



10 February 2011

Companies Announcements Office  
Australian Securities Exchange Limited  
10<sup>th</sup> Floor, 20 Bond Street  
SYDNEY NSW 2000

**DR. ROBIN SCAIFE TO PRESENT AT THE 4<sup>TH</sup> AUSTRALIAN HIGH CONTENT IMAGE MEETING**

BPH Energy Limited [ASX: BPH] is pleased to provide a copy of the presentation that the Principal Researcher Dr. Robin Scaife will present at the 4<sup>th</sup> Australian High Content Image Meeting in Lorne, Melbourne, today.

High Content Imaging has industrialised the field of microscopy, transforming process like fluorescence microscopy from the single glass slide to a fully automated high-throughput imaging process. Since its inception over ten years ago, high-content imaging and analysis has become a pivotal process in early-stage drug discovery. BPH investee company, Molecular Discovery Systems (MDS), uses high content imaging and analysis to screen for new oncology drugs.

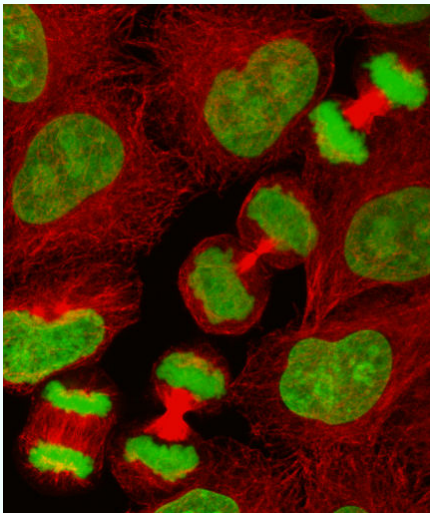
MDS has gained core expertise in high-content screening and high-throughput imaging and analysis. MDS' innovative high content imaging and analysis platform detects and quantifies cellular properties much faster than conventional methods, facilitating in the rapid optimisation and prioritisation of drug leads. MDS has effectively utilised high-content imaging and analysis to identify novel cancer drug candidates.

Yours Sincerely,

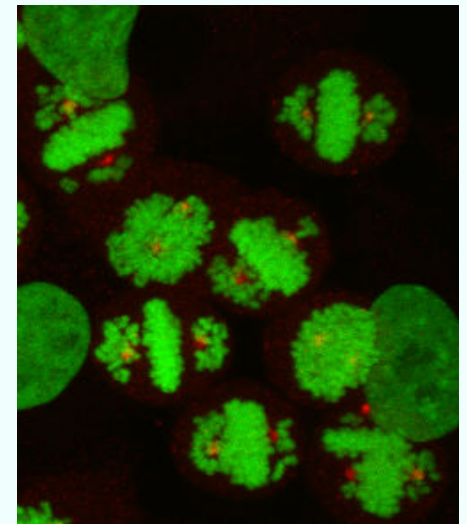
A handwritten signature in black ink, appearing to read "D. Breeze".

David Breeze  
Chairman

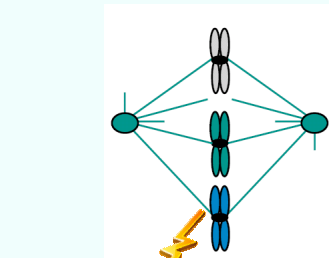
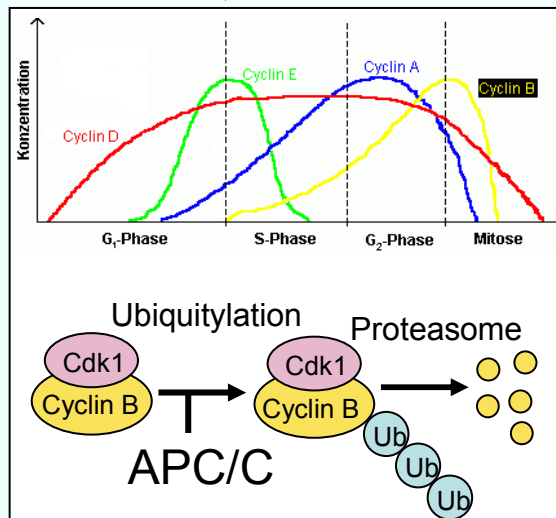
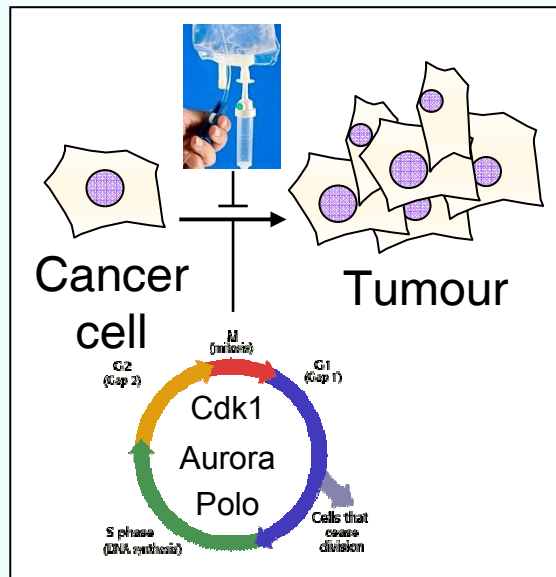
# Use of high-content imaging and analysis to screen for new oncology drugs



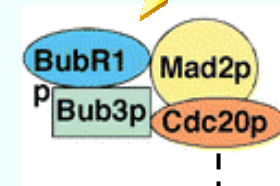
Robin Scaife  
Molecular Discovery Systems  
WAIMR



