

ReNeuron

Changing Patients' Lives

Shareholder Presentation

AGM Trading update

September 2018



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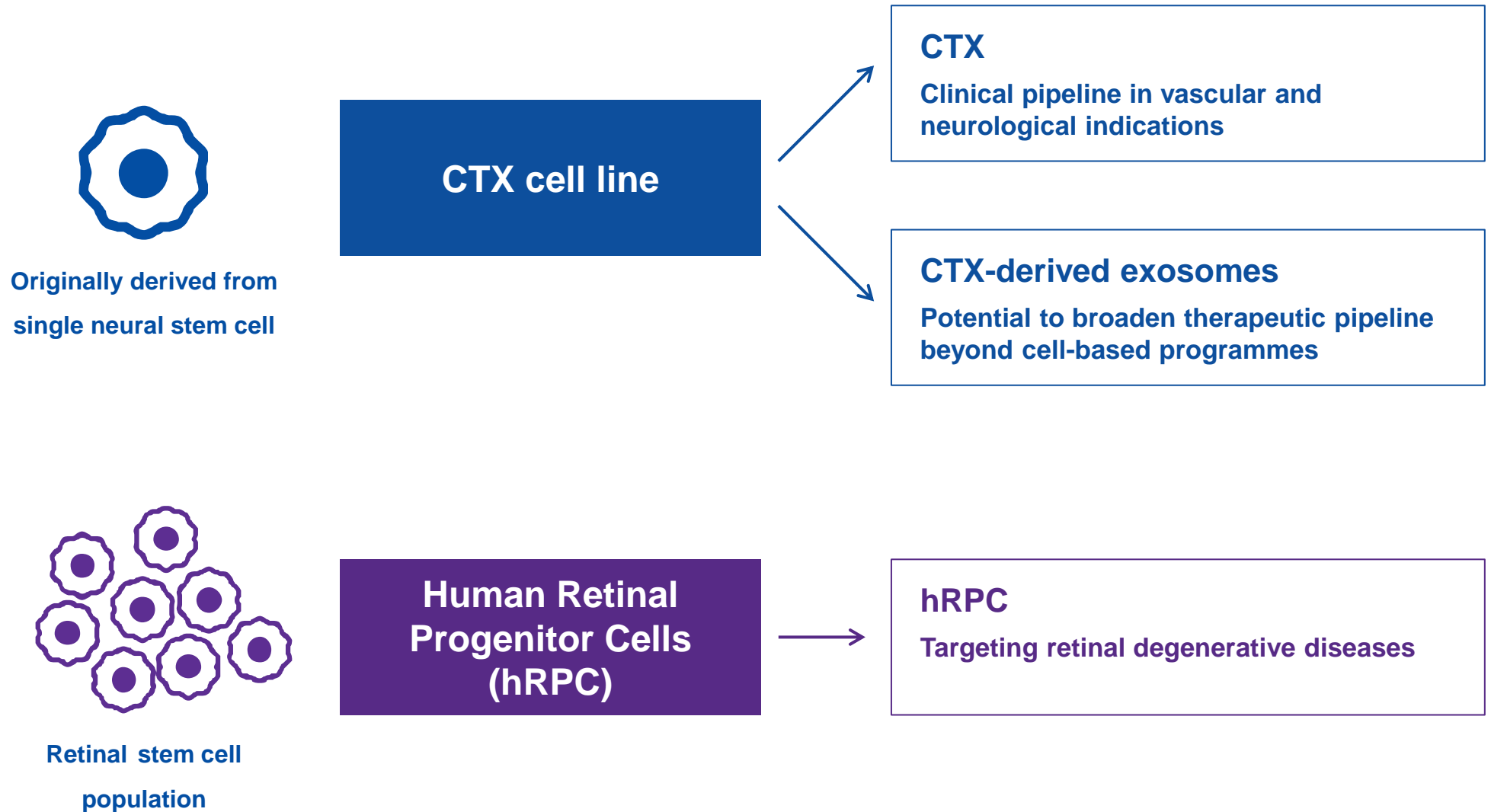
Clinical Highlights

