# **Medical Therapies Limited**

# **EGM Presentation**

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# Medical Therapies (ASX:MTY)

- Listed in December 2005
- © 185M shares on issue
- Midkine specialist and owner of the largest diagnostic and therapeutic patent portfolio around this novel cancer target globally
  - Developing early cancer diagnostic test
  - Developing therapeutic antibody portfolio with partners

# Recent developments

## Capital structure and funding

- •Retired \$1.1M convertible debt in February 2009
- •Raised \$790K for partial debt repayment and working capital in two private placements in February and April 2009
- •Retired \$1.5M debt in lieu of ordinary shares

#### Product development

•Commenced mycoplasma testing and antibody production for Midkine ELISA

•Selected cell banking and production facility for therapeutic monoclonal antibodies for pre-clinical validation trials



# **Recent developments**

#### **Business development**

• Completed licensing and partnering data packages in the following areas

- Therapeutic anti-midkine antibody (Multiple sclerosis and rheumatoid arthritis)
- Veterinary cancer diagnostics (Early cancer diagnosis in canines)
- Human early cancer diagnostics (Midkine as a novel cancer marker)

#### Advanced licensing and partnering discussions in the following areas

- Veterinary cancer diagnostics
- Anti-midkine antibody therapeutics for MS and RA
- Midkine ELISA distribution
- Midkine protein therapy clinical collaboration for post AMI treatment

### Business strategy – diagnostic portfolio

### Commercialise MK cancer diagnostic assets

- Complete optimisation of commercial Midkine ELISA
- Set up distribution of Midkine ELISA for research
- Complete clinical validation of early cancer diagnostic
- Complete validation of canine cancer diagnostic test
- Develop in vivo imaging assay for MK with partner
- Develop rapid MK diagnostic assay with partner
- License canine cancer diagnostic test

### Business strategy – therapeutic portfolio

#### Partner or license therapeutic assets

- Set up partnership for the humanisation, optimisation and preclinical POC of the anti-midkine antibody assets
- License antibody assets for selected indications (RA, MS, restenosis)
- Complete validation trials and license anti-inflammatory antibody for veterinary applications
- Complete pre-IND package for AMI in preparation for outlicensing of MK-protein therapeutics

# Key value inflection points



The Board and management consider the above a realistic timeline for achieving these milestones, it is subject to a number of external factors including availability of funding, external service providers and timely response from regulatory authorities.



# Midkine, a novel target

- Novel target for new therapies and diagnostics against cancer, inflammation and autoimmune diseases
- Small protein (13kD, 121 AA's) with two domains
- Heparin binding growth factor prominent in embryogenesis but largely undetectable in adults
- Acts by:
  - reducing apoptosis (cell death)
  - facilitating cell migration
  - modulating angiogenesis
  - promoting cell growth
- Has important role in cancer progression, onset of inflammatory diseases and preservation and repair of injured tissue
- Strong preservation across species (and substantial animal data) provides for veterinary applications



# Midkine, a novel cancer marker

#### Detection of Midkine in serum by using our proprietary Enzyme-linked Immunoassay (ELISA)





# Midkine levels in cancer patients

#### Midkine levels were found to be elevated in 87% of cancer patients in 10 different cancers



# Midkine in cancer prognosis

Strong correlation of elevated level of blood midkine with poor prognosis of neuroblastoma



# Midkine in cancer prognosis

Five year survival curves of 93 patients with oesophageal squamous cell carcinoma show statistically significant survival rate increase amongst patients with 300pg/mlor less Midkine in the blood



Hideaki, et al. Cancer Science 2003

# Midkine antibodies for the treatment of multiple sclerosis

The role of MK in the development and severity of multiple sclerosis



MK-/- mice show delay in disease onset and reduction in disease severity after immunisation with MOG35-55

- Standard MS animal model (EAE) was used by inducing MS with the administration of MOG35-55
- Significant delay of disease onset was observed in mice without the MK gene (MK-/-)
- Significantly less severe MS like symptoms were observed in MK-/- mice
- Further pre-clinical work on the same animal model confirmed therapeutic effect of two of MTY's antibodies

#### Wang et al, Proc Natl Acad Sci U S A, 2008

# Antisense Midkine in the treatment of rectal carcinoma

Anti-midkine antisense reduced growth of rectal carcinoma cells

Anti-midkine antisense dose curve against rectal carcinoma in mice





Takei, et al. Cancer Research 2001

# The role of Midkine in rheumatoid arthritis

#### Midkine levels observed in patients with rheumatoid arthritis



## Increased levels of Midkine detected in more severe cases of RA patients

	Arthritis		Incidence
	+	-	
WT Mice	6	1	86%
MK-/-	1	9	10%
MK pump	9	3	75%

• 6/7 (86%) WT mice show symptoms using antibody induced arthritis model

RA induced in wild type and MK-/- mice

• 1/10 of MK-/- (10%) mice show symptoms using antibody induced arthritis model

•When Midkine is injected into MK-/- mice 75% show symptoms of arthritis

#### Maruyama et al, Arthritis Rheum, 2004

# Midkine protein as therapeutic

Midkine therapy following Acute Myocardial Infarction (AMI)



#### Control

Midkine treated

Mortality rate after 24 hours:

- Control: 4/12 (33.3%)
- MK-treated: 1/9 (11.1%)

#### Pig model of AMI - balloon was placed distal to the first diagonal artery for 45 minutes

- Following ischemia and reperfusion 5 microgram per kilogram MK was injected directly into the ischemic area over 10 minutes
- Single dose MK reduced mortality to one third

# Summary

- Advanced diagnostic portfolio with near term revenue potential
- Veterinary diagnostic and therapeutic portfolio with lower regulatory hurdles and close to revenue
- Early licensing revenue opportunities for high value therapeutic anti-midkine antibody and anti-midkine oligonucleotide assets
- Early licensing revenue opportunities from MK protein to facilitate recovery after heart attack
- Actively targeting sustainable and significant returns for shareholders by selective partnering around MTY's blocking MK intellectual property portfolio
- Board and management is focused on increasing shareholder value through a transparent, milestone based growth strategy

# Thank you

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# Forward looking statements

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