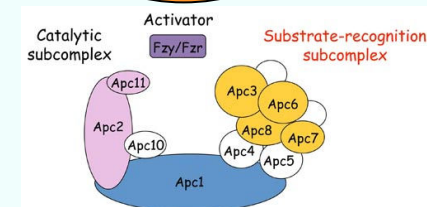
# Cyclin B Degradation by the UPS



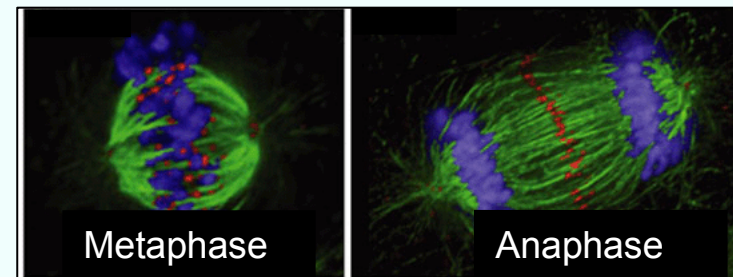
**Spindle Assembly Checkpoint**



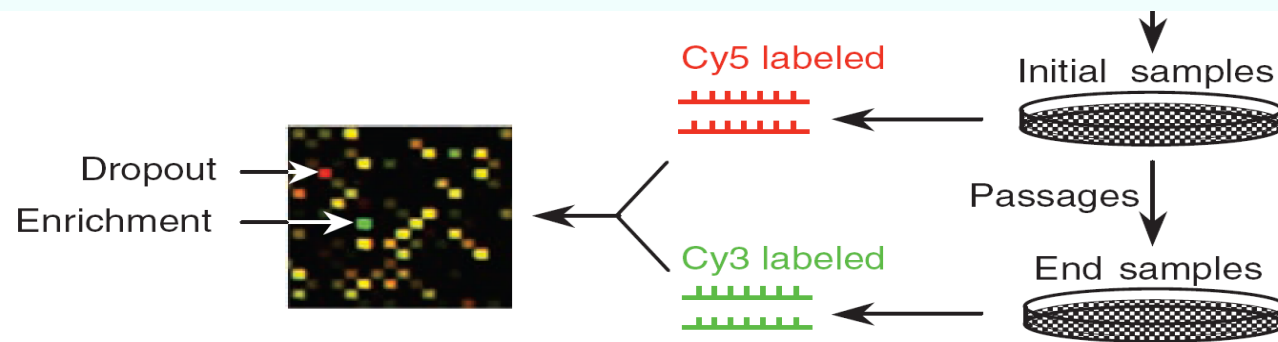
**Cdc20p**



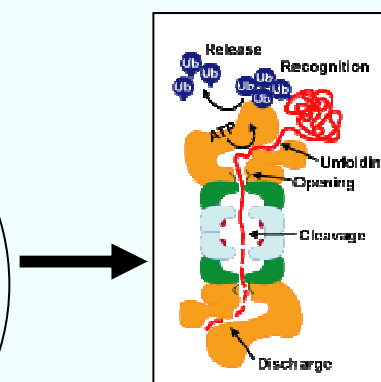
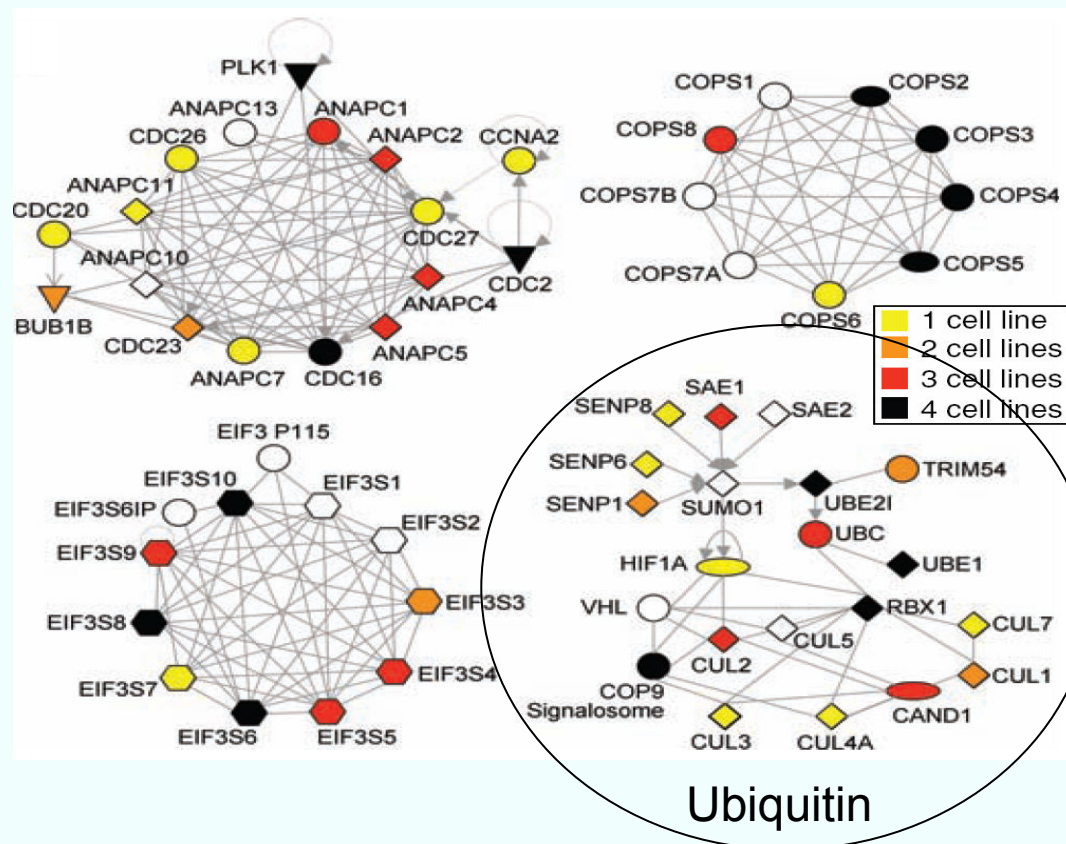
**Anaphase Promoting Complex**



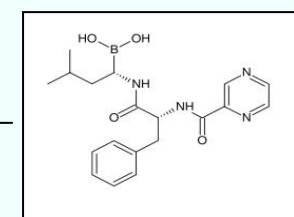
# Cell Proliferation shRNA Screen



Schlabach et al., Science (2008)

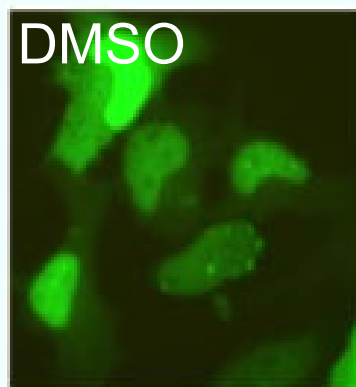


26S  
Proteasome

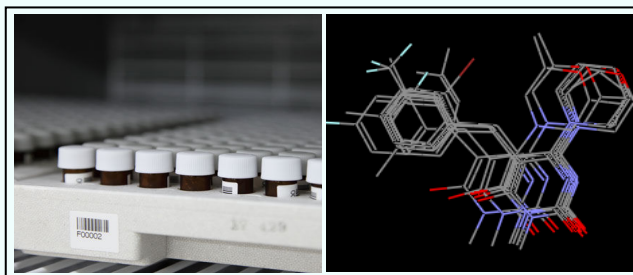
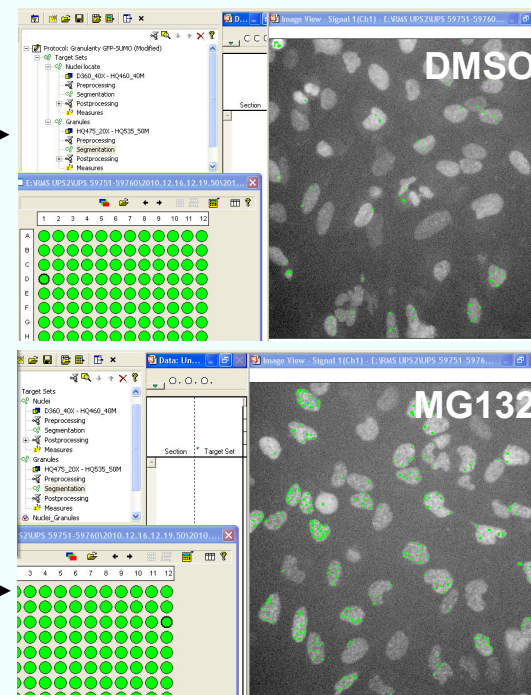


