



Melior Discovery has pioneered an efficient and systematic approach to find new uses for existing drugs and drug candidates. By applying the unique multiplexed *theraTRACE*® indications discovery platform, Melior can quickly evaluate a compound in multiple, high quality *in vivo* models in order to test the molecule's potential use in a variety of therapeutic areas. In response to growing demands, we are now able to offer this same expertise applied to the design and execution of *in vivo* studies utilizing individual efficacy models.

theraTRACE® Platform

Uniquely powerful approach
to Indications Discovery

- not "hypothesis driven"
- highly predictive *functional* animal models
- "systematic serendipity"
- multiplexing know-how makes this possible

THROMBOSIS

In vivo and ex vivo Blood Coagulation – bleeding time models can predict anti-thrombotic/anti-coagulant potential of test compounds

- » Validated with known anti-thrombotic/anti-coagulant compounds
- » Clotting measured and recorded

BLOOD PRESSURE AND HEART RATE

- » Validated with known clinically active compounds
- » Animals monitored through implantable telemetry probes or tail cuff apparatus
- » Parameters that can be monitored in animals home cage in real time:
 - Mean arteriole pressure
 - Heart rate
 - Pulse pressure
 - Diastolic and systolic pressure
 - Activity and activity related cardiac output
- » Vein and bladder pressure can also be monitored

HYPERTENSION

Aldosterone Levels – marker for mean arterial blood pressure and electrolyte homeostasis

- » Validated with compounds known to block secretion of aldosterone or antagonize aldosterone receptors
- » Serum samples assessed for aldosterone by EIA

Angiotensin II Levels – potent vasoconstrictor that leads to sustained increase in blood pressure and sodium retention

- » ACE inhibitors and angiotensin II receptor inhibitors are effective anti-hypertensive agents
- » Serum samples assessed for angiotensin II by EIA

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