RELATED INFORMATION DRUGS & BIOLOGICS EXPERIMENTAL PHARMACOLOGY EXPERIMENTAL MODELS

LATED INFORMATION

Heightman, T.D.; Berdini, V.; Braithwaite, H.; et al. Summary — Pill Test Fragment-based discovery of a highly potent, orally bioavailable ERK1/2 inhibitor that modulates the

activity of ERK1/2 29th EORTC-NCI-AACR Symp Mol Targets Cancer Ther (October 26-30, Philadelphia) 2017, Abst B161

Flemington, V.; Simpson, I.; Davies, E.; et al. Discovery and characterization of AZ6197, a potent and selective ERK1/2 inhibitor 29th EORTC-NCI-AACR Symp Mol Targets Cancer Ther (October 26-30, Philadelphia) 2017, Abst B156

ACOLOGY EXPERIMENTAL MODELS

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Search for specific information from a conference:



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Summary OpenURL

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Display References with associated text

Information via Ouick Storch

Printer Friendly Format

This search will retrieve all the citations in Integrity from the 2017 edition of the chosen conference.

If you are interested in only the citations associated with products with a certain mechanism of action, this can be included in the search using the **Product** section. This again retrieves a results list with the relevant citations.



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catalytic activity and phosphorylation were sustained for up to 24 hours.

	Reference
	29th EORTC-NCI-AACR Symp Mol Targets Cancer Ther (October 26-30, Philadelphia) 2017, Abst
	B154
	Title
l	Characterization of a novel ERK1/2 inhibitor, which modulates the phosphorylation and catalytic
ł.	activity of FRK1/2

(980502) Synthesis and optimization led to the discovery of a novel series of pERK modulating ERK1/2 inhibitors. The lead compound, compound [1], inhibited ERK1/2 catalytic activity with an IC50 of 3 nM when evaluated in an ERK TRF kinase assay. The compound bound to the active site of ERK2, and in a screen of 465 kinases it proved to be highly selective for ERK1/2. In A375 (BRAF-mutant melanoma) and HCT116 (KRAS-mutant colorectal cancer) cell lines, [1] potently inhibited ERK activity with IC values of 7.2 and 5.2 nM, respectively. Additionally, the compound inhibited the proliferation of cell lines harboring a range of MAPK aberrations. In vivo, [1] was evaluated in A375 (BRAF-mutant melanoma) and Calu-6 (KRAS-mutant lung) tumor xenografts, demonstrating that once-daily oral administration of 50 mg/kg of this compound resulted in significant antitumor activity in both models. A single dose at 50 mg/kg inhibited ERK catalytic activity in KRAS-mutant lung tumor xenografts and conferred a decrease in the phosphorylation of ERK itself. These effects on ERK

Records Retrieved 7 in D			n Drugs & Biolo	Drugs & Biologics Search Results				Options			
Drugs & E	Drugs & Biologics Search Results										
<u>Entry</u> Number	<u>Highest</u> Phase	Code Name	Generic Name	Brand Name	Product Category	Therapeutic Group	Mechanism of Action	Organizati			
355417*	Phase III	ARRY-142886 hydrogen sulfate ARRY-886 hydrogen sulfate AZD-6244 Hyd-Sulfate AZD-6244 hydrogen sulfate MK-5618 NSC- 748727	Selumetinib sulfate (Prop INN; USAN)			Multiple Myeloma Therapy Sarcoma Therapy Non-Small Cell Lung Cancer Therapy Endocrine Cancer Therapy Neurological Genetic Disorders, Treatment of Neurologic Cancer Therapy Melanoma Therapy Gastric Cancer Therapy Dvarian Cancer Therapy	Dual Specificity Mitogen- Activated Protein Kinase Kinase 2 (MAP2K2; MEK2) Inhibitors Signal Transduction Modulators Extracellular- Regulated	National Cancer Institute Array BioPharma (Originator) AstraZeneci University o Chicago Samsung Medical Center Merck & Co			

Advanced Search		Session History	Clear Form	Start
Reference				
Year	From201	7		Index AND
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Set up alerts for forthcoming conferences:

Advanced Search		Session History	Clear Form	Start
leference				
Source (Congress)	• "EORTC-	-NCI-AACR Symp Mol Targets Cancer Ther"		Index AND
Year	From201	18		Index AND
Optional Value				Index AND



Tip:

 Checkboxes can be used to select specific records which can then be further manipulated using Options such as Export to Excel or Integrity Reports that Integrity makes available for easy data management.

Where available, an *Integrity* Summary button will appear next to the reference citation. *Integrity* inhouse specialists have analyzed the information and published a summary of the findings. Click the button to access this information in a separate pop-up window.

To see information related to the citations that can be found in the different Knowledge Areas of *Integrity* go to **All Related Information via Quick Search**, which is one of the data management **Options**. Link to **Drugs & Biologics** to view the drug records associated with these references.

You can explore *Integrity* reference citations to see more literature associated with your area of interest.

For example, in the **Literature Knowledge Area** use the **Product** section to search for references associated with products that have a mechanism of interest. You can specify to see all the citations that have been published this year.

You can set up a **Save Query** alert to be notified when new citations from the latest edition of a particular conference are included in *Integrity*.

Set up a search for **Source (Congress)** [conference/congress of interest] and **Year** = From 2018. Your search will not retrieve results if the conference is in the future.

You will be given a Save Query option.



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Save Query				
To save your query, enter a Query Name and Description and click Save Query.				
Use the Optional e-mail alert field to change the frequency of alerts: never, daily, weekly or monthly.				
When new records are found by the query, you will receive notification by e-mail.				
Query Name	EORTC			
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Fill in the appropriate fields, select an alert frequency and save the alert. You will now be notified when new records that meet your search criteria enter *Integrity*.

Tip: For information about forthcoming conferences to be covered in Integrity, refer to the left-hand side of the advanced search form in the *Literature Knowledge Area*. The tabs contain information about recent *conferences* (left tab) and *forthcoming conferences* (right tab) covered by Integrity.