Velcade™

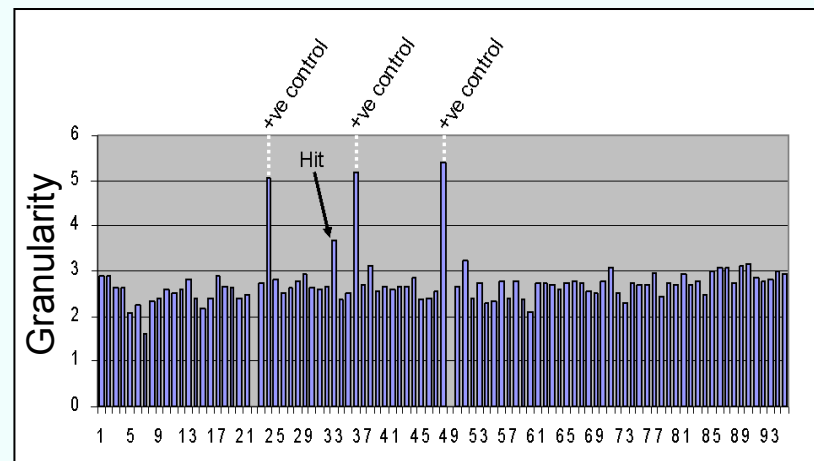
# GFP-SUMO Granularity Assay



High-content  
imaging and  
analysis

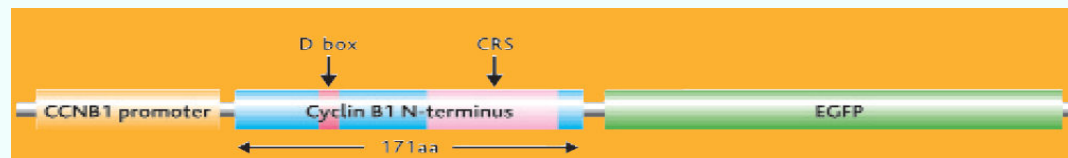
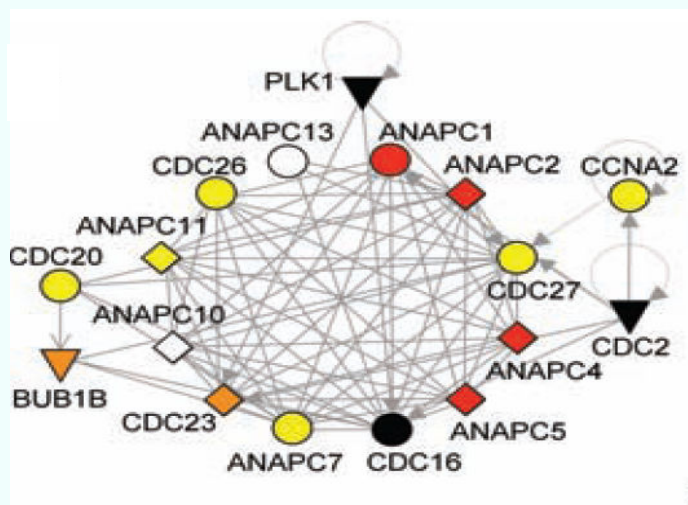
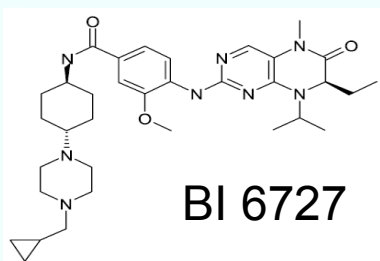


Pharmacologically Diverse

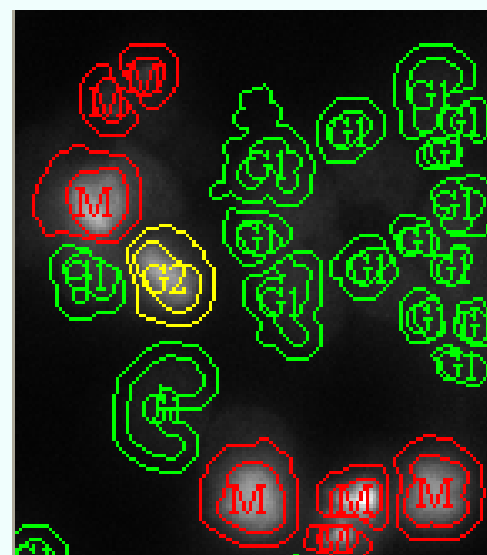
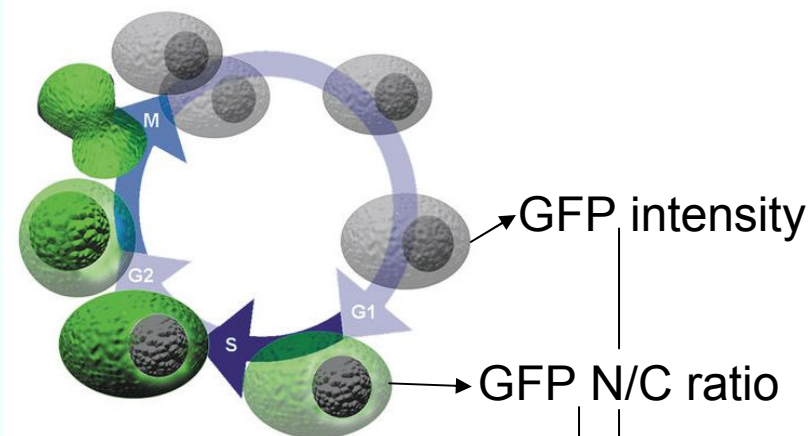




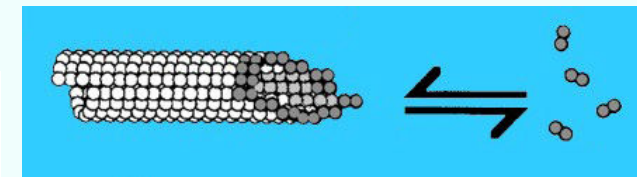
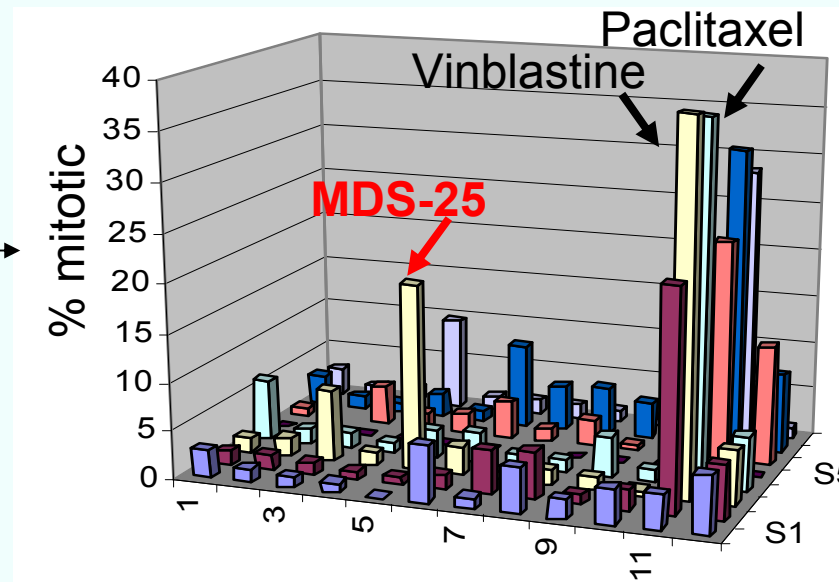
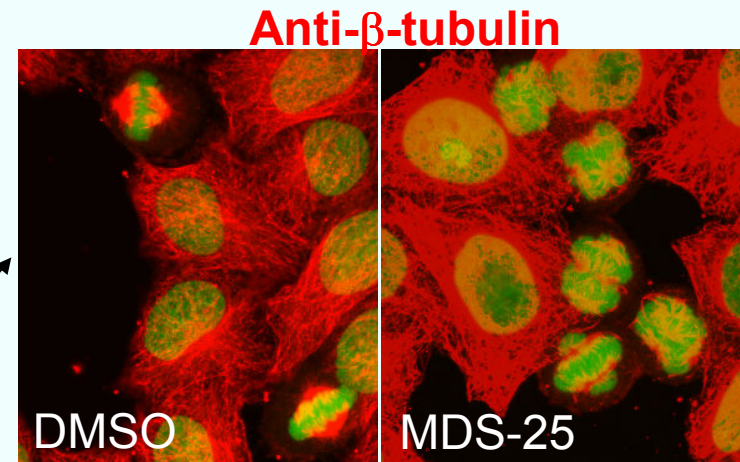
# Cyclin-B-GFP Cell Cycle Screen



