

## INFORME DE BÚSQUEDA DE PATENTES

**N° de solicitud:** NC2017/0011482 **Prioridad:** (22 de abril de 2015)

**CRITERIOS PARA LA BÚSQUEDA** (Anexar pantallazo o reporte en PDF, junto con los boléanos utilizados\*. Para el caso de secuencias, anexar alineamientos)

|                                                    |                                                                                                                                                                                                       |
|----------------------------------------------------|-------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| <b>PALABRAS CLAVE UTILIZADAS</b>                   | 1. "4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one" and "crystalline Form"<br>2. "Bromodomain inhibitor" AND XRPD before:publication:20150421 after:publication:19900101 |
| <b>CLASIFICACIONES UTILIZADAS PARA LA BÚSQUEDA</b> | <b>CIP:</b> A61K 31/47, C07D 217/24.<br><b>CPC</b> (Opcional):                                                                                                                                        |

### DOCUMENTOS CONSIDERADOS RELEVANTES (documentos patentes, literatura no patente, etc...)

| Informes/Base de datos | N° de solicitud/<br>Autor** | Fecha de<br>Publicación/Presentación<br>(DD/MM/AAAA) | Reiv. afectadas (Con indicación, si procede, de las partes relevantes o números de secuencias) | Categorías especiales de los documentos citados*** |
|------------------------|-----------------------------|------------------------------------------------------|------------------------------------------------------------------------------------------------|----------------------------------------------------|
| ISR/WO                 | US2012208814                | 16/Agosto/2012                                       |                                                                                                | A                                                  |
|                        | US2007213323                | 13/Septiembre/2007                                   |                                                                                                | A                                                  |
|                        | US2015111885                | 23/Abril/2015                                        |                                                                                                | P                                                  |
| ISR/EP                 | WO2015058160                | 23/Abril/2015                                        |                                                                                                | A                                                  |
|                        | WO2013097601                | 04/Julio/2013                                        |                                                                                                | A                                                  |
|                        | WO2018075796                | 26/Abril/2018                                        |                                                                                                | E                                                  |
| Google/patents         | US20160310423               | 27/Octubre/2016                                      |                                                                                                | P                                                  |
| SIPI                   | 12077201                    | 10/Mayo/2012                                         |                                                                                                | A                                                  |
|                        | 15245439                    | 15/Octubre/2015                                      |                                                                                                | A                                                  |
|                        | 15245450                    | 15/Octubre/2015                                      |                                                                                                | A                                                  |
|                        | 14011125                    | 21/Enero/2014                                        |                                                                                                | A                                                  |
|                        | 14160451                    | 24/Julio/2014                                        |                                                                                                | A                                                  |

**"A"** documento que define el estado general de la técnica no considerado como particularmente relevante.

**"E"** solicitud de patente o patente anterior pero publicada en la fecha de presentación internacional o en fecha posterior.

**"P"** documento publicado antes de la fecha de presentación internacional, pero con posterioridad a la fecha de prioridad reivindicada.

**"X"** documento particularmente relevante; la invención reivindicada no puede considerarse nueva o que implique una actividad inventiva por referencia al documento aisladamente considerado.

**"Y"** documento particularmente relevante; la invención reivindicada no puede considerarse que implique una actividad inventiva cuando el documento se asocia a otro u otros documentos de la misma naturaleza, cuya combinación resulta evidente para un experto en la materia.

\*\*\*\* Indicar entre paréntesis la Autoridad que realizó la búsqueda o el informe preliminar Ejs: EPO, JP, INAPI.

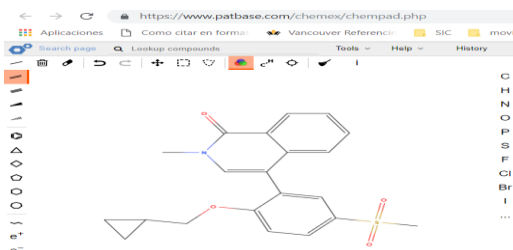
#### Literatura no patente

\*\*Se incluye entre otros, toda la referencia de la literatura no patente.