- **CTX stem cell therapy candidate for stroke disability:**
 - Long-term data from Phase II clinical trial presented, showing sustained improvements in motor function and reduced levels of disability and dependence
 - IND application approved by FDA to commence a Phase IIb, placebo-controlled clinical trial in the US
 - First clinical site initiated - top-line data expected in early 2020
- **hRPC stem cell therapy candidate for retinal diseases:**
 - Four cohorts of patients treated in ongoing US Phase I/II clinical trial in retinitis pigmentosa (RP)
 - hRPC drug product formulation successfully optimised for sub-retinal injection
 - Phase I/II study to be expanded to target patients with less impaired vision
 - Top line Phase I/II data expected in mid-2019
 - Phase II study planned in cone-rod dystrophy patients, to run in parallel with planned Phase IIb study in RP
- **Exosome nanomedicine platform:**
 - ExoPr0 exosome candidate demonstrates therapeutic potential
 - Positive pre-clinical data demonstrates that ExoPr0 exosome therapy candidate significantly reduces tumour volume in a variety of *in vivo* models of cancer
 - Initial clinical trial application planned for 2019 in oncology

Operational Highlights

- US office established in Boston area, reflecting the Company's increasing clinical activity in the US
- Increased business development activity in the period due to third party interest in Company's core therapeutic programmes
 - Active discussions ongoing with a number of commercial third parties
- Increased collaborative work in the period to exploit technology platforms beyond core therapeutic programmes
- Exclusivity agreement signed with US-based specialty pharmaceutical company regarding potential out-license of hRPC technology platform and therapeutic programmes:
 - Three month exclusivity period
 - \$2.5 million already received
 - Further \$2.5 million upon completion of certain due diligence activities
 - Definitive out-license agreement targeted for later this year

Unique Platform Technologies



Pipeline

Product/ Indication	Discovery	Pre-clinical	Phase I	Phase IIa	Phase IIb/pivotal	Market approval
CTX Cell line						
CTX Stroke Disability						
CTX Critical Limb Ischaemia						
Exosomes (CTX-derived) Cancer						
Human Retinal Progenitor Cells						
hRPC Retinitis Pigmentosa						
hRPC Cone-Rod Dystrophy						

Strong Investor Base and Balance Sheet

- Quoted on AIM
- Backed by major generalist and specialist life science institutional investors:

35.5%

Woodford Investment
Management

11.9%

Invesco

9.5%

Wales Life
Science Fund

- Cash on balance sheet at last reporting date (March 31, 2018) – £37.4 million (US\$48.6 million) – current cash resources for at least the next 12 months

Market Potential

based on analyst estimates*

Indication	Assumptions	Peak Annual Sales
CTX for stroke	<ul style="list-style-type: none"> 1.76 million strokes/year (total US/EU/Japan) 85% survival; 85% ischaemic; 75% long-term disability; 25% mRS score 3-4; 66% residual upper limb mobility Peak penetration 20% US/EU/Japan Treatment cost \$45,000 EU to \$75,000 US/Japan 	\$1.1bn - \$2.2bn
hRPC for RP	<ul style="list-style-type: none"> Prevalence 1:4000, ~244,000 cases (total US/EU/Japan) 15% advancing to severe vision loss Peak penetration 50% US/ EU Per-eye treatment cost \$50,000 EU to \$100,000 US/Japan 	\$0.5bn - \$1.6bn

- Applicability of hRPC in other hard-to-treat ophthalmic diseases could provide upside potential
- Longer-term upside from exosome platform

Chronic Stroke: A Large, Unmet Medical Need and Tremendous Economic Burden



- In the US, around 800,000 people experience a new or recurrent stroke each year ¹
- Stroke costs are \$34 billion annually in US, including healthcare costs, medications, lost productivity. ²
- Only one pharmaceutical treatment option available within 4.5 hours of symptom onset. ³
- No treatment options available for stroke patients months to years later
 - Rehabilitation provides limited benefit – most in 1st month, less thereafter, very little beyond 6 months⁴

CTX for chronic disability from stroke

CTX is a human neural stem cell product

Commercial scale manufacturing produces “off-the-shelf” drug product available on demand