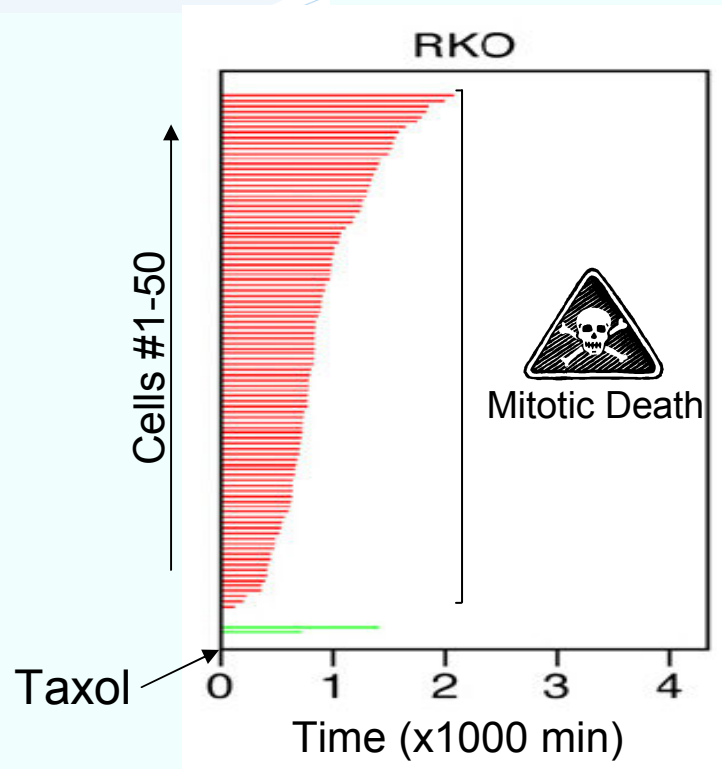
Courtesy of GE Healthcare



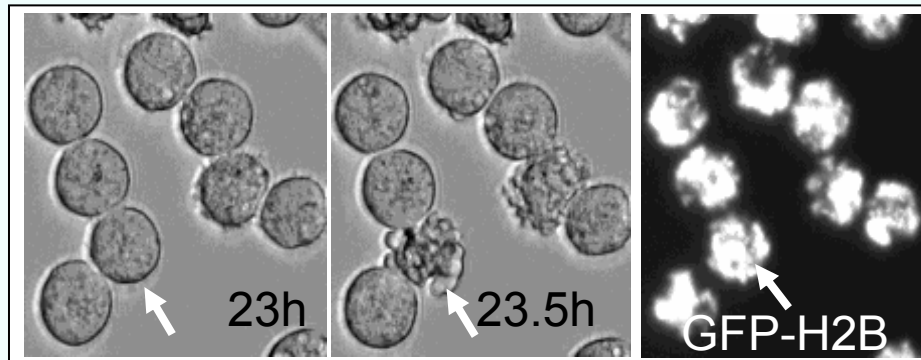
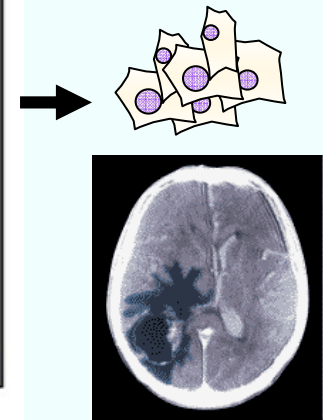
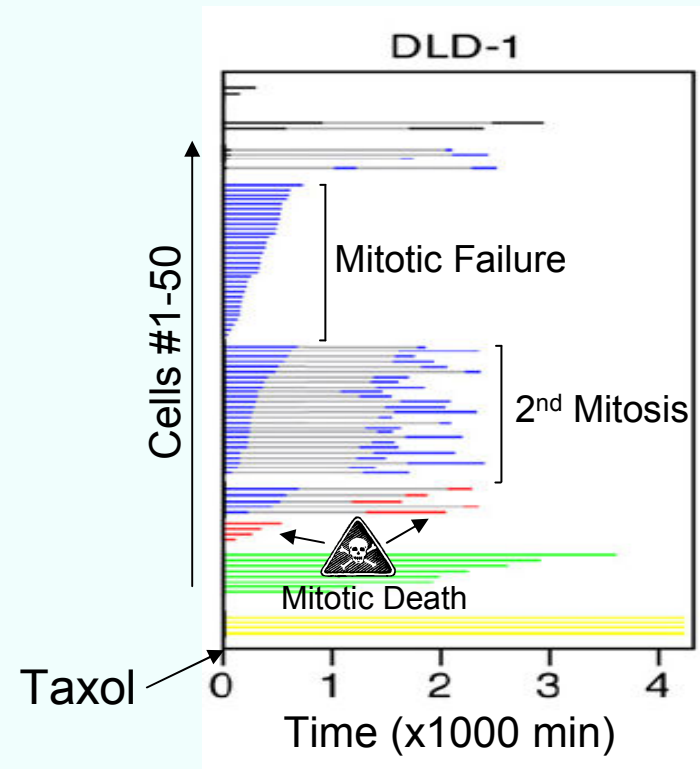
# Screening of Marine Natural Resources



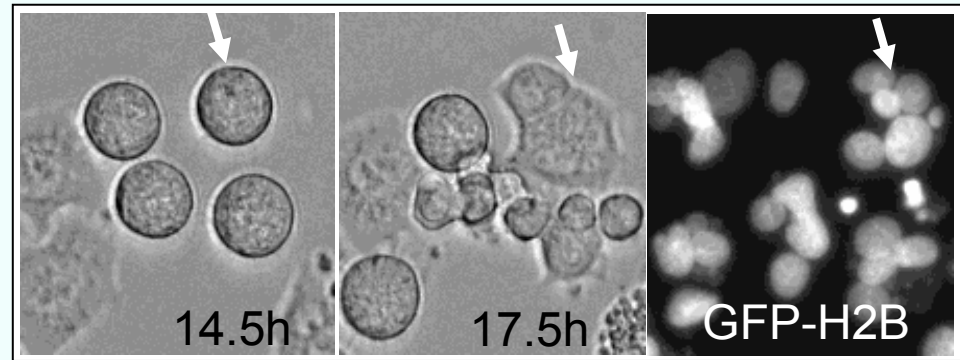
# Variation in Response to Mitotic Arrest



Gascoigne et al., Cancer Cell (2008)



