

The logo for ThromboGenics features a large, stylized 'TG' monogram. The 'T' is a solid red shape on the left, and the 'G' is a light gray shape on the right. The letters are partially overlapping. To the right of the monogram, the word 'ThromboGenics' is written in a red, sans-serif font.

# ThromboGenics

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An Emerging Force in Vascular Medicine

*Kempen & Co. Life Sciences Conference,*

*Brussels, April 3, 2008*

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## Disclaimer

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# Agenda

- ThromboGenics Today
- Microplasmin – a significant market opportunity
  - Microplasmin for the « back of the eye » diseases
  - Microplasmin for vascular occlusion
- Staphylokinase - a thrombolytic for developing countries
- Novel antibody programmes
  - TB-402, a long acting anticoagulant
  - TB-403, an innovative cancer therapeutic
- Key Financials
- Conclusion



## ThromboGenics Today

- A broad maturing pipeline of novel biopharmaceuticals for treatment of a range of vascular diseases
  - Six clinical programs – focused on back of the eye and thrombotic disease indications
  - One additional program will enter the clinic in Q108 - oncology
  - Two exciting earlier preclinical programs
- Experienced management team with world leading expertise in vascular disease
- Cash position of €49.3m as of 30<sup>th</sup> June 2007
- 50 people, with offices located in Belgium, Ireland, and the United States
- Listed on Euronext Brussels – ticker : THR



## An experienced management team

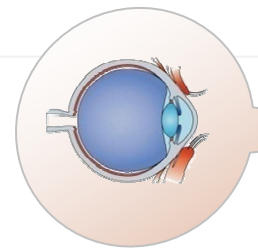
- Désiré Collen, MD, PhD      Chairman  
Chief Executive Officer
- Patrik De Haes, MD      Chief Operating Officer
- Chris Buyse      Chief Financial Officer
- Stuart Laermer, MSc, MBA      Chief Business Officer
- Steve Pakola, MD      Chief Medical Officer
- Jean Marie Stassen, PhD      Senior Director, R&D



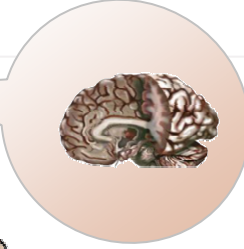
# Strategic Focus

## Back of the Eye Disease

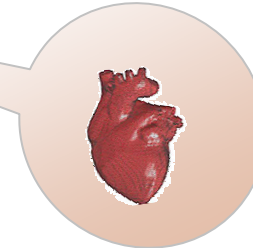
- Vitrectomy
- Diabetic Retinopathy



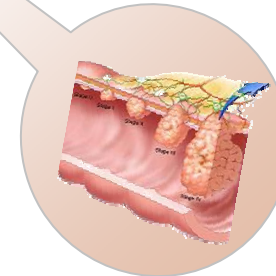
## Ischemic stroke



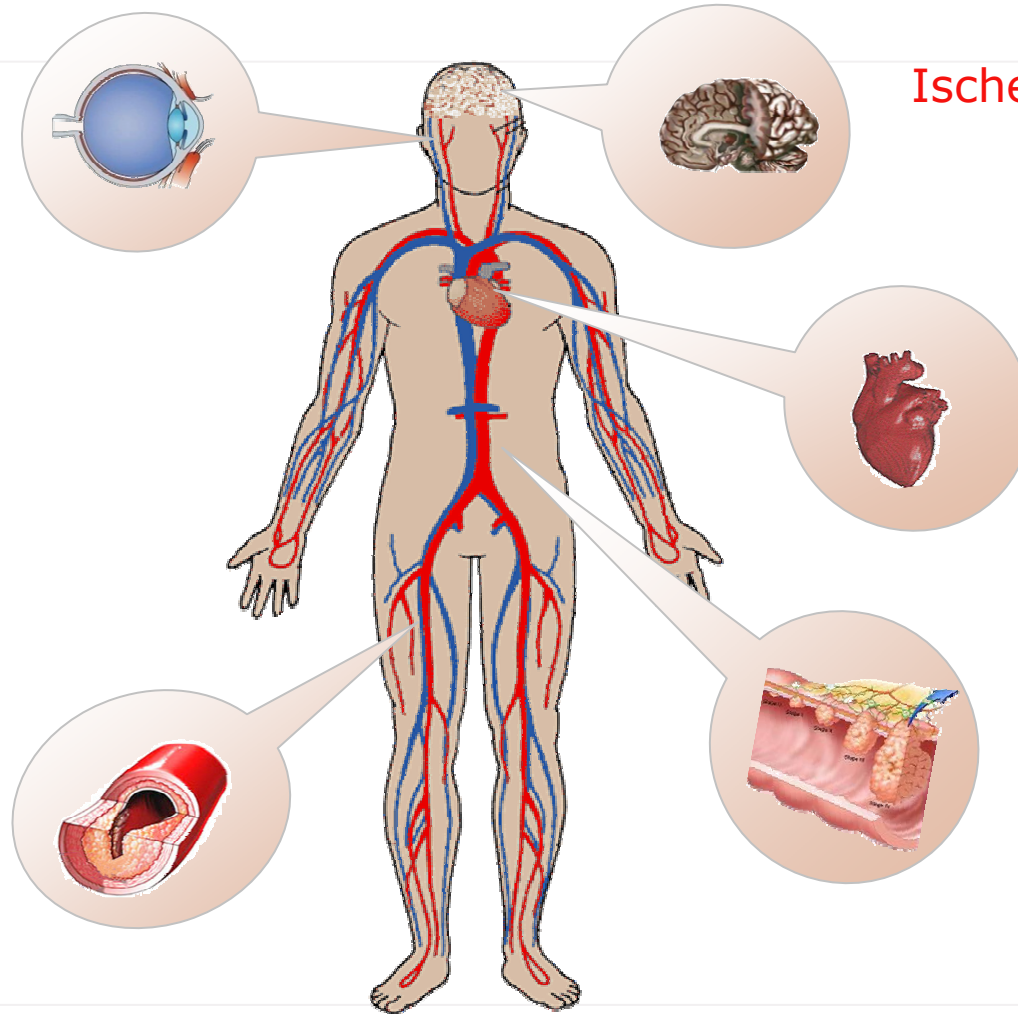
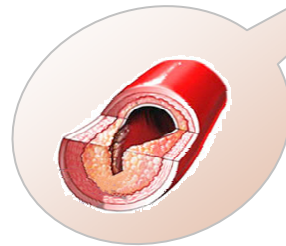
## Heart attack