CTX is safe and well-tolerated

One-time, stereotactic, intracerebral injection up to 20 million cells

Clinically meaningful improvements in key measures of
functional disability post-stroke with CTX

Based on well-established and widely accepted modified Rankin Scale disability assessment

Market potential for CTX in stroke disability is estimated to
have peak annual sales in **\$1-2 billion** range

US/EU/Japan

CTX Clinical Development in Stroke Disability

Completed PISCES trials

PISCES I:

- Phase I, first-in-human, safety study
- 11 disabled, stable stroke patients
 - 6 months to 5 years post-stroke
- CTX dose escalation (2, 5, 10, 20 million cells) administered by stereotactic, intracerebral injection
- No cell-related or immunological adverse events



CTX treatment is safe and well-tolerated

PISCES II:

- Phase II, single arm, open label
 - 20 million CTX cell dose
- 23 disabled, stable stroke patients
 - 2 to 12 months post-stroke
- Clinically meaningful improvements in disability scales were measured out to 12 months post-implantation
- No cell-related safety issues identified



Very promising results for chronic stroke disability; warrants a larger, randomised, placebo-controlled Phase IIb study

PISCES II efficacy measure

PISCES II

Modified Rankin Scale (mRS) – measure of functional disability

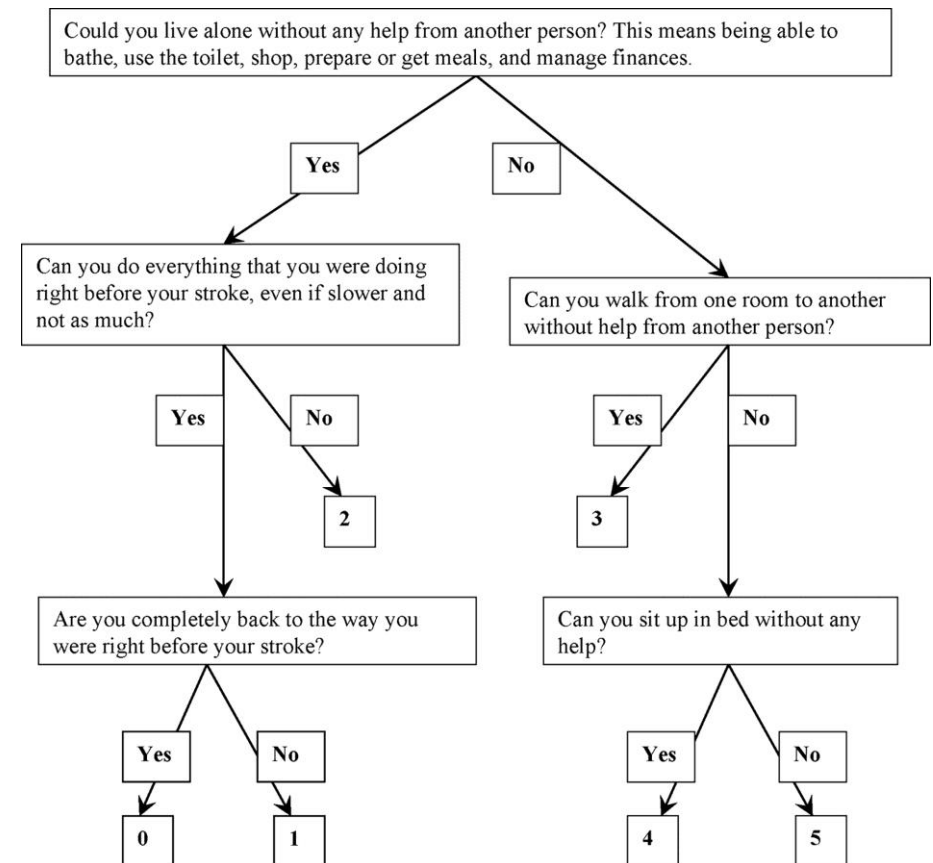
Total Subjects			Patients with NIHSS upper limb score < 4 at baseline		Patients with NIHSS upper limb score = 4 at baseline	
Month	N	n* (%)	N	n* (%)	N	n* (%)
Baseline	23	-	14	-	9	-
3	23	7 (30.4%)	14	6 (42.9%)	9	1 (11.1%)
6	22	6 (27.3%)	13	5 (38.5%)	9	1 (11.1%)
12	20	7 (35.0%)	12	6 (50.0%)	8	1 (12.5%)