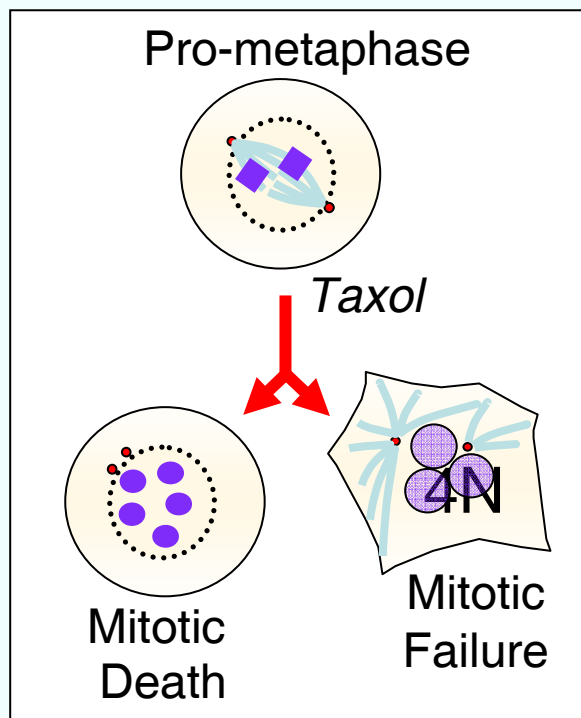
Mitotic Death



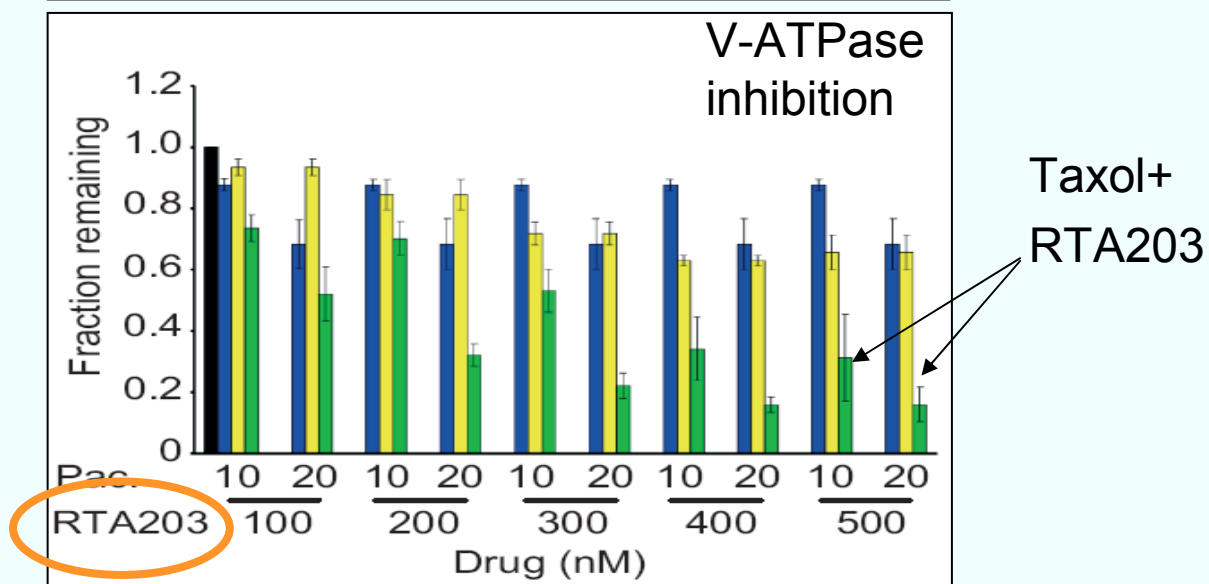
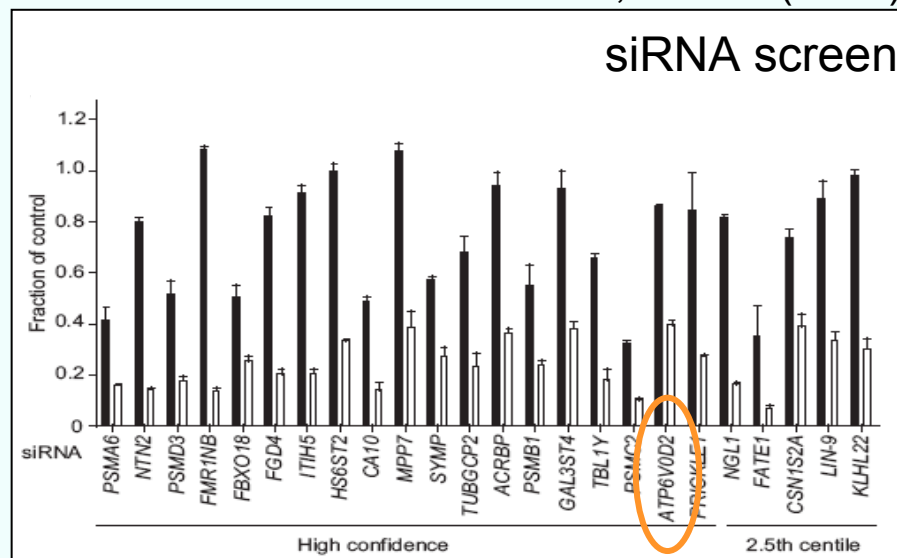
Mitotic Failure



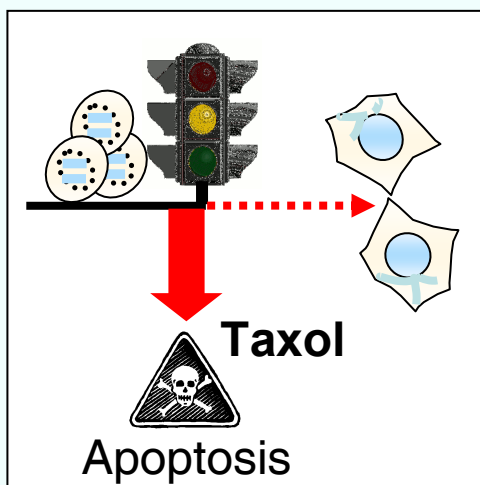
# Taxol Sensitization



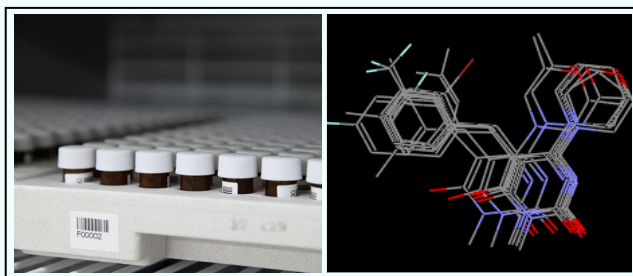
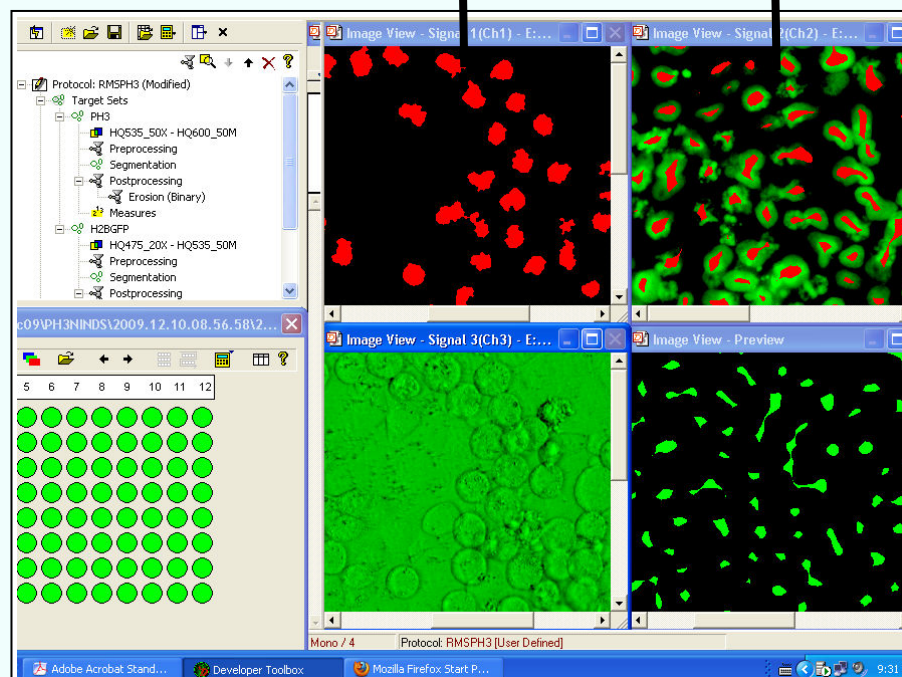
Whitehurst et al., Nature (2007)



