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(54) Title **BILE ACID DERIVATIVES AS FXR/TGR5 AGONISTS AND METHODS OF USE THEREOF**

(56) References Cited: WO-A1-02/28881  
WO-A1-2014/184271  
WO-A2-2014/036379  
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US-A1- 2014 206 657

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PELLICCIARI R ET AL: "Back Door Modulation of the Farnesoid X Receptor: Design, Synthesis and Biological Evaluation of a Series of Side Chain Modified Chenodeoxycholic Acid Derivatives", JOURNAL OF MEDICINAL CHEMISTRY, AMERICAN CHEMICAL SOCIETY, vol. 49, 15 June 2006 (2006-06-15), pages 4208-4215, XP008092111, ISSN: 0022-2623, DOI: 10.1021/JM060294K

DATABASE CA [Online] CHEMICAL ABSTRACTS SERVICE, COLUMBUS, OHIO, US; AOGO, TOSHIKI ET AL: "Positive-working photosensitive composition", XP002784471, retrieved from STN Database accession no. 1999:260860 & JP H11 109628 A (FUJI PHOTO FILM CO LTD) 23 April 1999 (1999-04-23)

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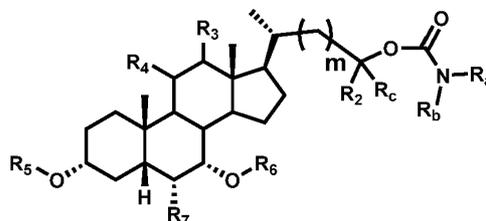
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## 5 Patentkrav

1. En forbindelse representert ved formel I, eller et farmasøytisk akseptabelt salt derav:

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(I)

hvor:

R<sub>a</sub> er valgt fra gruppen som består av:

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- 1) Hydrogen;
- 2) Substituert eller usubstituert -C<sub>1</sub>-C<sub>6</sub> alkoksy;
- 3) Substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl;
- 4) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl;
- 5) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl;
- 25 6) Substituert eller usubstituert -C<sub>3</sub>-C<sub>6</sub> -cykloalkyl;
- 7) Substituert eller usubstituert arylalkyl;
- 8) Substituert eller usubstituert aryl; og
- 9) heteroaryl;

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R<sub>b</sub> er valgt fra gruppen som består av:

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- 1) -SO<sub>2</sub>R<sub>1</sub>;
- 2) -C(O)R<sub>1</sub>;
- 3) -C(O)NHSO<sub>2</sub>R<sub>1</sub>;
- 4) -C(O)NR<sub>10</sub>R<sub>11</sub>;
- 5) Hydrogen;
- 6) Substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl;
- 7) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl;
- 8) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl;
- 9) Substituert eller usubstituert arylalkyl; og
- 40 10) Substituert eller usubstituert aryl;

eller R<sub>a</sub> og R<sub>b</sub> er tatt sammen med nitrogenatomet som de er bundet til for å danne en heterosyklisk; R<sub>1</sub> er valgt fra gruppen som består av:

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- 1) Halogen;
- 2) Hydroksyl;
- 3) Substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl;
- 4) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl;
- 5) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl;
- 6) Substituert eller usubstituert -C<sub>3</sub>-C<sub>8</sub> cykloalkyl;
- 50 7) Substituert eller usubstituert aryl;
- 8) Substituert eller usubstituert arylalkyl;
- 9) Substituert eller usubstituert heterosykloalkyl;
- 10) Substituert eller usubstituert heteroaryl;
- 11) Substituert eller usubstituert heteroarylalkyl; og
- 55 12) -NR<sub>10</sub>R<sub>11</sub>;

R<sub>2</sub> er valgt fra gruppen som består av:

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- 1) Hydrogen;
- 2) Substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl;
- 3) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl;

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- 5           4) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl;  
 5) Substituert eller usubstituert arylalkyl; og  
 6) Substituert eller usubstituert aryl;

R<sub>c</sub> er valgt fra gruppen som består av:

- 10           1) Hydrogen;  
 2) Substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl;  
 3) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl;  
 4) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl;  
 5) Substituert eller usubstituert arylalkyl; og  
 15           6) Substituert eller usubstituert aryl;

eller R<sub>2</sub> og R<sub>c</sub> er tatt sammen med karbonatomet som de er bundet til for å danne en syklisk ring; m er valgt fra 0, 1, 2 og 3;