*number of subjects with ≥ 1 point improvement in mRS (% of N observed at day of visit)

Improvements in mRS greatest in subjects with residual movement of the affected arm (NIHSS UL <4)

Modified Rankin Score

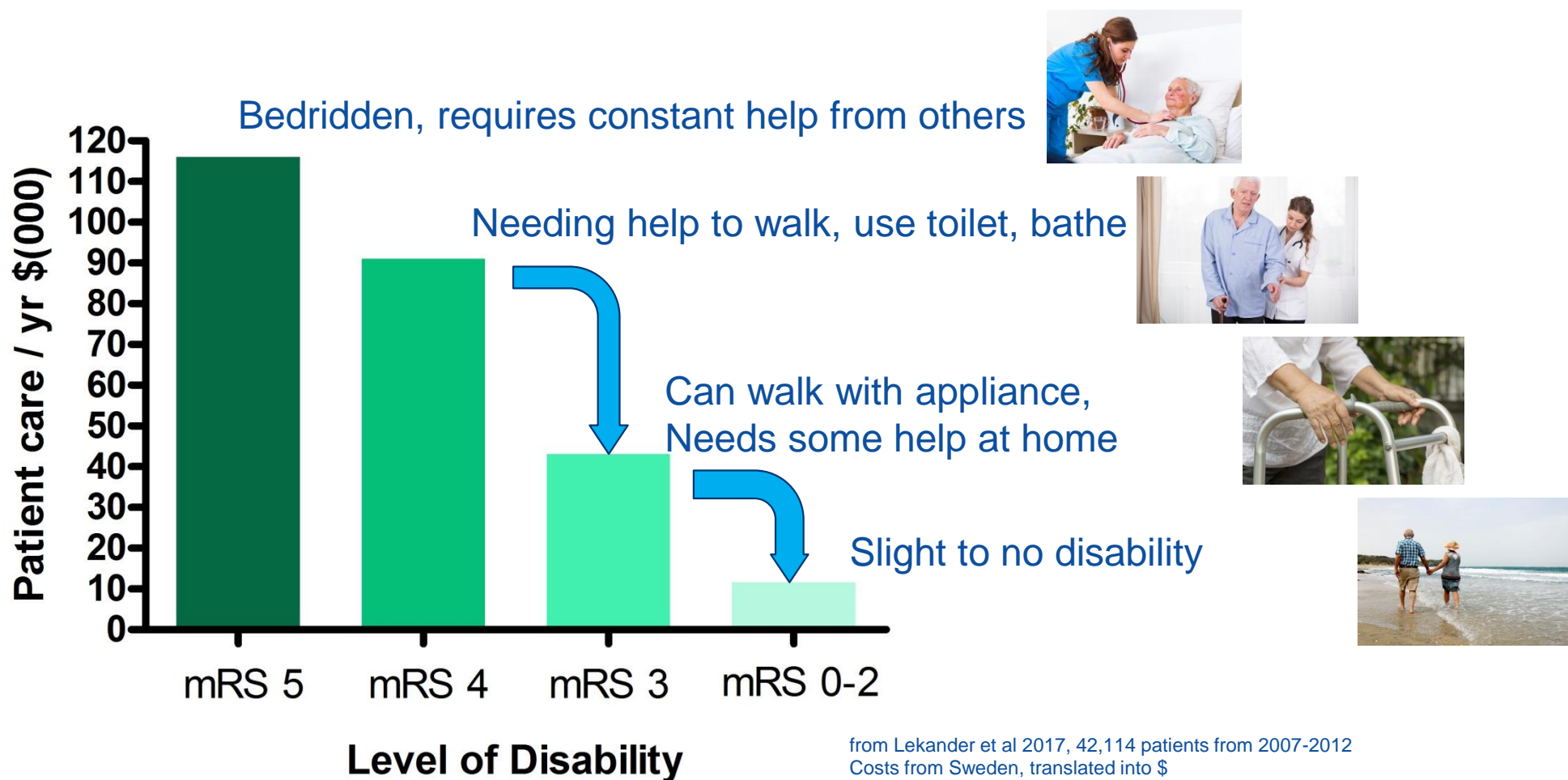
Category
5 Bedridden, completely dependent on others
4 Needing help to walk, use toilet, bathe
3 Can walk, but need still need help at home
2 Mostly recovered, but still has limitations
1 Slower than before, but no limitations
0 Back to pre-stroke life



