



**EpiCypher**<sup>®</sup>

Bringing Epigenetics to Life



**Stratech**

EpiDyne<sup>®</sup>-FRET  
For Nucleosome  
Remodeling Assays

[www.stratech.co.uk/epicypher](http://www.stratech.co.uk/epicypher)

## Nucleosome Remodeling Assay by EpiDyne®-FRET

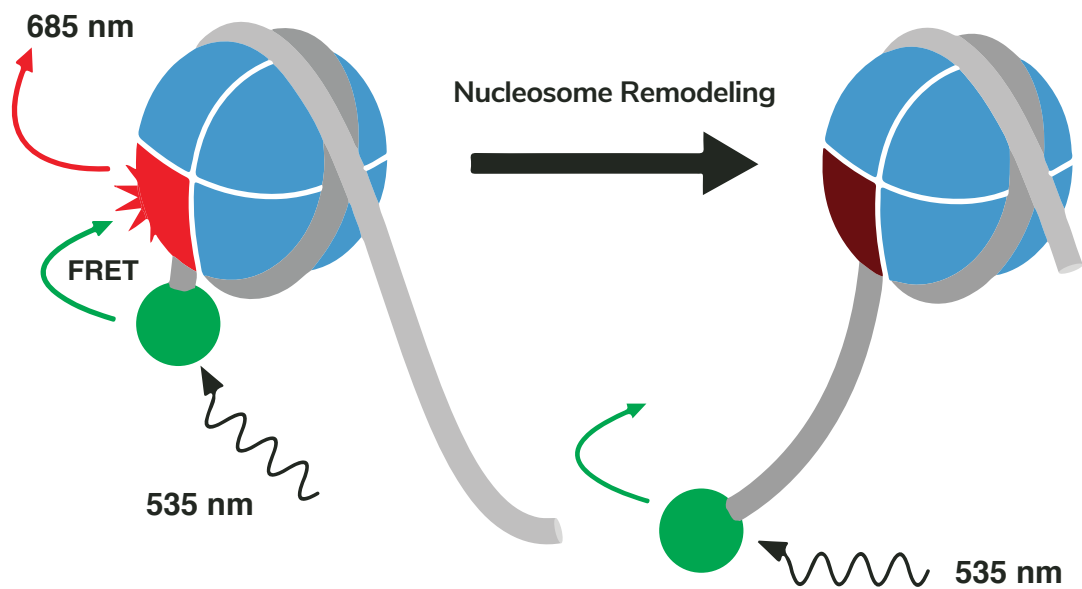
### EpiDyne®-FRET: a functionalized recombinant nucleosome-based assay platform for chromatin remodeling studies

Chromatin remodeling, or the repositioning of nucleosomes, regulates DNA access and thus gene expression and genome repair. Many ATP-dependent remodeling enzyme complexes are associated with human disease but are challenging study targets due to the requirement for nucleosome-based substrates. EpiCypher® has addressed this need by developing the EpiDyne® platform of fully recombinant remodeling substrates to monitor nucleosome repositioning along DNA using Fluorescence Resonance Energy Transfer (FRET) readout (Figure 1).

FIGURE 1

EpiDyne®-FRET Nucleosome Remodeling Substrates consist of a Cy5-labeled human histone octamer (H2A T120C-Cy5; shown as red section of octamer) wrapped by 5' Cy3-labeled DNA (217bp; green ball) comprising a terminal nucleosome positioning sequence (147bp Widom 601 adjacent to a TGGA-repeat region refractory to nucleosome assembly). In its assembled starting state, Cy3- Cy5 FRET is at a maximum. The activity of an ATP-dependent remodeler (e.g. RSC or another SWI/SNF ATPase) is detected by a reduction in FRET signal as the Cy3-labeled DNA 5' end is moved away from the Cy5-labeled octamer. EpiDyne®-FRET is a one-step no-wash method immediately compatible with HTS applications.

### EpiDyne®-FRET



### Useful for

- Inhibitor screening and development
- Structure-Activity Relationship assays
- Biochemical profiling of ATPase family proteins

## Nucleosome Remodeling Assay by EpiDyne®-FRET

### Chromatin Remodeling Enzymes As Therapeutic Targets

Aberrant nucleosome organization can severely disrupt gene expression, DNA repair and cellular differentiation, and it also plays a major role in human disorders, including cancer, inflammation, autoimmunity, schizophrenia, cardiovascular disease, and intellectual disability. Remarkably, nearly 20% of all cancers contain mutations in subunits from the SWI/SNF family of ATP-dependent chromatin remodeling complexes. These enzyme complexes regulate local genome access by ‘pumping’ the DNA around histone octamers, thus ‘sliding’ nucleosomes.