**Fecha del reporte de búsqueda:**

04/11/2018

### PATBASE



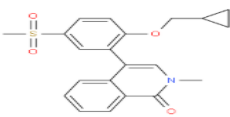
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| #  | Structure                                                                         | Details                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     |
|----|-----------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| 1) |  | <b>4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one</b><br>IUPAC Name: 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one<br>Molecular Weight: 383.461 g/mol<br>InChIKey: UWZAJPTKGMFJ-UHFFFAOYSA-N<br>SMILES: C1(CC1)COC1=C(C=C(C=C1)S(=O)(=O)C)C1=CN(C(C2=CC=CC=C2)=O)C<br>Formula: C <sub>21</sub> H <sub>21</sub> NO <sub>4</sub> S<br><br>26 patent publications found   View in PatBase   Stats   Analytics<br>Links: PubChem   ChemSpider   Wikipedia   Save MOL |

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| 24 | crystalline forms                                                                                                        | 2403    | View   Browse   Hits   Optimise   more... |
| 23 | BROMODOMAIN INHIBITORS                                                                                                   | 115     | View   Browse   Hits   Optimise   more... |
| 22 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methyl-8-(3-methylpyridin-3-yl)isoquinolin-1-one | 1       | View   Browse   Hits   Optimise   more... |
| 21 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-ethylsulfonylphenyl]-7-fluoro-2-methylisoquinolin-1-one                  | 1       | View   Browse   Hits   Optimise   more... |
| 20 | [Chemical Explorer] 8-chloro-4-[2-(cyclopropylmethoxy)-5-ethylsulfonylphenyl]-2-methylisoquinolin-1-one                  | 1       | View   Browse   Hits   Optimise   more... |
| 19 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-ethylsulfonylphenyl]-2-methyl-8-(1-methylpyrazol-4-yl)isoquinolin-1-one  | 1       | View   Browse   Hits   Optimise   more... |
| 18 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methyl-8-(trifluoromethyl)isoquinolin-1-one      | 1       | View   Browse   Hits   Optimise   more... |
| 17 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methyl-8-(4-methylpyrazol-1-yl)isoquinolin-1-one | 1       | View   Browse   Hits   Optimise   more... |
| 16 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methyl-8-(trifluoromethoxy)isoquinolin-1-one     | 1       | View   Browse   Hits   Optimise   more... |
| 15 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-7-fluoro-2-methylisoquinolin-1-one                 | 1       | View   Browse   Hits   Optimise   more... |
| 14 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-propan-2-ylsulfonylphenyl]-2-methylisoquinolin-1-one                     | 1       | View   Browse   Hits   Optimise   more... |
| 13 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-propan-2-ylsulfonylphenyl]-8-methoxy-2-methylisoquinolin-1-one           | 1       | View   Browse   Hits   Optimise   more... |
| 12 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-8-fluoro-2-methylisoquinolin-1-one                 | 1       | View   Browse   Hits   Optimise   more... |
| 11 | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one                          | 3       | View   Browse   Hits   Optimise   more... |
| 10 | [Chemical Explorer] 4-[2-(cyclopropyl-8-methylsulfonyl-3,4-dihydro-2H-chromen-8-yl)-2-methylisoquinolin-1-one            | 1       | View   Browse   Hits   Optimise   more... |
| 9  | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-ethylsulfonylphenyl]-2-methylisoquinolin-1-one                           | 1       | View   Browse   Hits   Optimise   more... |
| 8  | [Chemical Explorer] 4-[2-(cyclopropyl-5-methylsulfonyl-2,3-dihydro-1-benzofuran-7-yl)-2-methylisoquinolin-1-one          | 1       | View   Browse   Hits   Optimise   more... |
| 7  | [Chemical Explorer] 2-methyl-4-(5-methylsulfonyl-2-propoxyphenyl)isoquinolin-1-one                                       | 1       | View   Browse   Hits   Optimise   more... |
| 6  | [Chemical Explorer] 4-[2-(cyclobutylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one                           | 1       | View   Browse   Hits   Optimise   more... |
| 5  | [Chemical Explorer] 2-methyl-4-(5-methylsulfonyl-2-oxetan-3-ylmethoxyphenyl)isoquinolin-1-one                            | 1       | View   Browse   Hits   Optimise   more... |
| 4  | [Chemical Explorer] 2-methyl-4-[2-(1-methylcyclopropylmethoxy)-5-methylsulfonylphenyl]isoquinolin-1-one                  | 1       | View   Browse   Hits   Optimise   more... |
| 3  | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-(methylsulfonylphenyl)-8-methoxy-2-methylisoquinolin-1(2H)-one           | 1       | View   Browse   Hits   Optimise   more... |
| 2  | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one                          | 3       | View   Browse   Hits   Optimise   more... |
| 1  | [Chemical Explorer] 4-[2-(cyclopropylmethoxy)-5-methylsulfonylphenyl]-2-methylisoquinolin-1-one                          | 3       | View   Browse   Hits   Optimise   more... |