## Cancer

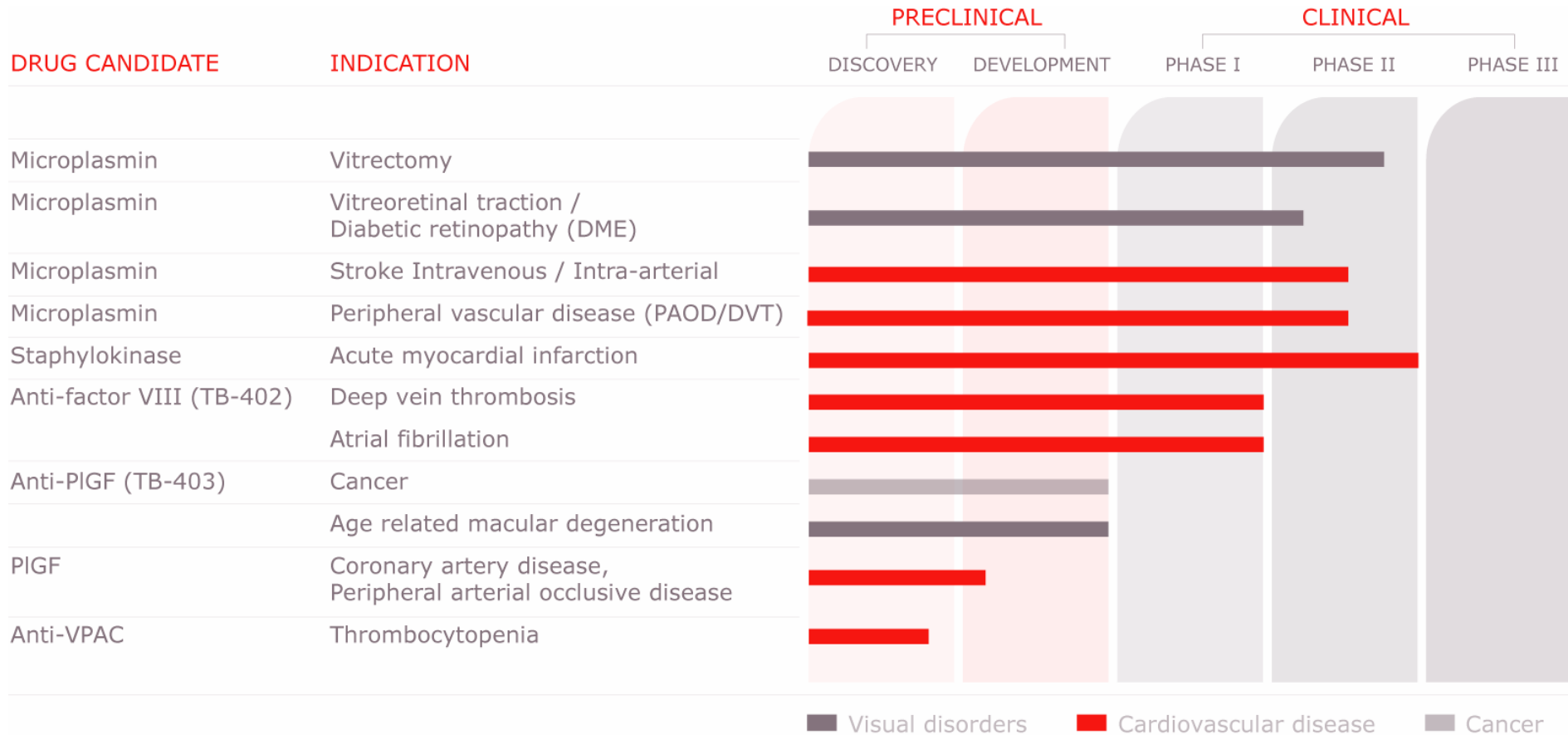


## Vascular Thrombosis





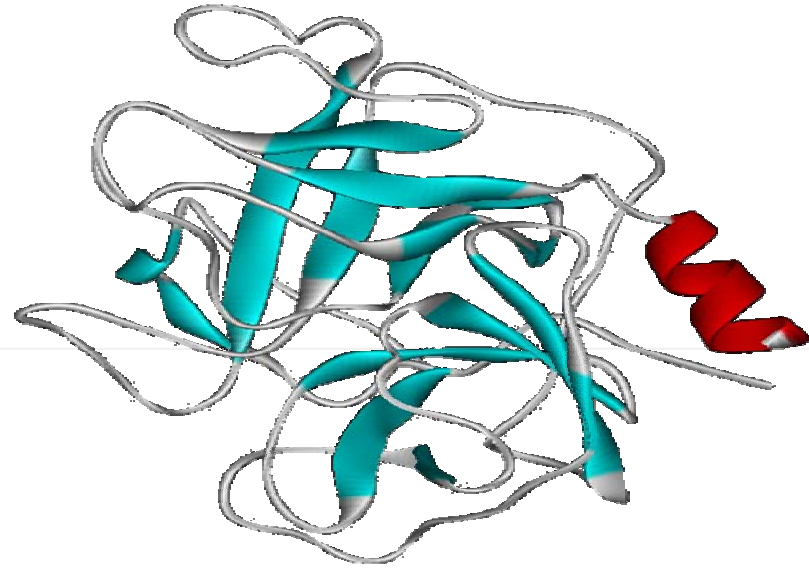
# Our broad pipeline





# ThromboGenics

Microplasmin



*A product with significant commercial potential  
via multiple indications*





## Microplasmin – Significant commercial opportunities

- A truncated and stable form of plasmin
- Microplasmin is a proteolytic enzyme
- Back of the eye diseases
  - Dissolves the protein structure linking the vitreous to the retina
- Thrombotic diseases (stroke, peripheral vascular disease)
  - Dissolves the fibrin structure which is a fundamental element of blood clots

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# TG ThromboGenics

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Microplasmin for “back of the eye” diseases