Algorithm from Bruno et al, 2010

Improvement by one category is a significant change in a patient's life

Costs of disability – mRS scale



Reductions in disability result in substantial reductions in patient care costs

PISCES III trial



PHASE IIb
INVESTIGATION
OF STEM CELLS
IN STROKE

Primary Objective:

Assess efficacy of intracerebral CTX by change in degree of dependency and disability as measured by mRS

Primary Endpoint:

- ≥ 1 pt improvement from baseline in mRS at 6 months post-treatment

Secondary Endpoints (1, 3, 6, 9, 12 months post-tx):

- Barthel Index (ADL independence)
- Timed Up and Go test (lower limb and trunk function)
- Chedoke Arm/Hand Activity Inventory (upper limb function)
- NIHSS
- Fugl-Meyer Assessment
- EQ-5D-5L (QoL)

Subjects (n=110):

- Age 35-75 inclusive
- Ischemic stroke that includes supratentorial region (CT/MRI confirmed)
- 6-12 months post-stroke
- mRS 3 and 4
- Some residual arm movement

First site initiated - patient enrolment commencing shortly

Retinal Platform

- The intrinsic regenerative capacity of cells in retina is limited^{1,2}.
- Any preservation of retinal structure/function balance can greatly impact vision loss associated with retinal disease
- Our programme is based on subretinal injection of hRPCs (human retinal progenitor cells)
- Collaborations:
 - Schepens Eye Research Institute (Harvard Medical School)
 - Massachusetts Eye and Ear Infirmary (MEEI)
 - University College London – Institute of Ophthalmology, UK
- Initially targeting inherited retinal degenerative diseases
 - Characterised by progressive loss of photoreceptors



Broad application across a range of retinal diseases

¹Ader et al (2014) Regenerative Biology of the Eye, A Pebay (Ed), doi: 10.1007/978-1-4939-0787-8_8; ²So and Yip (1998) Vis Res 38, 1525-1535.

Retinitis Pigmentosa (RP)

- RP is an inherited, degenerative eye disease^{1,2}
 - Onset varies from early childhood to 20s/30s
 - Early stage main symptom is night blindness
 - Progressive loss of peripheral vision (ie tunnel)
 - Incidence of RP is 1:4000 in US and worldwide
 - Estimated treatment population of 275,000 in the US and EU
- Orphan Drug Designation in EU and US
- FDA Fast Track Designation in US
- Phase I/IIa study ongoing in the US
 - Phase I dosing complete
 - Phase IIa commenced
 - Top line Phase IIa readout mid-2019