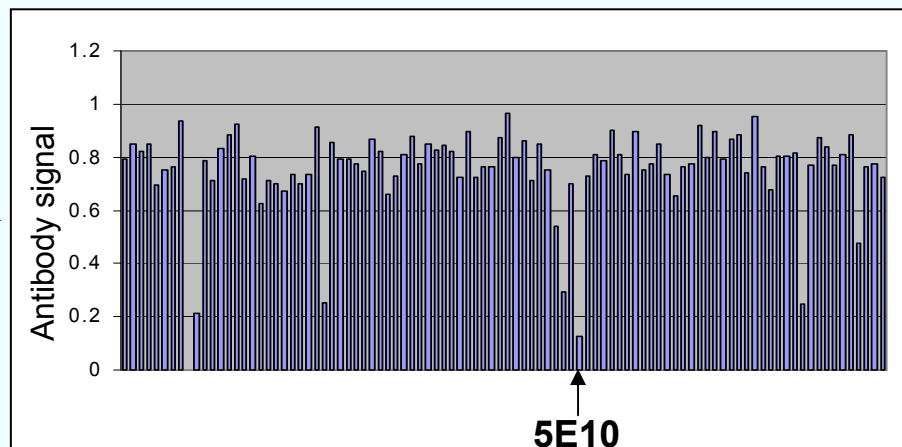
# Induction of Mitotic Death



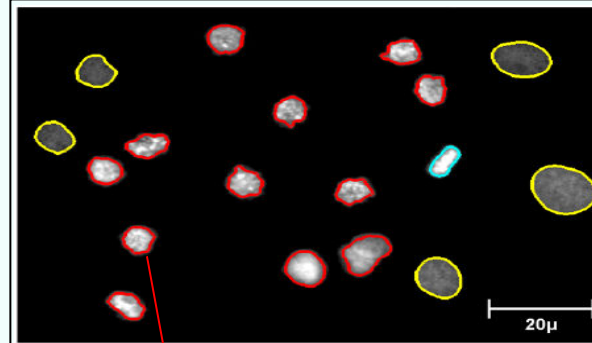
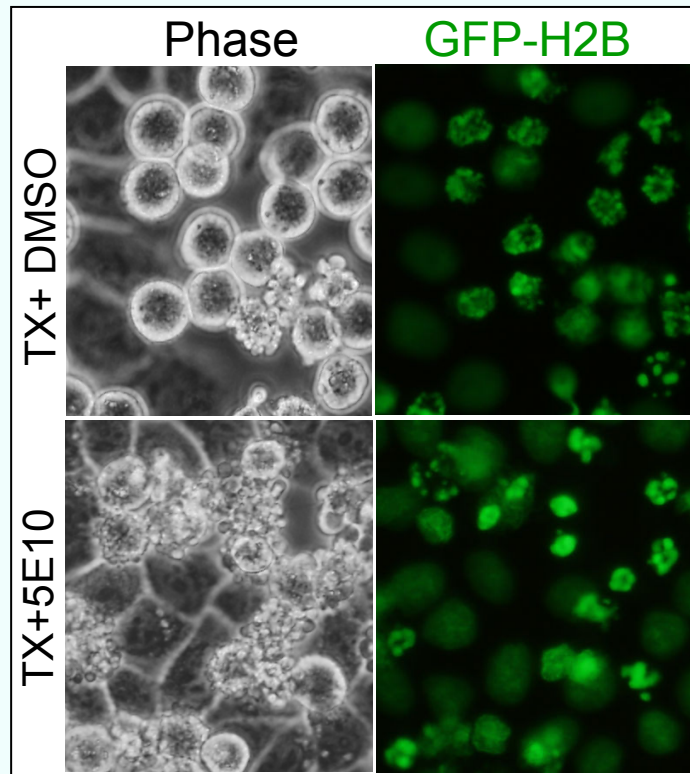
$\alpha$ PH3 segmentation      GFP-H2B segmentation



Pharmacologically Diverse



# HCS of Mitosis vs Apoptosis

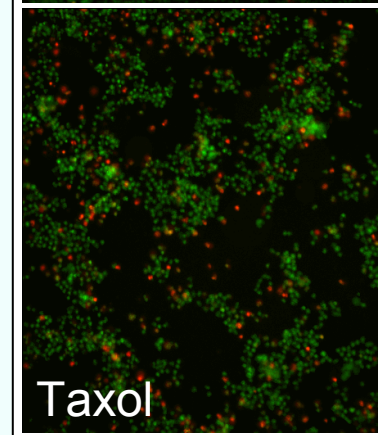
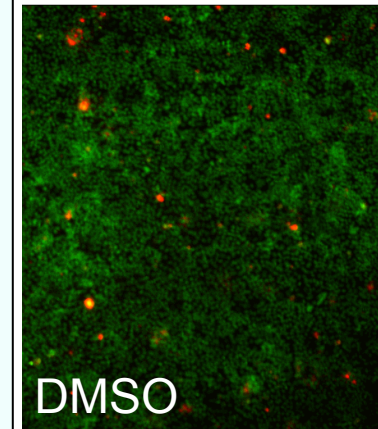


Tsui et al., PLoS (2009)

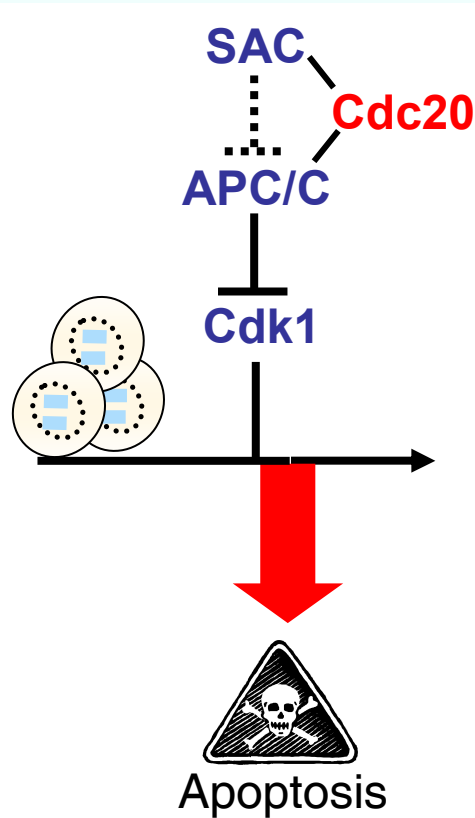
Monopolar spindles:

- Small form factor (Area/Perimeter)
- High SD of intensity

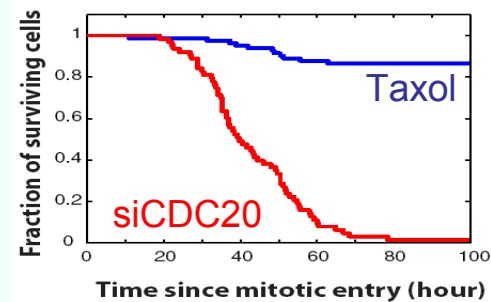
GFP-H2B  
Propidium



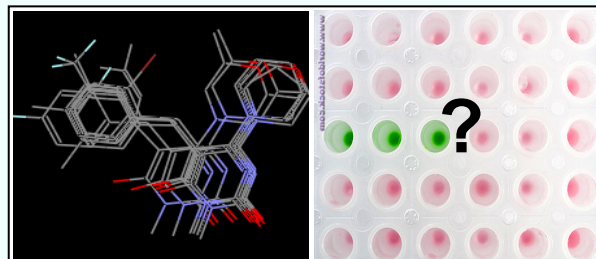
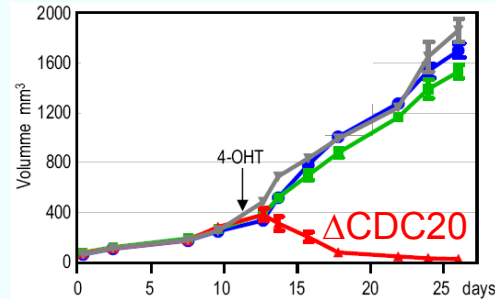
# Inhibition of Mitotic Exit



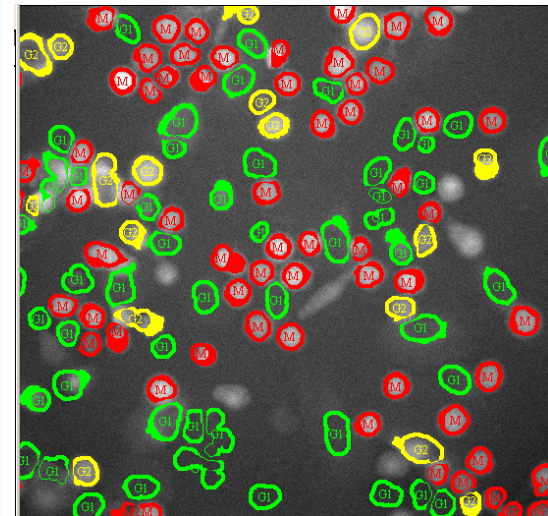
Huang et al., Cancer Cell (2009)  
MCF7



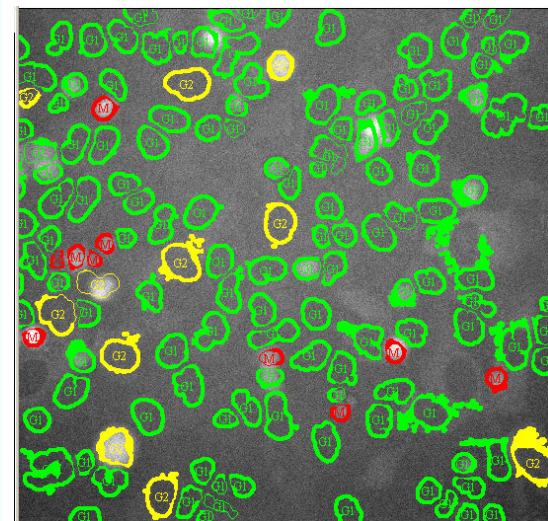
Manchado et al., Cancer Cell (2010)



Cyclin-B-GFP



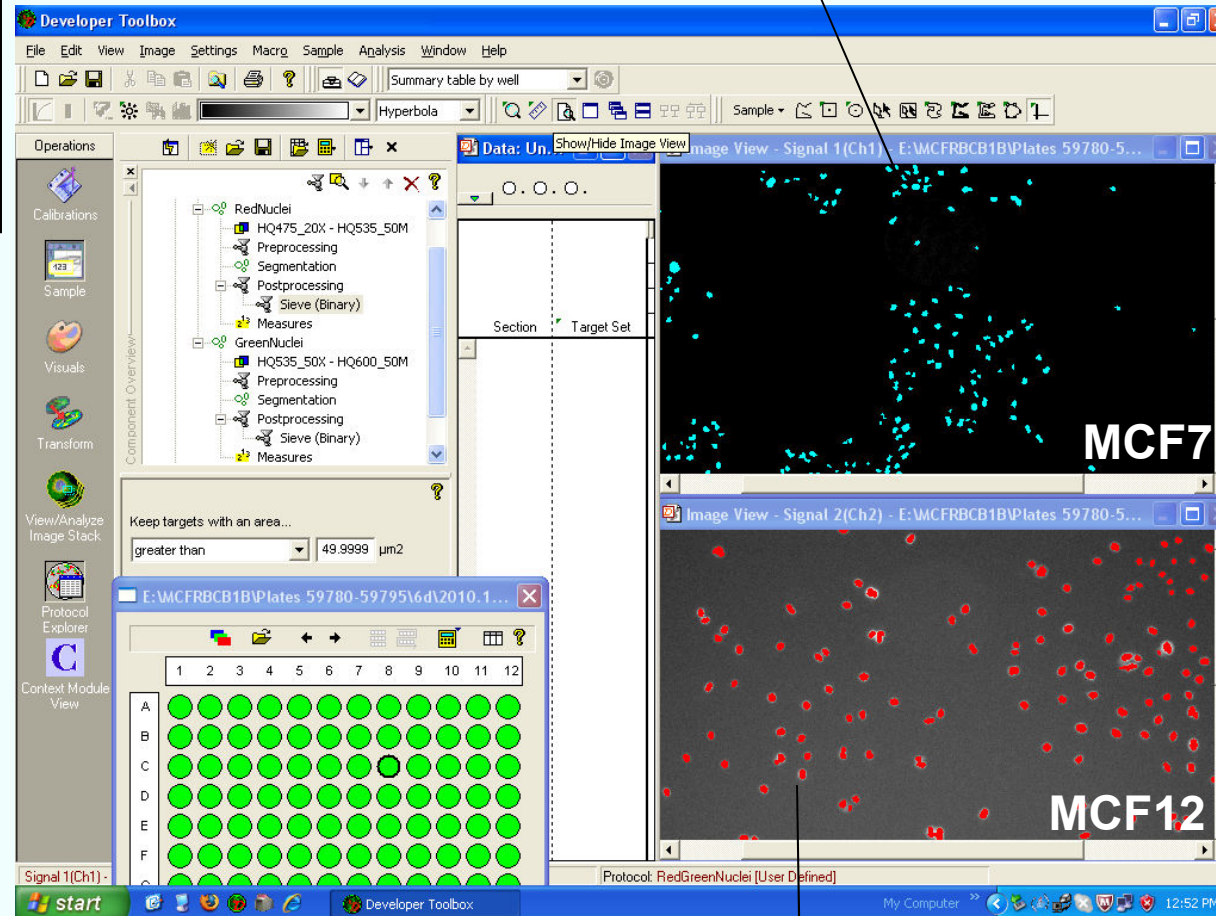
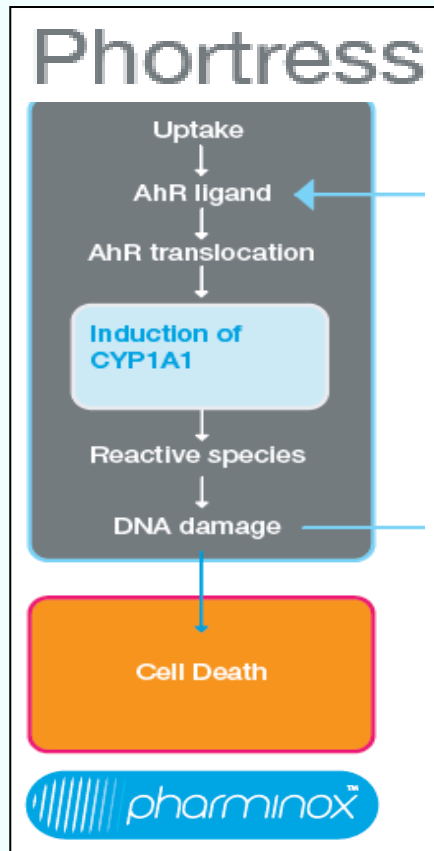
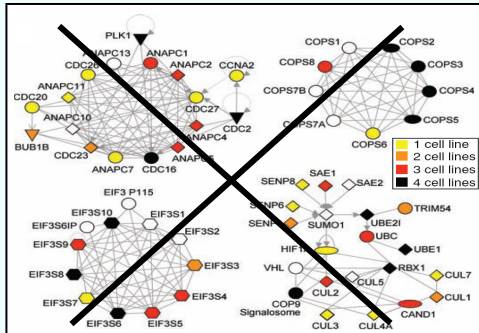
Nocodazole (35%M)