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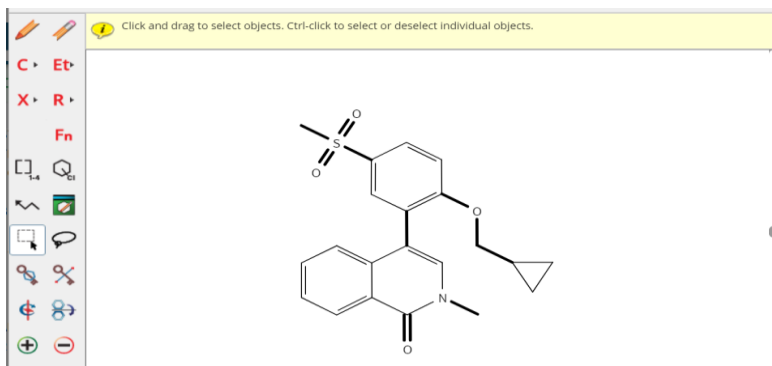
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## ORBIT

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| #  | Title                                                                                                                                                          | Publication number | 1st app. date | Applicant/Assignee                                                     | Relevance score | Litigations & oppositions |
|----|----------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------------|---------------|------------------------------------------------------------------------|-----------------|---------------------------|
| 1. | 4-(8-methoxy-1-((1-methoxypropan-2-yl)-2-(tetrahydro-2h-pyran-4-yl)-1h-imidazo[4,5-c]quinolin-7-yl)-3,5-dimethylisoxazole and its use as bromodomain inhibitor | GB201114103        | 2011-08-17    | GLAXOSMITHKLINE*; GLAXOSMITHKLINE A LIABILITY; ZOE SMITH KLEIN GALLERY | 100 %           |                           |
| 2. | Bromodomain inhibitor                                                                                                                                          | WO2018075796       | 2017-10-19    | CELGENE QUANTICEL RESEARCH*                                            | 98 %            |                           |

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| Application number | Title                                                                                                                                                         | Applicant(s) | Inventor(s)                                                                              | Filing date | Application status |
|--------------------|---------------------------------------------------------------------------------------------------------------------------------------------------------------|--------------|------------------------------------------------------------------------------------------|-------------|--------------------|
| 2013351146         | Method of preparation of Crystal Forms of 4-(cyclopropylmethoxy)-N-(3,5-dichloro-1-oxido-4-pyridyl)-5-methoxypyridine-2-carboxamide and Crystal Forms thereof | Sanofi       | Billot, Pascal; Brige, Didier; Giuliani, Alexandre; Elmaleh, Hagit; Perrin, Marc-Antoine | 2013-11-28  | LAPSED             |
| 2013351145         | Process for preparing 4-(cyclopropylmethoxy)-N-(3,5-dichloro-1-oxido-4-pyridyl)-5-methoxypyridine-2-carboxamide                                               | Sanofi       | Casimir, Jean-Paul; Rossey, Guy; Wehrey, Christian                                       | 2013-11-28  | LAPSED             |
| 2000079218         | Polymorphic b form of 3-(cyclopropylmethoxy)-4-(4-(methylsulfonyl)phenyl)-5,5-dimethyl-5h-furan-2-one                                                         | Meril        | Calais, Beatrice; Chassagneux, Evelyne; Bonard, Jean-Michel                              | 2000-10-09  | GRANTED            |