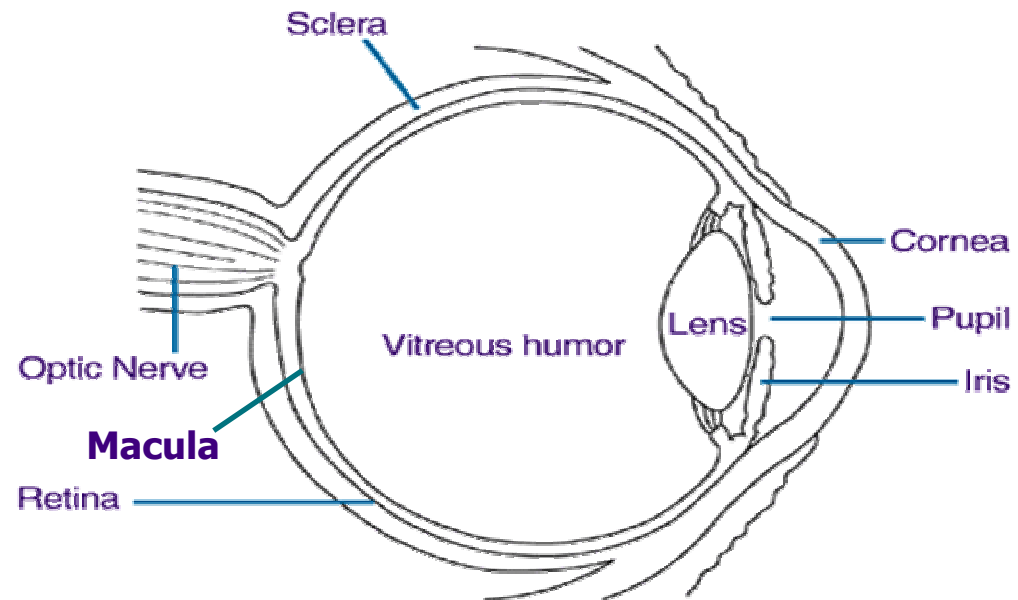
*New approach towards major retina diseases*

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# Anatomy of the eye



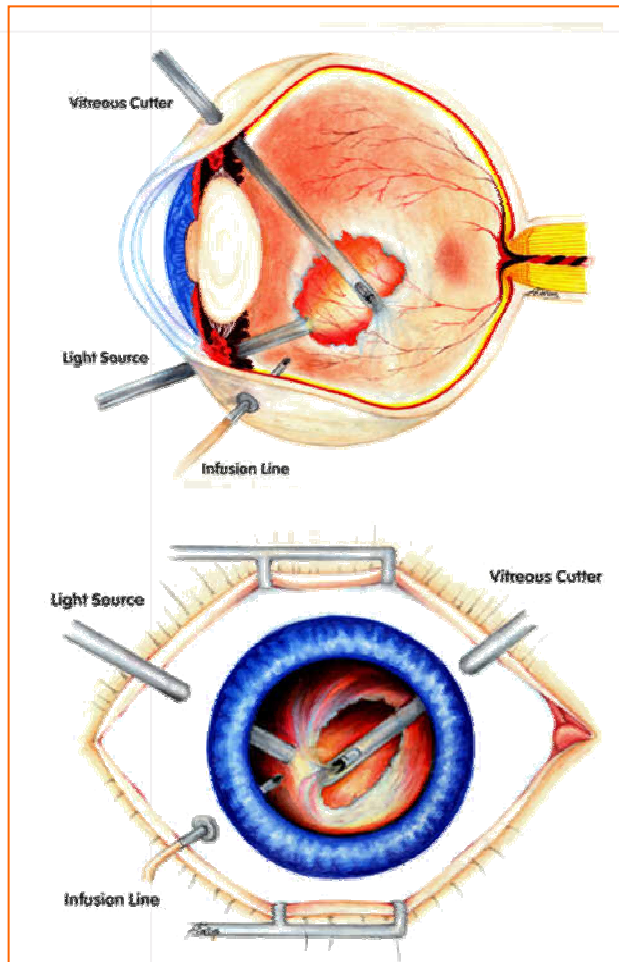


## Significant market opportunity

- PVD (posterior vitreous detachment) is condition where the vitreous is disconnected from the retina
- Currently PVD is induced surgically in a procedure called a vitrectomy to treat a range of back of the eye diseases
  - Vitrectomy - 600,000 cases worldwide annually
  - 6-8% growth per year
- Microplasmin is initially being developed as an adjunct to vitrectomy and maybe used to achieve a PVD non-surgically
- Potential non-surgical preventative for Diabetic Macular Edema and Proliferative Diabetic Retinopathy - \$1 billion plus market opportunity



## Vitrectomy : a surgical procedure to induce PVD



- The procedure is performed in an operating room under general or (occasionally) local anesthesia
- The procedure involves the removal via suction of the vitreous gel through a very small ( $\sim 1.4\text{mm}$ ) incision in the eye wall, hence the name "vitrectomy" – the vitreous gel is replaced by an isotonic liquid solution
- Vitrectomy is a microsurgical procedure used to repair retinal disorders:
  - Diabetic vitreous hemorrhage
  - Retinal detachment
  - Epiretinal membrane
  - Macular hole

