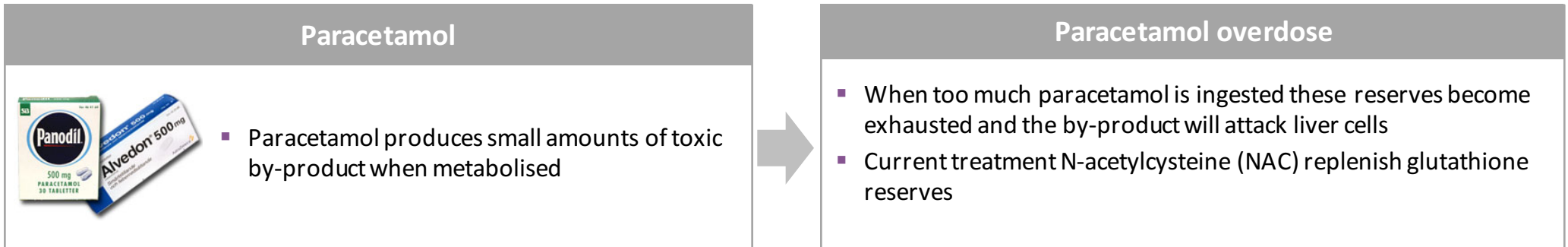
# Aladote® – Paracetamol/acetaminophen overdose is one of the most common forms of poisoning in the US



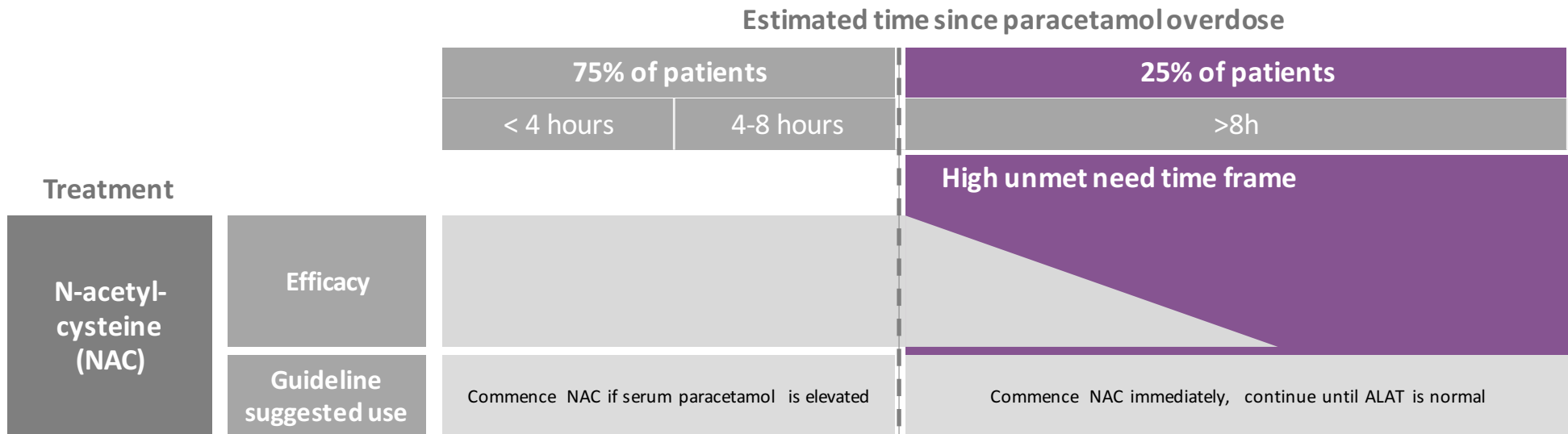
Source:

1. Schilling, A. M. Y., et al. "Acetaminophen: old drug, new warnings." *Cleveland Clinic journal of medicine* 77.1(2010): 19-27.
2. IMS, MIDAS DATA, 2013
3. Manthripragada AD, Zhou EH, Budnitz DS, Lovegrove MC, Willy ME. Characterization of acetaminophen overdose-related emergency department visits and hospitalizations in the United States. *Pharmacoepidemiol Drug Saf.* 2011;20(8):819-26.
4. FDA: "Acetaminophen Overdose and Liver Injury - Background and Options for Reducing Injury", (2009).