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|   | Application number         | Title                                                                                                                                                           | Applicant(s)                                                                                  | Inventor(s)                                                                                                                  | Filing date | Application status | Results per page 10 ▼ |                          |
|---|----------------------------|-----------------------------------------------------------------------------------------------------------------------------------------------------------------|-----------------------------------------------------------------------------------------------|------------------------------------------------------------------------------------------------------------------------------|-------------|--------------------|-----------------------|--------------------------|
| 1 | <a href="#">2016379347</a> | Bromodomain and extra-terminal protein inhibitor combination therapy                                                                                            | Celgene Quantical Research, Inc.                                                              | Nikolova, Zariana; Cho, Robert; Stafford, Jeffrey Alan                                                                       | 2016-12-20  | FILED              |                       | <input type="checkbox"/> |
| 2 | <a href="#">2016252992</a> | Bromodomain inhibitor                                                                                                                                           | Celgene Quantical Research, Inc.                                                              | Betancort, Juan Manuel; Stafford, Jeffrey Alan; Stansfield, Ryan; Veal, James Marvin                                         | 2016-04-22  | FILED              |                       | <input type="checkbox"/> |
| 3 | <a href="#">2016249273</a> | Bromodomain inhibitor                                                                                                                                           | Celgene Quantical Research, Inc.                                                              | Bennett, Michael John; Betancort, Juan Manuel; Boloor, Amogh; Kaldor, Stephen W.; Stafford, Jeffrey Alan; Veal, James Marvin | 2016-04-15  | FILED              |                       | <input type="checkbox"/> |
| 4 | <a href="#">2015358334</a> | Bromodomain inhibitor as adjuvant in cancer immunotherapy                                                                                                       | Dana-Farber Cancer Institute, Inc.; H. Lee Moffitt Cancer Center and Research Institute, Inc. | Sotomayor, Eduardo M.; Bradner, James; Tao, Jianguo                                                                          | 2015-12-04  | FILED              |                       | <input type="checkbox"/> |
| 5 | <a href="#">2015203454</a> | BENZODIAZEPINE BROMODOMAIN INHIBITOR                                                                                                                            | GlaxoSmithKline LLC                                                                           | Bailey, James; Gosmini, Romain Luc Marie; Mirguet, Olivier; Witherington, Jason                                              | 2015-06-23  | GRANTED            |                       | <input type="checkbox"/> |
| 6 | <a href="#">2012296914</a> | 4-(8-methoxy-1-((1-methoxypropan-2-yl)-2-(tetrahydro-2H-pyran-4-yl)-1 H-imidazo(4,5-c)quinolin-7-yl)-3,5-dimethylisoxazole and its use as bromodomain inhibitor | GlaxoSmithKline LLC                                                                           | Demont, Emmanuel Hubert; Jones, Katherine Louise; Watson, Robert J.                                                          | 2012-08-15  | GRANTED            |                       | <input type="checkbox"/> |
| 7 | <a href="#">2010317096</a> | Benzodiazepine bromodomain inhibitor                                                                                                                            | GlaxoSmithKline LLC                                                                           | Bailey, James; Gosmini, Romain Luc Marie; Mirguet, Olivier; Witherington, Jason                                              | 2010-11-03  | GRANTED            |                       | <input type="checkbox"/> |
| 8 | <a href="#">2010314395</a> | Benzodiazepine bromodomain inhibitor                                                                                                                            | GlaxoSmithKline LLC                                                                           | Gosmini, Romain Luc Marie; Mirguet, Olivier                                                                                  | 2010-08-06  | GRANTED            |                       | <input type="checkbox"/> |

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| #  | Patent Number | Patent Title                                                                                     | Score |
|----|---------------|--------------------------------------------------------------------------------------------------|-------|
| 1. | 2983446       | BROMODOMAIN INHIBITOR                                                                            | 64%   |
| 2. | 2902225       | METHODS OF TREATMENT OF HUMAN CYTOMEGALOVIRUS INFECTION AND DISEASES WITH BROMODOMAIN INHIBITORS | 50%   |

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| 1      |           | 168       | BROMODOMAIN 1W INHIBITOR |
| 2      |           | 7.980     | CRYSTALLINE 1W FORMS     |
| 3      |           | 11        | 1 AND 2                  |
| 4      |           | 1.108.015 | CRYST+                   |
| 5      |           | 13        | 1 AND 4                  |
| 6      |           | 34.848    | POLYMO+                  |
| 7      |           | 1         | 1 AND 6                  |
| 8      |           | 852       | ISOQUINOLIN              |
| 9      |           | 0         | 1 AND 8                  |
| 10     |           | 327       | ISOQUINOLIN 3W ONE       |
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4 0 (2 Facets) (1 AND 2)