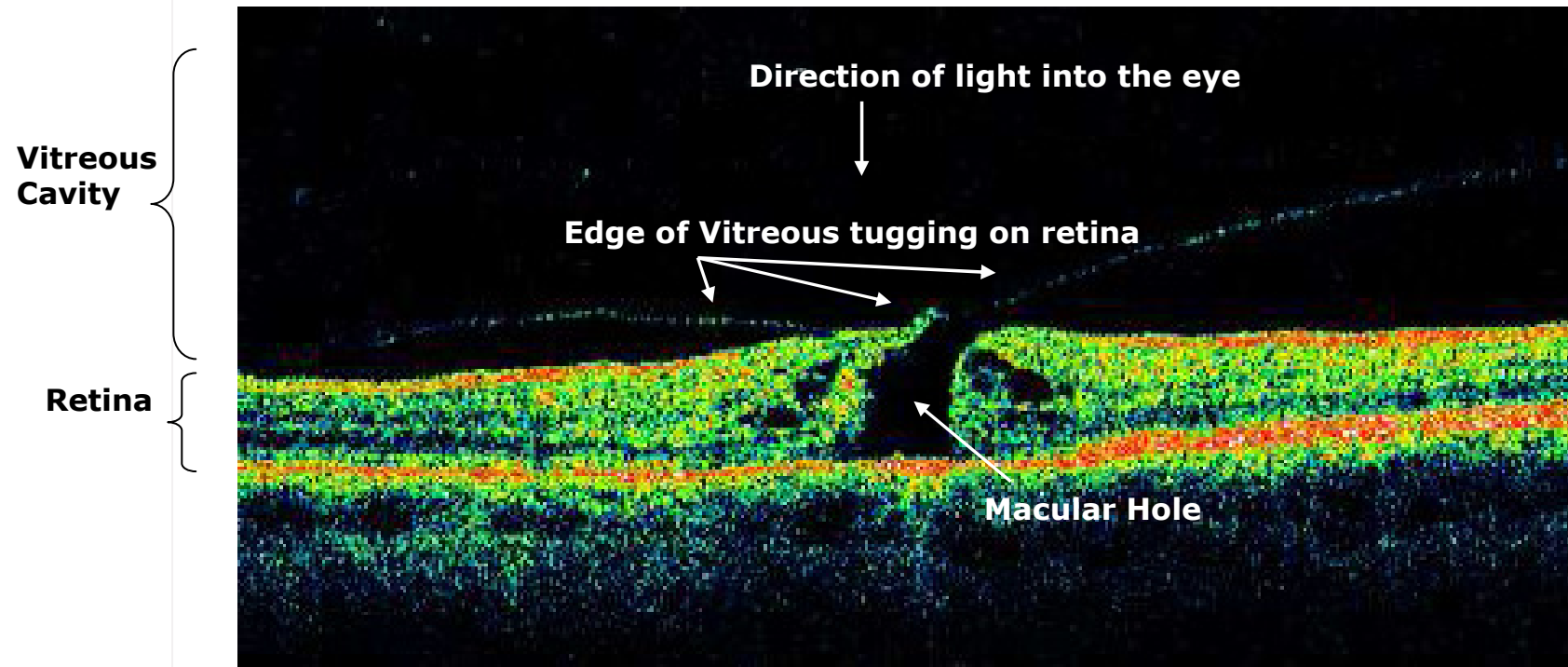


# Microplasmin for Eye disease clinical development overview

	Vitreotomy setting		Non surgical setting	
	MIVI-I	MIVI-III	MIVI- IIT	MIVI-II (DME)
<b>Clinical</b>	Phase IIa	Phase IIb	Phase IIa	Phase IIa
<b>Objectives</b>	Safety and efficacy (PVD induction with reduced suction)	Safety and efficacy	Safety and efficacy in vitreomacular traction	Safety and efficacy in DME
<b># Patients</b>	60	120	45 (one cohort added)	60
<b>Protocol</b>	<ul style="list-style-type: none"> <li>- open label</li> <li>- dose ranging</li> <li>- exposure time</li> <li>- multi center in EU</li> </ul>	<ul style="list-style-type: none"> <li>- randomized, placebo controlled</li> <li>- double masked</li> <li>- dose ranging</li> <li>- exposure time</li> <li>- multi center in US</li> </ul>	<ul style="list-style-type: none"> <li>- Sham injection controlled</li> <li>- dose ranging</li> <li>- exposure time</li> </ul>	<ul style="list-style-type: none"> <li>- Sham injection controlled</li> <li>- dose ranging</li> <li>- exposure time</li> </ul>
<b>Results</b>	Successfully completed <b>50% of patients at 25 µg showed PVD on day 7</b>	Due in 2008 <b>Encouraging patient cases observed</b>	First 2 cohorts completed <b>Treatment effect shown</b>	Due in 2008