# Aladote<sup>®</sup> – targets patient population currently lacking effective antidote treatment



Although NAC efficacy declines after ~8 hours, it is still recommended for use even after 24 hours because no other treatments are available

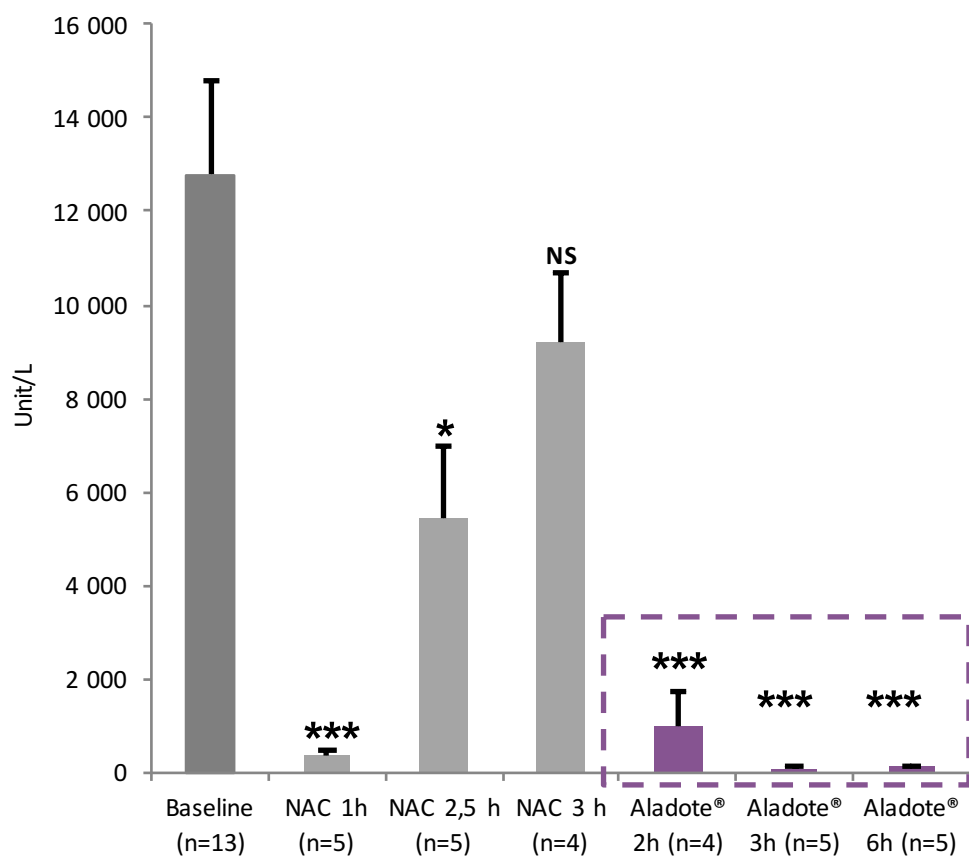


Source: Company information



# Aladote<sup>®</sup> has demonstrated a protective effect beyond the NAC therapeutic window in animal models

## Level of liver damage (ALAT test in mice)



- Male B6C3F1 mice were fasted 8-10h, injected with 300 mg/kg APAP i.p.
- Test compounds were injected i.v. at time point indicated in graph (NAC 300 mg/kg, Aladote<sup>®</sup> 10 mg/kg).
- Serum concentration of ALAT determined at 12h

Source: Company information  
Note: \*\*\* =  $p \leq 0.01$ , \*\* =  $p \leq 0.05$ , \* =  $p \leq 0.1$ , NS = not significant

## Summary of results

- Aladote<sup>®</sup> appears to be more effective in lowering the level of ALAT especially in later time frames compared to NAC
- As mice have much faster metabolism than humans, the comparable time frames are many times higher demonstrating that Aladote<sup>®</sup> could be effective in treating late arrivals (>8h)



# News flow up to top-line data 2020



- Aladote®:
  - PoP study initiated
  - PoP results
  - PoC study initiated
  - PoC results
- Corporate:
  - Financing closed
  - Main market listing
  - IP updates
  - Strengthening of organization
  - Continued BD activities
- PledOx®:
  - PLIANT publication
  - Selection of CRO
  - FDA type C meeting – phase III design
  - EMA - phase III design
- Start of phase III:
  - Approval of phase III studies
  - Centers recruited and initiated
  - First patient in
  - DSMB status reports
  - Last patient in