6h Nocodazole release (4%M)



# Selective Inhibition of Proliferation



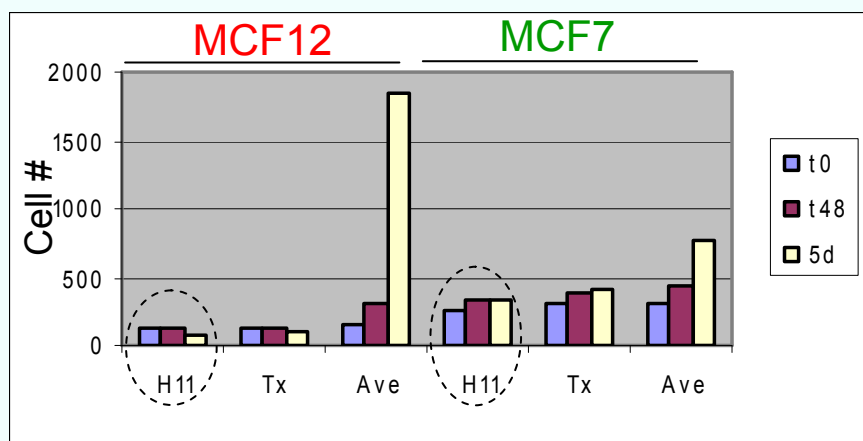
$\lambda 475\text{nm}$  segmentation

$\lambda 535\text{nm}$  segmentation

# Screening for Selective Growth Inhibition

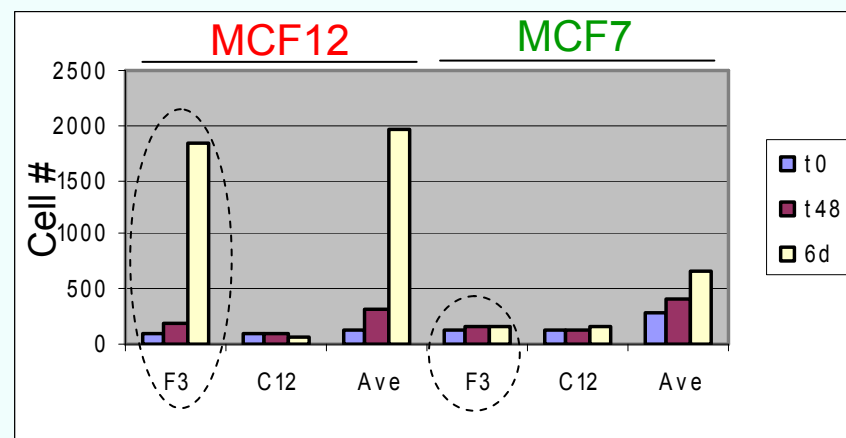
~50/10,000 compounds

## Non-selective inhibition

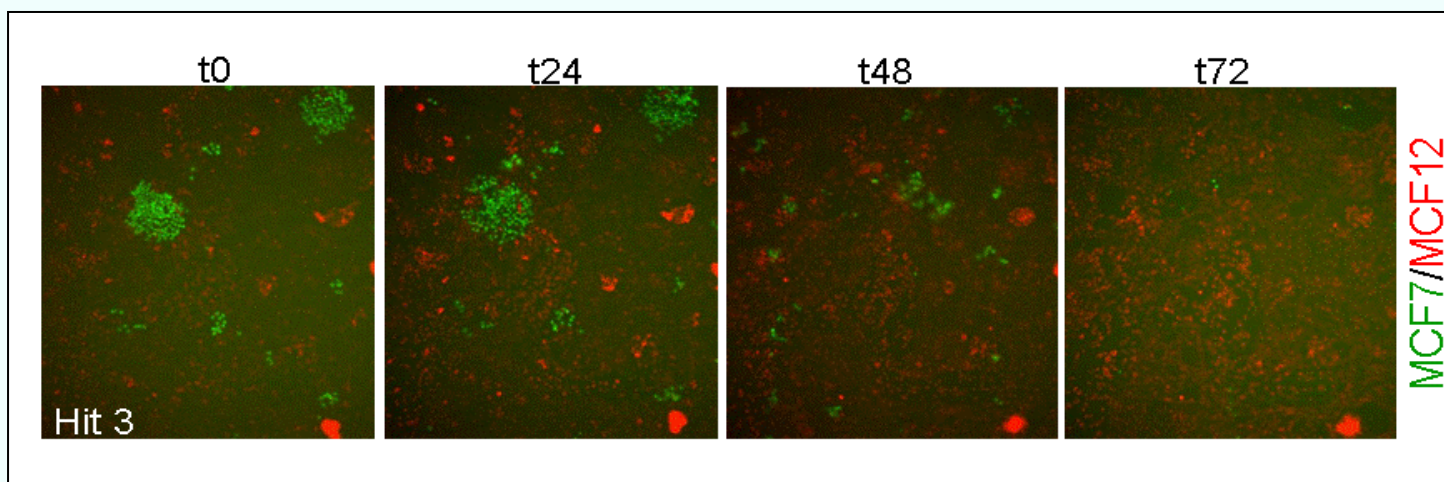


~2/10,000 compounds

## Selective inhibition



## Selective induction of cell death





Rachel Ramsdale



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AUSTRALIAN INSTITUTE OF MARINE SCIENCE



Catherine Liptrot