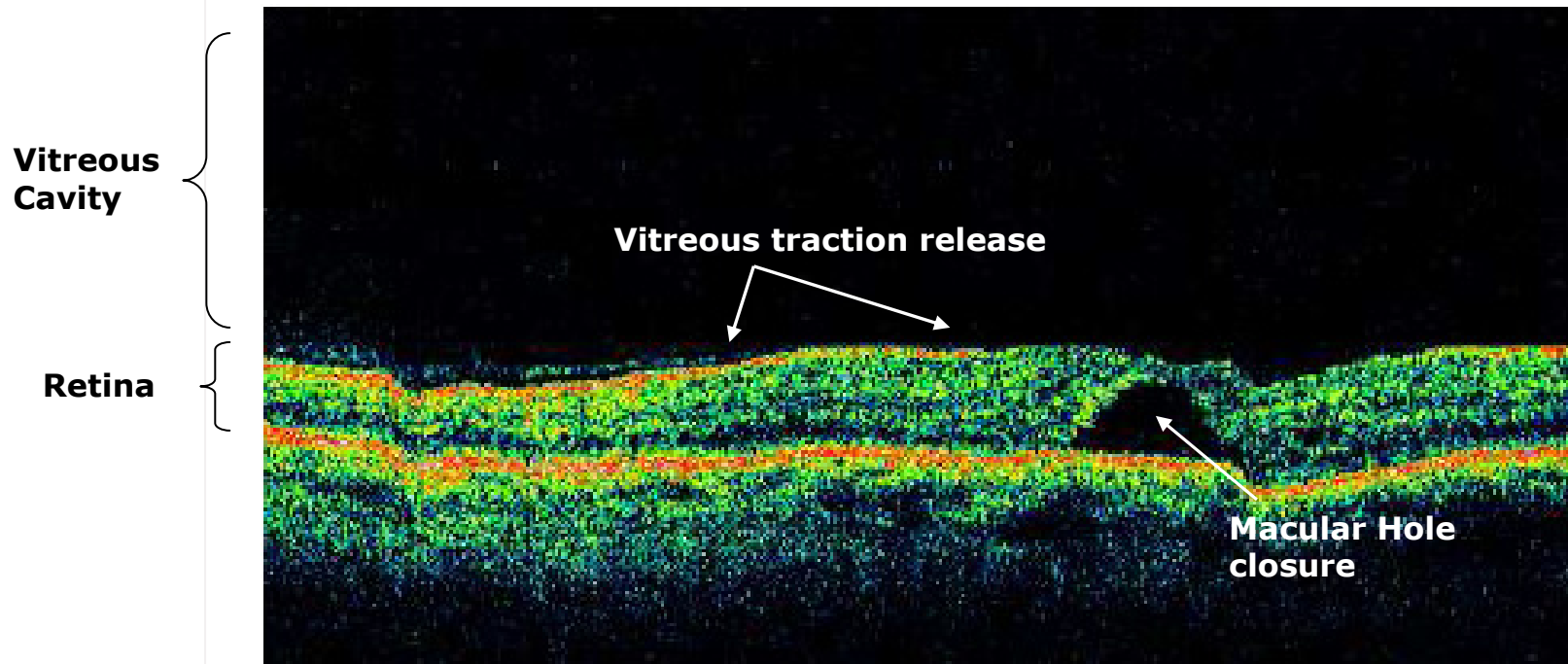


## MIVI-I - Pre injection of microplasmin





## MIVI-I - 48hrs post injection of microplasmin

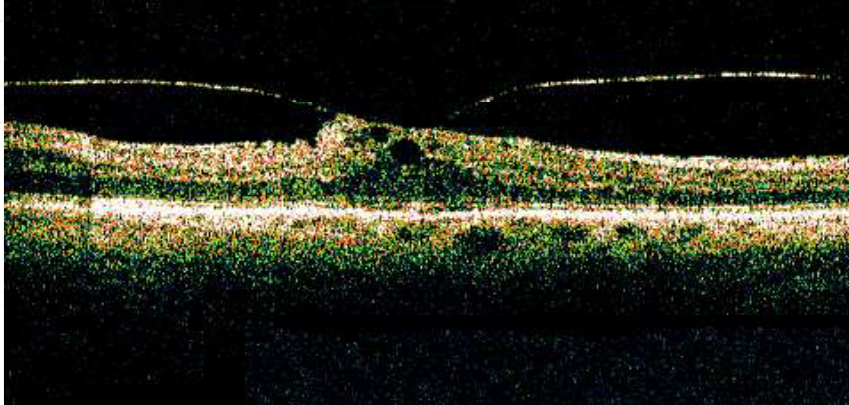




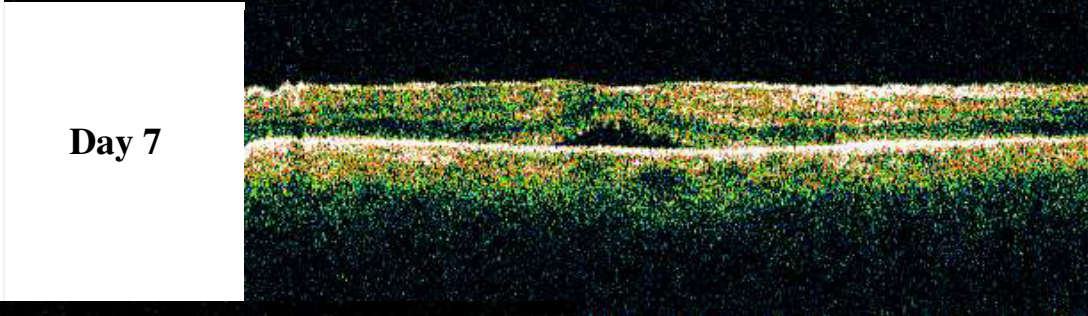
# MIVI II results

## Traction case

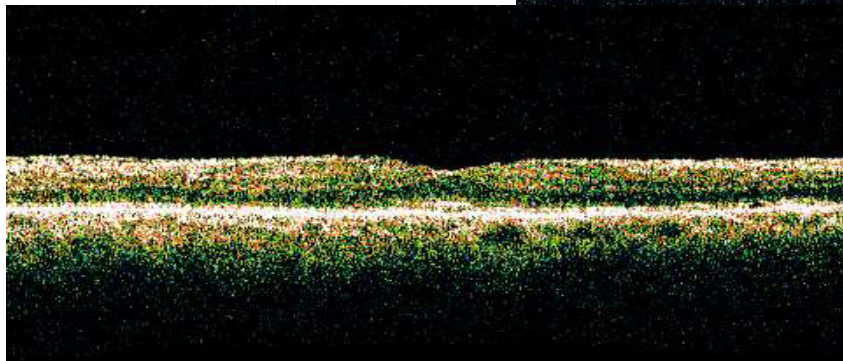
**Baseline**



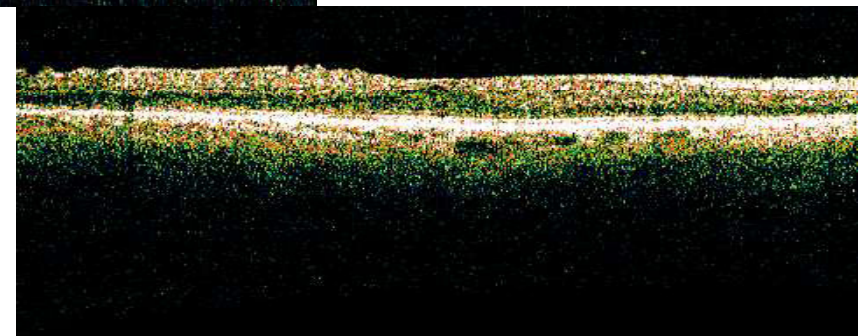
**Day 7**



**Day 28**



**3 Mths**

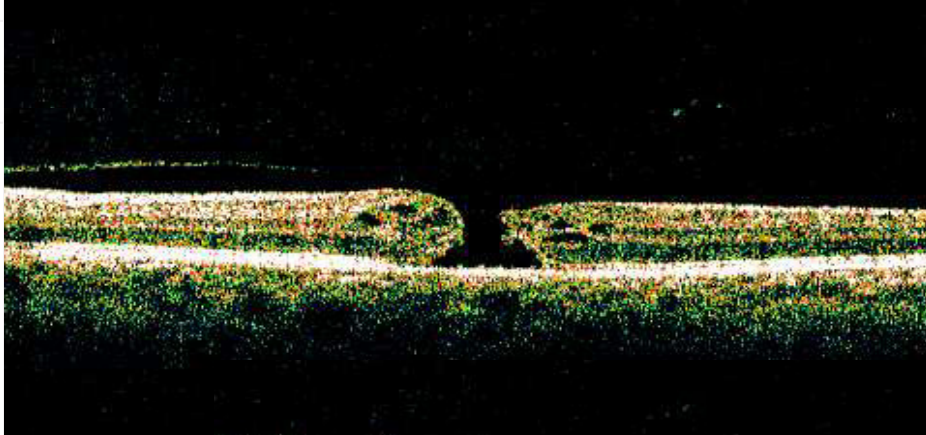




# MIVI II results

## Macular hole case

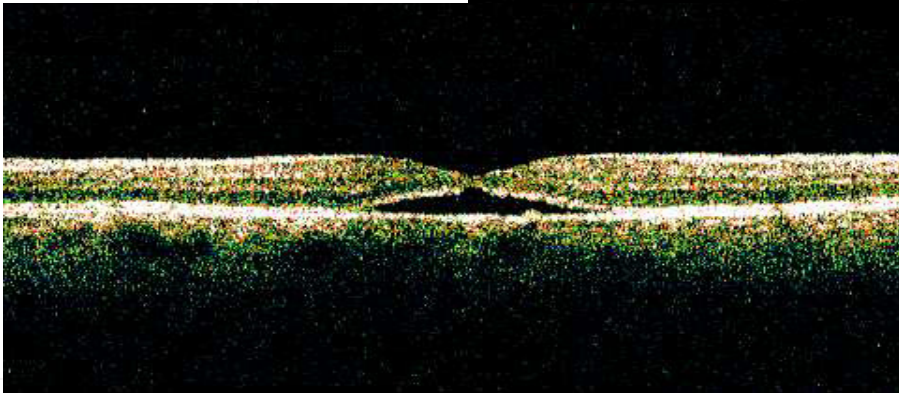
**Baseline**



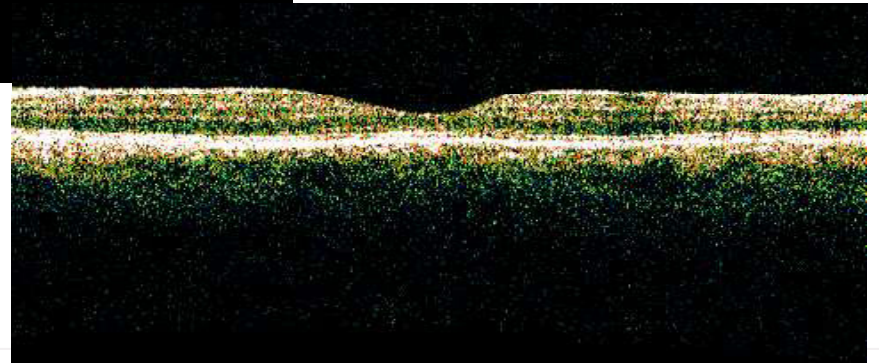
**Day 7**



**Day 28**



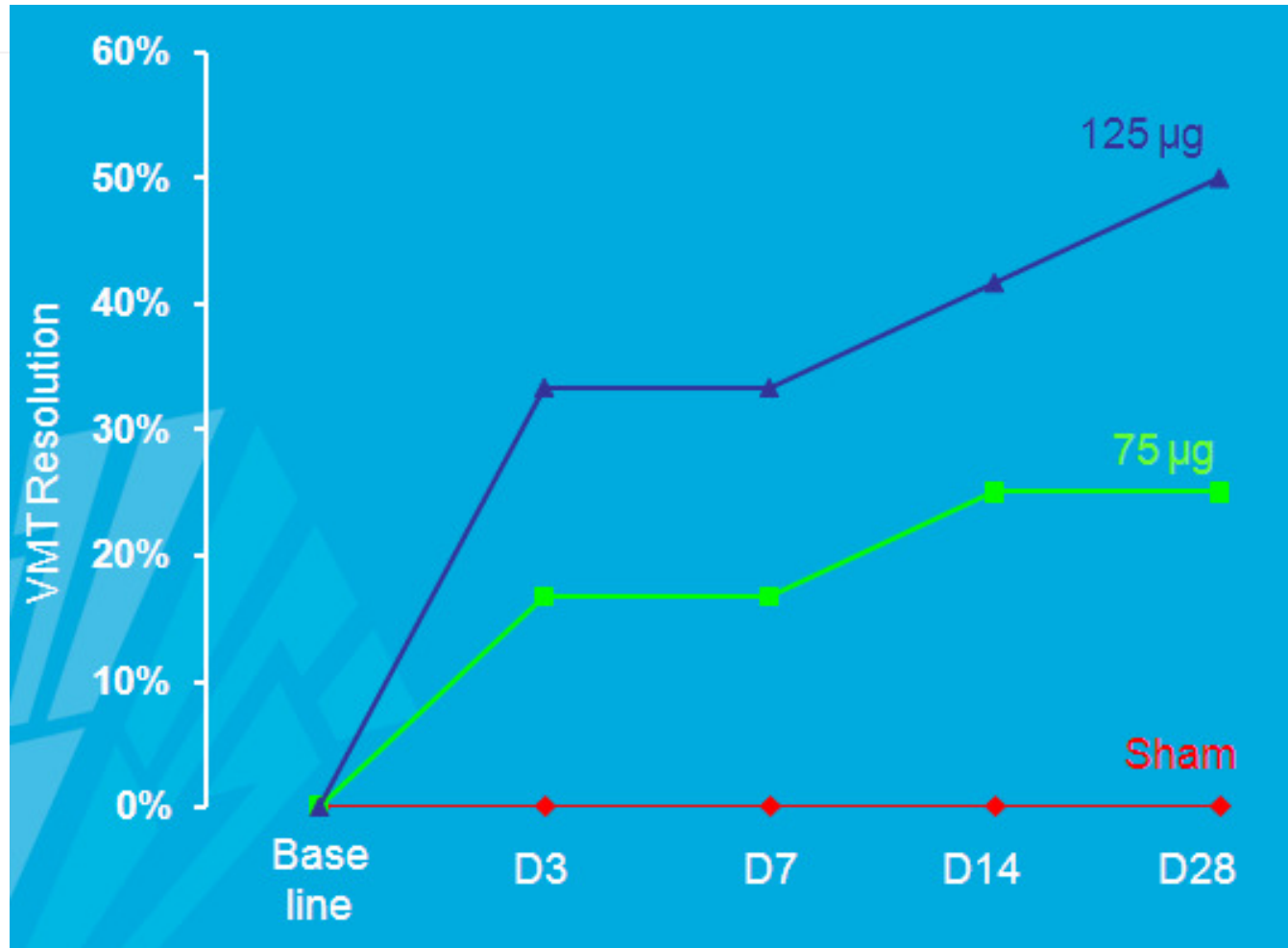
**6 Mths**





# MIVI II results

## Efficacy results





## Microplasmin - a potential alternative to vitrectomy

### **Encouraging data**

- Ability to achieve spontaneous PVD without need of suction (MIVI-I)
- Well tolerated (MIVI-I)
- Some patients cases showed non surgical traction release and macular hole closure (MIVI-IIT and MIVI-III)
- Microplasmin alone maybe sufficient to induce PVD

### **Next steps**

- Continue to Phase III by second half of 08 possibly with co-development partner

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# ThromboGenics

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Microplasmin for vascular occlusion