### SWI/SNF Remodeling Complex

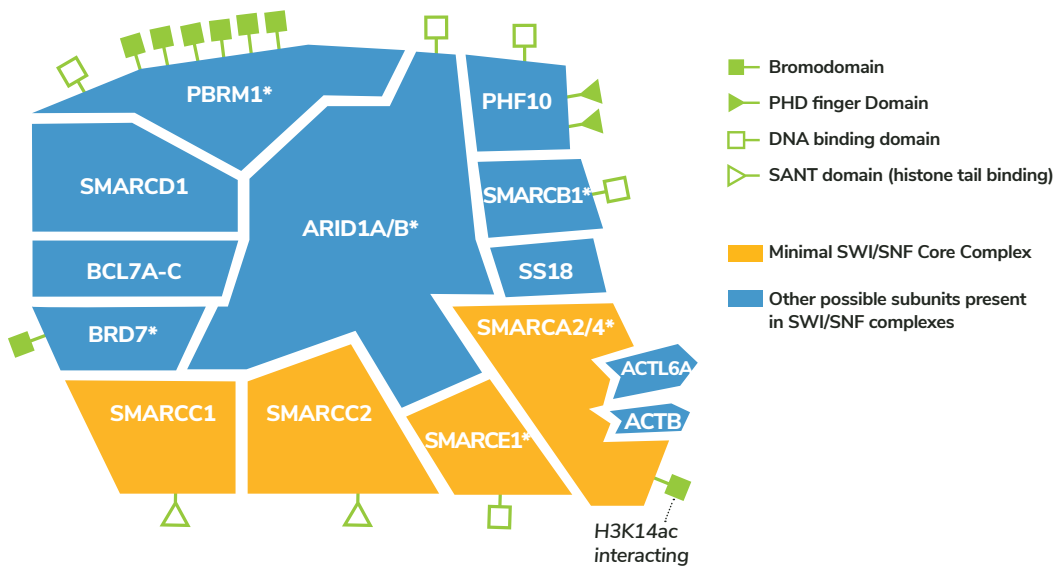


FIGURE 2

Schematic of SWI/SNF remodeling complex. Yellow subunits denote minimal core complex that recapitulates activity of full complex.

Recurrent somatic mutations in SWI/SNF subunits are observed in multiple cancers, supporting a driver role in tumorigenesis. The mutated remodeling proteins are attractive therapeutic targets, since further compromising their ATPase activity promotes cancer cell death but spares normal cells.

SUBUNIT	CANCER
ARID1A	Ovarian, Hepatocellular, Bladder, Gastric, Endometrioid, Pancreatic, Colon, Lung, Neuroblastoma, Burkitt Lymphoma
ARID1B	Melanoma, Neuroblastoma, Hepatocellular, Pancreatic, Liver
PBRM1	Renal cell carcinoma, Breast, Gastric, Pancreatic
ARID2	Melanoma, Hepatocellular, Pancreatic
SMARCA2	Lung, Colon, Breast
SMARCA4	Lung, Medulloblastoma, Burkitt Lymphoma, SCCOHT
SMARCB1	Rhabdoid tumor, Familial Schwannomatosis
SMARCE1	Spinal meningitis
BRD7	Breast