NORMAL VIEW



VIEW WITH
RETINITIS PIGMENTOSA

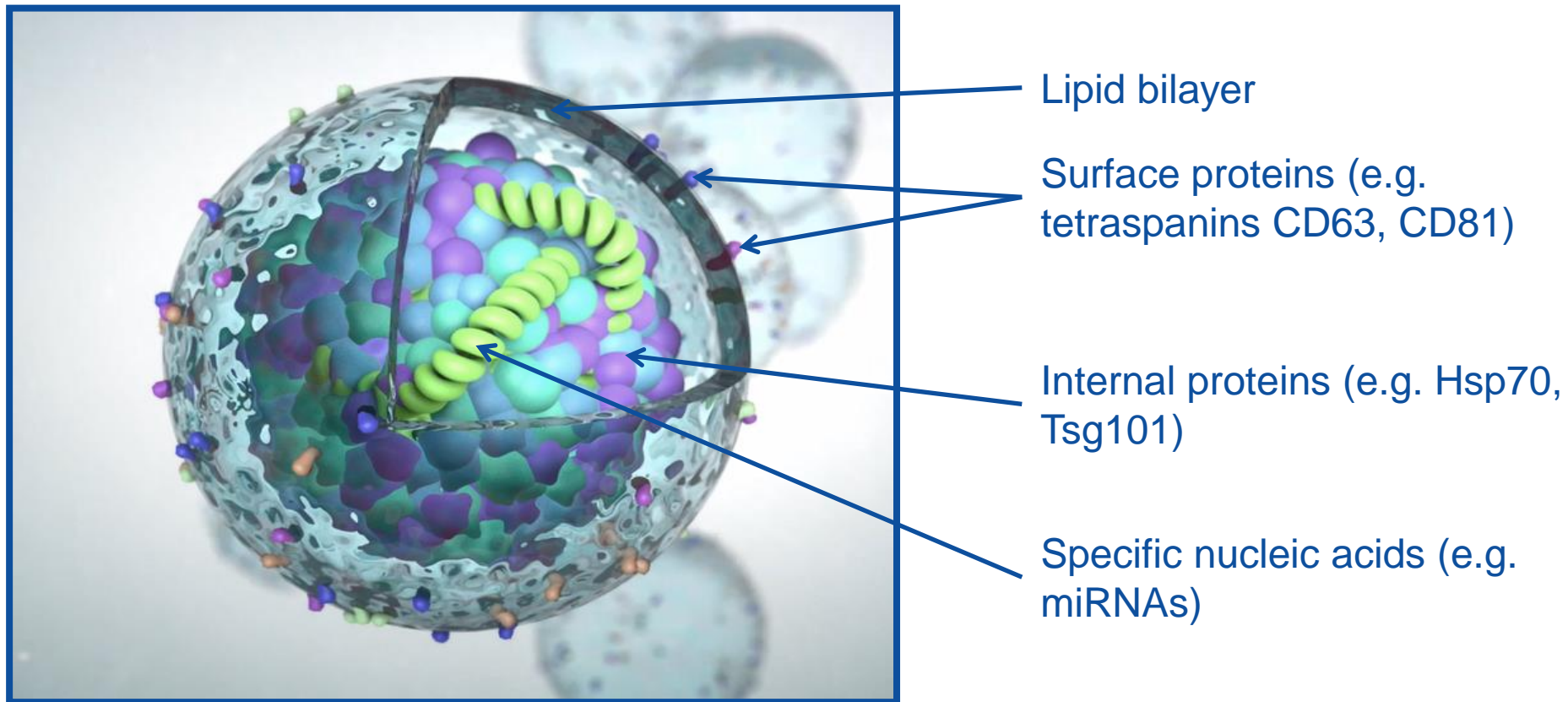
www.eyehelthweb.com/retinitis-pigmentosa

There is no treatment for the vast majority of patients with RP

¹Hamel (2006) Orphanet J Rare Disease 1, 40;
²https://nei.nih.gov/health/pigmentosa/pigmentosa_facts

What are Exosomes?

- Nano-scale vesicles (30-100nm) released by most cell types as a means of intercellular communication
- Considered to be a naturally-occurring liposomal delivery system
- Contain and transport bio-active lipids, proteins and nucleic acids