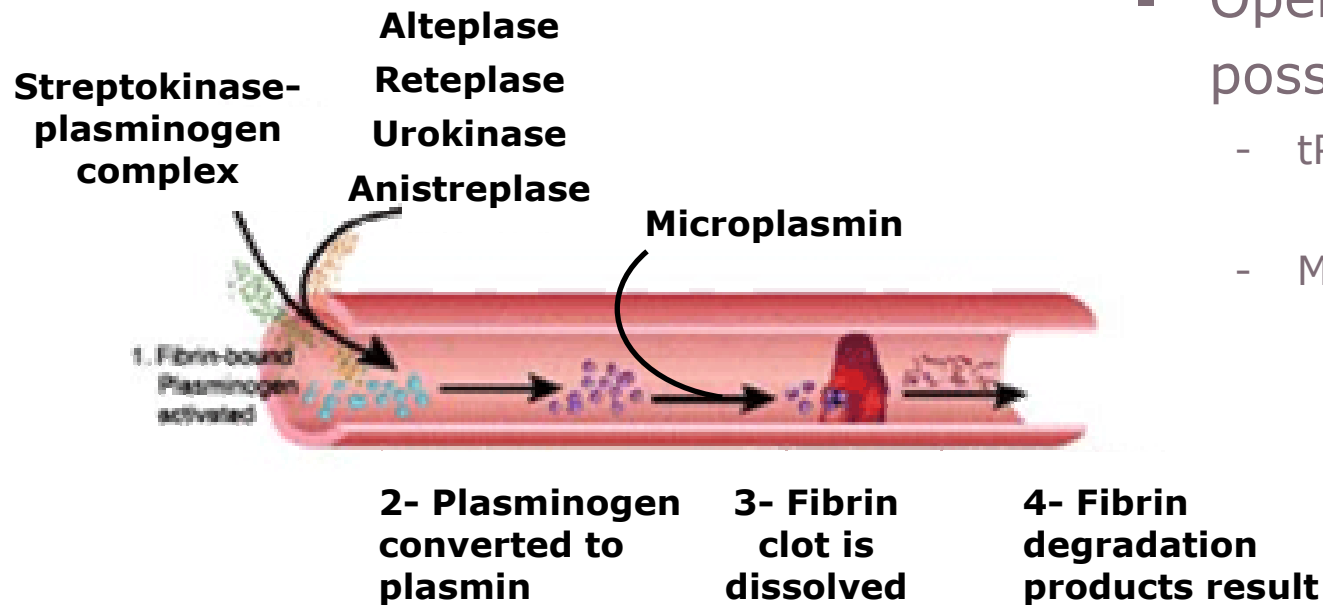
*A direct acting trombolytic*

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# Microplasmin for vascular occlusion



- Open the vessel as soon as possible
  - tPA a.o.
    - Indirect via plasminogen
  - Microplasmin
    - Direct - plasminogen independent



## Microplasmin for vascular occlusion

- No relevant side effects in Phase I up to 4 mg/kg
- Improved risk/benefit: rapid inactivation reduces risk of systemic / intra-cranial bleeding events
  - Broad applicability: Stroke and DVT
  - Potentially longer therapeutic time window than offered by tPA in Stroke



# Microplasmin for vascular occlusion

## Overview clinical development

	<b>Stroke IV</b>	<b>DVT</b>
<b>Clinical</b>	Phase IIa	Phase IIa
<b>Objectives</b>	Assess safety and efficacy	Assess safety and efficacy
<b># Patients</b>	40	Up to 15 patients
<b>Protocol</b>	- Multi centre - Double blind placebo controlled	- Single centre - Open label ascending dose study
<b>Results by</b>	2008	2008



ThromboGenics

Staphylokinase

*A thrombolytic for developing countries*





## Staphylokinase (THR 100) for heart attack

- Completed Phase II clinical trials
- Administered to over 900 patients
- Much more fibrin selective than tPA
- Efficacy as good as tPA
- Much less expensive than tPA
- More effective than streptokinase
- Aim to replace streptokinase as the “standard of care” in developing markets



## Staphylokinase (THR 100) for heart attack

- Approval in U.S./Europe requires large, expensive non-inferiority, mortality trials vs. tPA
- In contrast, clinical trials in less developed markets will only need to demonstrate that SAK restores arterial patency in small number of patients
- Developing countries represent a significant commercial opportunity, of the order of \$200 million
- Signed license agreement with Bharat in December 2006  
Technology transfer finished: August 2007
- Launch 2008/2009

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# ThromboGenics

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Novel antibody programmes

*Productive collaboration with BioInvent*

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## BioInvent collaboration

- Signed in 2004 for the co-development of antibody-based drugs for vascular indications
- Novel antibody drugs based on ThromboGenics research
- Investment on a 50/50 basis
- Return and profit sharing on a 60/40 basis in favour of ThromboGenics



## Anti-Factor VIII (TB-402)

### Long acting anti-coagulant

- Fully human antibody against Factor VIII, an essential blood clotting factor
- Long acting anti-coagulant for treatment and prevention of Deep Vein Thrombosis. Also been developed for the prevention of emboli in Atrial Fibrillation
- Only partially inhibits coagulation: expected to lead to a lower risk of spontaneous bleeding
- Phase I enrolment completed – presentation at American Society of Hematology, Dec 10
- Preliminary clinical data supports one-time treatment for acute indications; monthly for chronic indications
- Phase II expected to start in H2 2008



## Anti-PIGF (TB-403)

An innovative cancer therapeutic

- Monoclonal antibody which inhibits blood vessel growth
- Inhibits PIGF (placental growth factor), a homologue of VEGF (vascular endothelial growth factor)
- PIGF activates VEGFR-1 (Flt-1) whereas VEGF stimulates VEGFR-2 (Flk-1)
- Inhibition of VEGF proven commercially: Avastin (Genentech) - Sales of \$2.5 billion in 2006
- TB-403 acts via inhibition of VEGFR-1 (Flt-1) instead of VEGFR-2 (Flk-1)
- Targeting favorable risk/benefit ratio compared to Avastin
- To be developed for cancer and age-related macular degeneration (AMD)
- Preclinical development completed and published in Cell, Nov 2, 2007
- Start phase I-studies planned for Q1 2008