TABLE 1

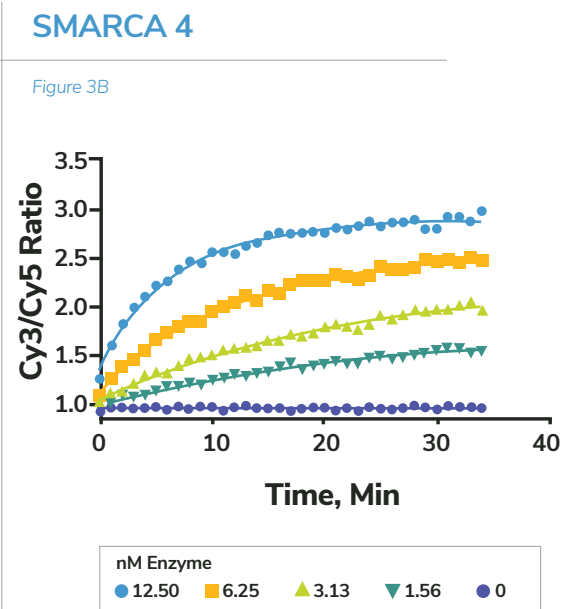
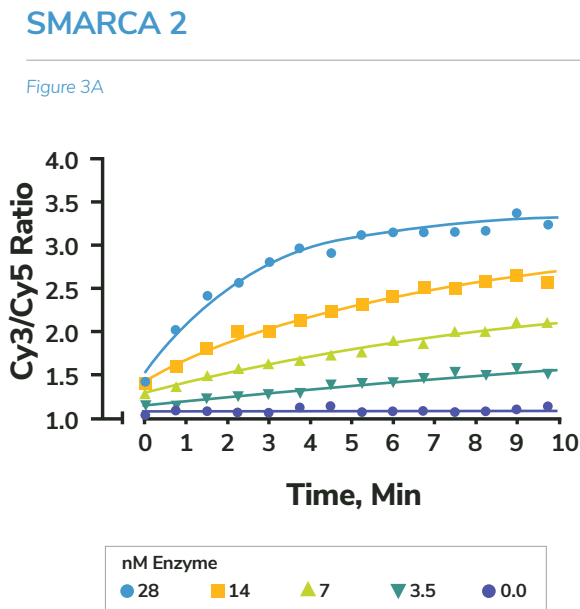
List of cancers associated with various SWI/SNF subunit mutations adapted from Helming et al. Cancer Cell 26, 309-317 (2014). Asterisks in Figure 2 indicate associated cancers in Table 1.

## Nucleosome Remodeling Assay by EpiDyne®-FRET

# EpiDyne®-FRET allows unprecedented access to disease-relevant ATP-dependent chromatin remodeling complexes

FIGURE 3

EpiDyne®-FRET nucleosomes (20 nM) were incubated with chromatin remodeling enzyme (Panel 3A, SMARCA2; panel 3B, SMARCA4) at the indicated concentration in the presence of ATP (2 mM). Upon ATP addition, reactions were immediately read in an Envision Multi-label plate reader. Data are presented as the mean of the Cy3-Cy5 ratio (N=2).



## ORDERING INFO

### Chromatin Remodeling Substrate, Fluorescent Readout

#### EpiDyne®-FRET Nucleosome Remodeling Assay Substrate

Catalog No. 16-4201  
Pack Size: 50 µg

### Chromatin Remodeling Substrates, Non-Fluorescent Readout

#### ST601-GATC1

Cat. No. 16-4101  
Pack Size: 50 µg

#### ST601-GATC1, 50-N-66, Biotinylated

Cat. No.: 16-4114  
Pack Size: 50 µg

#### ST601-GATC1, Biotinylated

Cat. No.: 16-4111  
Pack Size: 50 µg

#### ST601-GATC1,2, 50-N-66, Biotinylated

Cat. No.: 16-4115  
Pack Size: 50 µg

#### ST601-GATC1,2, Biotinylated

Cat. No.: 16-4112  
Pack Size: 50 µg

#### ST601-GATC1,2,3, 50-N-66, Biotinylated

Cat. No.: 16-4116  
Pack Size: 50 µg

#### ST601-GATC1,2,3, Biotinylated

Cat. No.: 16-4113  
Pack Size: 50 µg

Website: [epicypher.com/epidyne](http://epicypher.com/epidyne)

### Enzymes

#### SMARCA2 Chromatin Remodeling Enzyme (Human BRM)

Catalog No. 15-1015  
Pack Size: 100 remodeling rxns

#### SMARCA4 Chromatin Remodeling Enzyme (Human BRG1)

Catalog No. 15-1014  
Pack Size: 100 remodeling rxns

#### ACF Chromatin Remodeling Enzyme Complex

Catalog No. 15-1013  
Pack Size: 100 remodeling rxns

Website: [epicypher.com/epidyne-enzymes](http://epicypher.com/epidyne-enzymes)

EpiDyne® Chromatin Remodeling HTS Assay Services are available

Inquire at: [info@stratech.co.uk](mailto:info@stratech.co.uk)



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