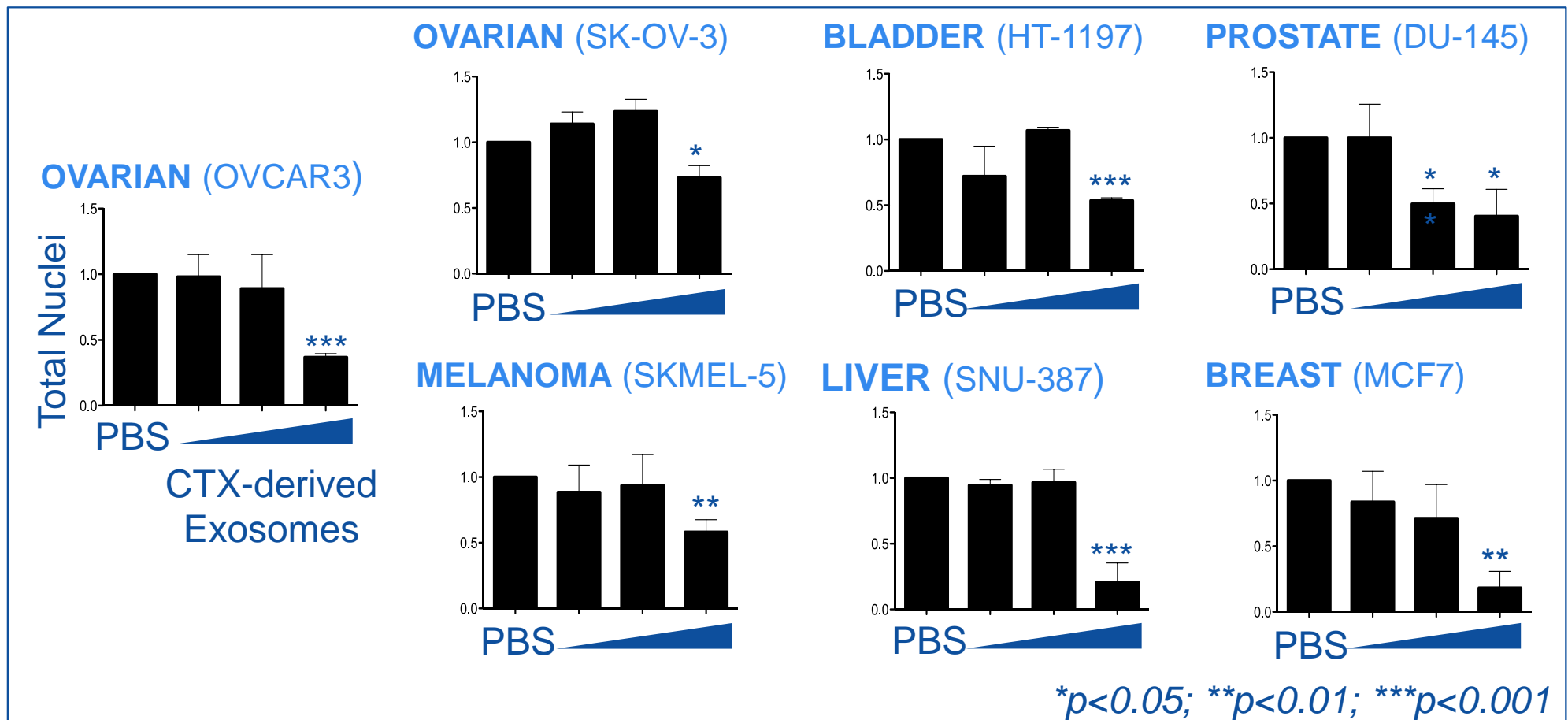


A Global Leader in Stem Cell-Derived Exosome Manufacture

- Exosome platform established at ReNeuron in 2011
- Significant IP portfolio established
- Qualified, scalable GMP process
- Proprietary clinical-grade producer cell line (CTX), giving high yields
- Stable and consistent product
- Established analytics
- Broad anti-cancer properties identified in lead candidate (ExoPr0)
- Scope to tailor endogenous exosomes to specific targets via loading and/or producer cell modification

Broad Anti-Cancer Efficacy of ExoPr0 *in vitro*

- *In vitro* screen for ExoPr0 efficacy in a truncated NCI60 cell line panel
- Anti-proliferative responses observed across multiple tumour types – approx. 1/3 of all lines tested showed evidence of positive response to ExoPr0



Expected Clinical Milestones

CTX for stroke disability

- Early 2020 – Phase IIb data

hRPC for retinitis pigmentosa

- Mid-2019 – Top line Phase I/IIa data
- 2019 – Phase IIb commencement

hRPC for cone-rod dystrophy

- 2019 – Phase II commencement

Exosomes for cancer (solid tumours)

- 2019 – Phase I commencement

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