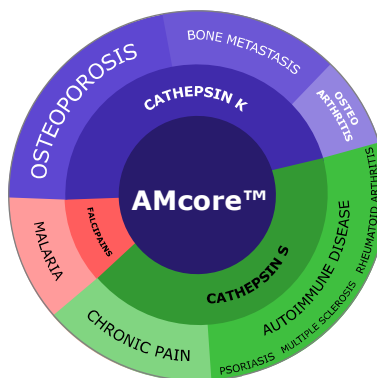




Background

Amura Therapeutics Limited is a drug discovery business specialising in the development of inhibitors of CAC1 cysteine peptidases. Cysteine peptidases are involved in several commercial and therapeutically important diseases such as **osteoporosis, bone metastasis, osteoarthritis, rheumatoid arthritis, multiple sclerosis, atherosclerosis, chronic pain** and **malaria**. Amura's compounds are derived from the proprietary **AMcore™** scaffold, which provides a turnkey solution for inhibitor design against cysteine peptidases of the CAC1 family. Amura has discovered novel chemical entities (NCE's) that are readily synthesised and exhibit excellent *in vitro* activity, primary DMPK stability, rodent and non-human primate oral pharmacokinetics, safety pharmacology, functional cell-based activity and efficacy in *in vivo* disease related animal models. Amura intends to out-license its programmes for clinical development.



Cathepsin K Inhibitors

Amura's **AMcore™** derived inhibitors have proven efficacious in a variety of industry gold standard animal models targeting blockbuster therapeutic indications. All compounds tested are via oral administration and key highlights include:-

Osteoporosis

- In a non-human primate study, multiple compounds significantly reduced serum bone markers (sCTXI and sNTx); improving upon the profile of competitor compounds that have entered the clinic. The pharmacodynamic effect is maintained for many hours upon single oral dose.
- In an ovariectomized rat model of osteoporosis, multiple compounds demonstrated a dose dependant and significant decrease in serum bone markers (sCTXI) As a result, turnover index (sCTXI/osteocalcin) was significantly decreased by all six test compounds.
- Compounds show excellent pharmacokinetic profiles, safety pharmacology and toxicity profile.

Bone metastasis

- In a human breast cancer cell-induced mouse model of bone metastasis, compound demonstrated a reduction in lesion size and total tumour burden associated with metastasis compared to the control group.
- Compounds show excellent pharmacokinetic profiles, safety pharmacology and toxicity profile.

Osteoarthritis

- In a rat surgical menisectomy model of osteoarthritis, compound showed significant beneficial effects on tibial cartilage degeneration and total joint score compared to the control group.
- Additionally, compound demonstrated a significant decrease in synovial fluid collagen markers (CTxI and CTxII) in a rat surgical menisectomy model.
- Compounds show excellent pharmacokinetic profiles, safety pharmacology and toxicity profile.

SENIOR MANAGEMENT

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Cathepsin S Inhibitors

Amura's **AMcore™** derived inhibitors have proven efficacious in a variety of industry gold standard animal models targeting blockbuster therapeutic indications. All compounds tested are via oral administration and key highlights include:-

Rheumatoid arthritis

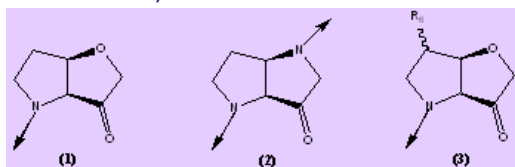
- In a collagen-induced arthritis animal model, compound demonstrated a marked reduction in collagen-specific antibodies.
- Additionally, a significant decrease in joint swelling score was observed, along with a significant improvement in joint histopathology.
- Compounds show excellent pharmacokinetic profiles, safety pharmacology and toxicity profile.

Chronic neuropathic pain

- In a rat sciatic nerve ligature model, multiple compounds demonstrated a rapid and marked inhibition of mechanical hyperalgesia.
- Inhibition was dose-dependant and fully reversible.
- Compounds show excellent pharmacokinetic profiles, safety pharmacology and toxicity profile.

AMcore™ intellectual property

The key proprietary **AMcore™** bicyclic scaffolds are:



	AU	BR	CA	CN	EU	HK	IN	ID	IL	JP	KR	MX	NZ	NO	PH	RU	SA	SG	UA	US
WO02057270	G	U	U	G	G	G	G	G	G	U	U	G	G	P	P	G	G	G	G	G
WO04007501	P	U	P	U	G	P	U		P	U	P		G		P		G	G		G

G Granted
 U Under examination
 P Pending

	2001	2002	2003	2004	2005	2006	2007	2008	Exp. date
WO02057248	Cysteine protease inhibitor								01-2021
WO02057270	Cysteine protease inhibitor								01-2021
WO02057246	Cysteine protease inhibitor								01-2021
WO02057249	Cysteine protease inhibitor								01-2021
WO04007501	Cysteine protease inhibitor								07-2022
WO07017698							AMcore™ Process1		08-2025
WO07023281							Mimetics		08-2025
WO0807132							AMcore™ Process2		07-2026
WO0807109							Selection		07-2026
WO0807103							Selection		07-2026
WO0807130							Selection		07-2026
WO0807114							Selection		07-2026
WO0807127							Selection		07-2026
WO0807107							Selection		07-2026
WO0807112							Selection		07-2026
GB0800338.6									01-2028
GB0804701.1									03-2028
GB0804792.9									03-2028
GB0809776.8									05-2028

The granted patents protect the use of **AMcore™** derived 5,5-bicyclic ketones as inhibitors of CAC1 cysteine peptidases to provide new medicines for diseases such as **osteoporosis, bone metastasis, osteoarthritis, rheumatoid arthritis, chronic pain and malaria.**

Amura intends to out-license/partner its programmes for clinical development.



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LICENSING OPPORTUNITIES

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