ThromboGenics

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Key Financials

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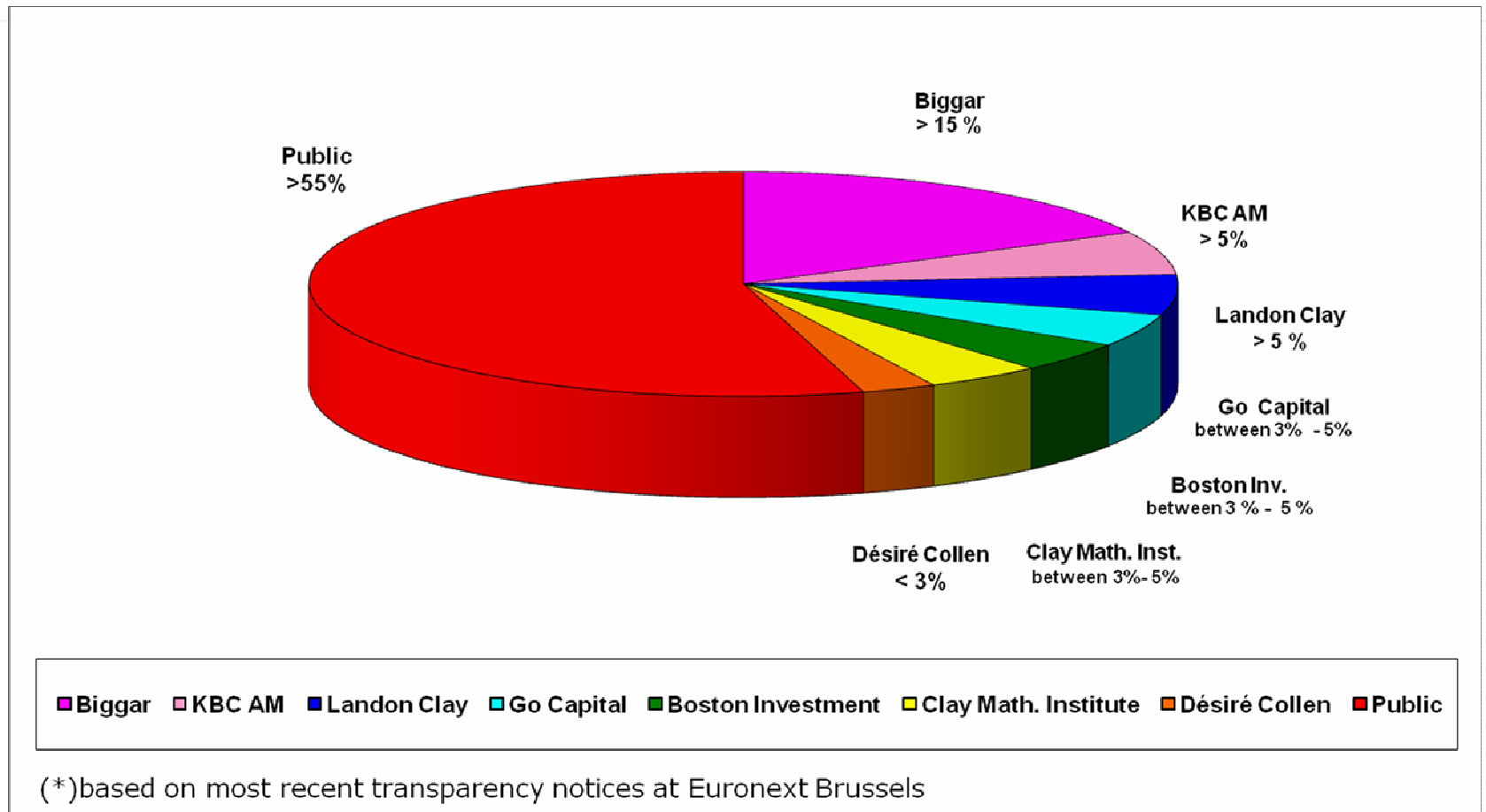


## Pro-active financial management

July '06:	<b>Listing</b> Euronext Brussels Raising of <b>€35m</b>
December 2006:	<b>First Bel Small Award 2006</b>
February 2007:	Increase Free float (GO Capital takes 5%)
May 2007:	Raising of <b>€24m</b> Broadening shareholders base <b>Increase Free float &gt; 70 %</b>
July 2007:	<b>Merger</b> ThromboGenics NV – Thromb-X NV
September 2007:	<b>Capital Increase</b> – Warrant exercise raises <b>€5.1m</b>



# Our current shareholders update December 2007 (\*)





## Cash situation

- Cash Position as of end June 2007: €49.3m
- Warrant exercise raised an additional €5.1m in September 2007

Sufficient cash to support the Company's Operating Plan for at least the next 2 years



## Financial calendar

- Full year results 2007: **13 March 2008**  
(Period covering May '06 – December 2007)
- Annual Shareholders' Meeting: **06 May 2008**



ThromboGenics

Conclusion





## Key forthcoming milestones

<b>Drug Candidate</b>	<b>Event</b>	<b>Timing</b>
Microplasmin in Eye	Completion of patient enrolment of MIVI-III Completion of patient enrolment of MIVI-II DME GMP production for Phase III ready	Q1 08 mid-2008 mid-2008
Microplasmin in Vascular	Completion of patient enrolment of Stroke-IV Completion of patient enrolment in DVT	H1 08 H2 08
Staphylokinase	Initiation of Phase III clinical trials	H1 08
TB-402	Initiation of Phase II clinical trials	H2 08
TB-403	Initiation of Phase I clinical trials	Q1 08



# ThromboGenics : market potential

TG Product and Indication	Gepland	Aantal patienten of behandelingen	Kost Markt (Mio)	
<b>Microplasmin/Eye</b>				
Vitrectomy	H1/2011	600,000 vitrectomies	€ 900	€ 540
Diabetic retinopathy	2013	5,200,000 pts (G7) x 1.3 eyes	€ 900	€ 6,084
Diabetic macular edema	2012	1,875,000 pts x 1.3 eyes	€ 900	€ 2,194
<b>Microplasmin/Vascular</b>				
Stroke	2012	1,050,000 pts	€ 2,500	€ 2,625
Deep Vein Thrombosis	2013	1,125,000 pts	€ 2,500	€ 2,813
<b>Staphylokinase</b>				
Heart Attach	2010	4,500,000 pts in developing countries	€ 50	€ 225
<b>TB-402 (Anti-Factor VIII)</b>				
Deep Vein Thrombosis	2014	(Lovenox sales €2.1 bn)	€ 100	€ 840
Atrial fibrillation	TBD	4,400,000 pts x 12 treatments/yr	€ 100	€ 5,280
<b>TB-403 (Anti-PIGF)</b>				
Cancer	2014	100,000 pts - cfr Avastin	€ 50,000	€ 5,000
				€ 25,600



## ThromboGenics – Wrap-up

- Exciting clinical portfolio progressing according to plan
- Microplasmin offers a significant opportunity in back of the eye disease – clinical data to-date have been very encouraging
- Microplasmin as a thrombolytic targets indications with clear unmet medical needs
- Staphylokinase – clear route to value creation
- Exciting opportunities with novel antibody therapeutics - TB-402 and TB-403
- Experienced management focused on delivery

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# ThromboGenics

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An Emerging Force in Vascular Medicine

*[www.thrombogenics.com](http://www.thrombogenics.com)*

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