## Rights issue details

- Subscription price: 20 SEK/share
- Record date: Nov 11, 2016
- Subscription period: Nov 15 – 29, 2016
  - Payment according to instructions from your bank or asset manager
- Seven (7) subscription rights plus 100 SEK =  
Five (5) new shares
- Trading period for subscription rights: Nov 15 – 29, 2016
- "BTA" trading period: Nov 15 – Dec 1, 2016

# Summary

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# Income statement Q3 report

Tkr	2016 jan-sept	2015 jan-sept	2015 jan-dec	2014 jan-dec
<b>Intäkter</b>				
Övriga rörelseintäkter	985	321	378	233
	<b>985</b>	<b>321</b>	<b>378</b>	<b>233</b>
<b>Rörelsens kostnader</b>				
Projektkostnader	-13 142	-22 100	-26 093	-29 459
Övriga externa kostnader	-9 793	-8 235	-11 274	-13 086
Personalkostnader	-4 631	-5 089	-6 909	-6 271
Av- och nedskrivningar av anläggningstillgångar	-0	-2	-2	-2
Övriga rörelsekostnader	-287	-91	-128	-146
<b>Rörelseresultat</b>	<b>-26 868</b>	<b>-35 196</b>	<b>-44 028</b>	<b>-48 731</b>
<b>Resultat från finansiella poster</b>				
Ränteintäkter	106	165	203	312
Räntekostnader och liknande resultatposter	-	-10	-10	-1
<b>Resultat efter finansiella poster</b>	<b>-26 761</b>	<b>-35 041</b>	<b>-43 836</b>	<b>-48 420</b>
<b>Periodens resultat</b>	<b>-26 761</b>	<b>-35 041</b>	<b>-43 836</b>	<b>-48 420</b>

## Balance sheet - Assets Q3 report

Tkr	2016-09-30	2015-09-30	2015-12-31	2014-12-31
<b>TILLGÅNGAR</b>				
<b>Anläggningstillgångar</b>				
<i>Materiella anläggningstillgångar</i>				
Inventarier, verktyg och installationer	0	1	0	2
<b>Summa anläggningstillgångar</b>	<b>0</b>	<b>1</b>	<b>0</b>	<b>2</b>
<b>Omsättningstillgångar</b>				
Kortfristiga fordringar				
Övriga fordringar	405	783	788	2 732
Förutbetalda kostnader och upplupna intäkter	2 044	2 380	1 213	430
	<b>2 449</b>	<b>3 164</b>	<b>2 001</b>	<b>3 162</b>
Kassa och bank	23 590	59 313	50 360	100 304
<b>Summa omsättningstillgångar</b>	<b>26 040</b>	<b>62 477</b>	<b>52 361</b>	<b>103 466</b>
<b>Summa tillgångar</b>	<b>26 040</b>	<b>62 478</b>	<b>52 361</b>	<b>103 468</b>

# Balance sheet- Equity and liabilities Q3 report

Tkr	2016-09-30	2015-09-30	2015-12-31	2014-12-31
<b>EGET KAPITAL OCH SKULDER</b>				
<b>Eget kapital</b>				
Aktiekapital	1 494	1 494	1 494	1 492
Övrigt tillskjutet kapital	46 538	90 374	90 374	137 586
Balanserat resultat inklusive årets resultat	-26 761	-35 041	-43 836	-48 420
<b>Summa eget kapital</b>	<b>21 271</b>	<b>56 827</b>	<b>48 032</b>	<b>90 658</b>
<b>Kortfristiga skulder</b>				
Leverantörsskulder	2 271	2 481	1 766	9 967
Skatteskulder	0	0	0	0
Övriga skulder	202	164	177	292
Upplupna kostnader och förutbetalda intäkter	2 295	3 006	2 386	2 551
<b>Summa kortfristiga skulder</b>	<b>4 769</b>	<b>5 651</b>	<b>4 329</b>	<b>12 810</b>
<b>Summa eget kapital och skulder</b>	<b>26 040</b>	<b>62 478</b>	<b>52 361</b>	<b>103 468</b>