R<sub>3</sub> er hydrogen, hydroksyl, -OSO<sub>3</sub>H, -OSO<sub>3</sub><sup>-</sup>, -OAc, -OPO<sub>3</sub>H<sub>2</sub> eller -OPO<sub>3</sub><sup>2-</sup>;

- 20           R<sub>4</sub> er hydrogen, halogen, CN, N<sub>3</sub>, hydroksyl, -OSO<sub>3</sub>H, -OSO<sub>3</sub><sup>-</sup>, -OAc, -OPO<sub>3</sub>H<sub>2</sub>, -OPO<sub>3</sub><sup>2-</sup>,  
 -SR<sub>2</sub> eller -NHR<sub>2</sub>;

eller R<sub>3</sub> og R<sub>4</sub> er tatt sammen med karbonene de fester fra -CH = CH- eller cykloalkylring eller heterocy- cloalkylring;

- 25           R<sub>5</sub> og R<sub>6</sub> er uavhengig hydrogen eller hydroksyl beskyttelsesgruppe; R<sub>7</sub> er valgt fra gruppen som består av:

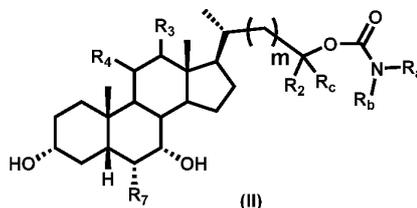
- 1) Halogen;  
 2) Substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl;  
 3) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl;  
 30           4) Substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl; og  
 5) Substituert eller usubstituert -C<sub>3</sub>-C<sub>8</sub> cykloalkyl; og

R<sub>10</sub> og R<sub>11</sub> er hver uavhengig valgt fra hydrogen, substituert eller usubstituert -C<sub>1</sub>-C<sub>8</sub> alkyl, substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkenyl, substituert eller usubstituert -C<sub>2</sub>-C<sub>8</sub> alkynyl, substituert eller usubstituert -C<sub>3</sub>-C<sub>8</sub> cykloalkyl, eller R<sub>10</sub> og R<sub>11</sub> er tatt sammen med nitrogenatomet som de er bundet til for å danne en heterosyklisk ring.

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2. Forbindelsen ifølge patentkrav 1, representert ved formel II eller et farmasøytisk akseptabelt salt derav:

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- 50           hvor i R<sub>a</sub>, R<sub>b</sub>, R<sub>c</sub>, R<sub>2</sub>, R<sub>3</sub>, R<sub>4</sub>, R<sub>7</sub> og m er som definert i patentkrav 1.

3. Forbindelsen ifølge patentkrav 1, representert ved en av formlene (III-1-III-18) eller et farmasøytisk akseptabelt salt derav:

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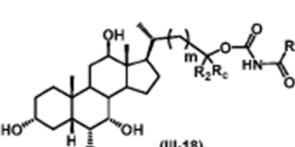
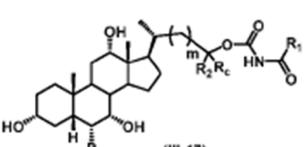
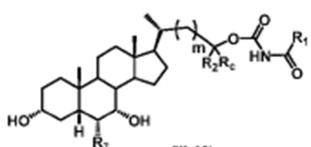
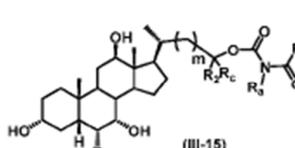
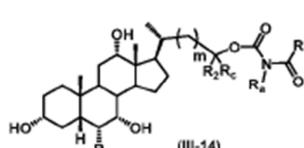
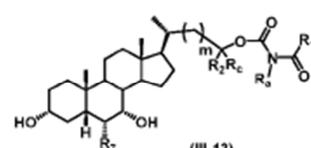
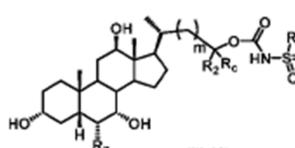
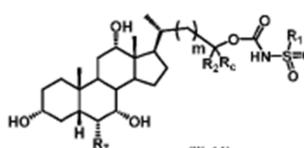
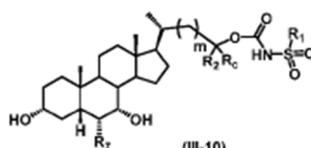
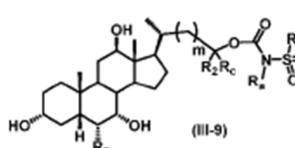
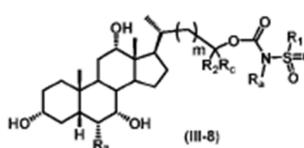
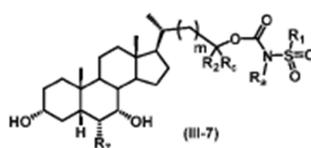
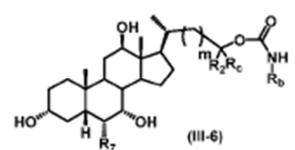
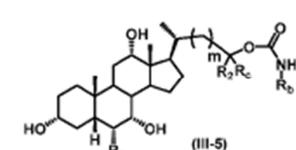
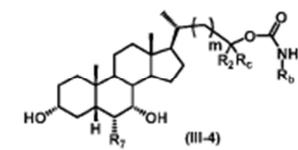
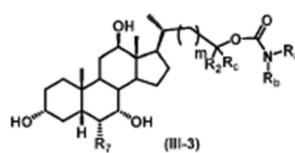
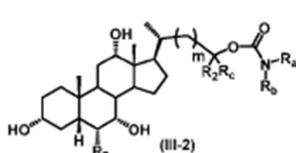
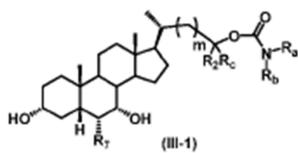
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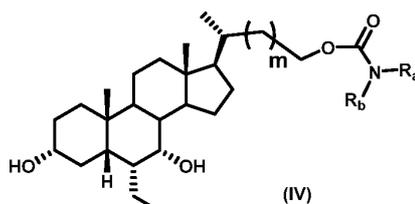


hvor,  $R_a$ ,  $R_b$ ,  $R_c$ ,  $R_1$ ,  $R_2$ ,  $R_7$  og  $m$  er som definert i patentkrav 1.

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4. Forbindelsen ifølge patentkrav 1, representert ved formel IV eller et farmasøytisk akseptabelt salt derav:

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hvor,  $R_a$ ,  $R_b$  og  $m$  er som definert i patentkrav 1.

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5. Forbindelse ifølge patentkrav 4, valgt fra forbindelser med formel IV hvor,  $R_a$ ,  $R_b$  og  $m$  er avgrenset for hver forbindelse i tabell 1 eller et farmasøytisk akseptabelt salt derav:

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**Tabell 1**

	Eksempel	m	R <sub>a</sub>	R <sub>b</sub>	Eksempel	m	R <sub>a</sub>	R <sub>b</sub>	Eksempel	m	R <sub>a</sub>	R <sub>b</sub>
	1	0	Metyl	H	34	1	Metyl	H	67	2	Metyl	H
10	2	0	Etyl	H	35	1	Etyl	H	68	2	Etyl	H
	3	0	Klorprofam	H	36	1	Klorprofam	H	69	2	Klorprofam	H
	4	0	Butyl	H	37	1	Butyl	H	70	2	Butyl	H
15	5	0	t-Butyl	H	38	1	t-Butyl	H	71	2	t-Butyl	H
	6	0	Propyl	H	39	1	Propyl	H	72	2	Propyl	H
	7	0	Benzyl	H	40	1	Benzyl	H	73	2	Benzyl	H
	8	0	Vinyl	H	41	1	Vinyl	H	74	2	Vinyl	H
20	9	0	Allyl	H	42	1	Allyl	H	75	2	Allyl	H
	10	0	CF <sub>3</sub>	H	43	1	CF <sub>3</sub>	H	76	2	CF <sub>3</sub>	H
	11	0		H	44	1		H	77	2		H
25	12	0		H	45	1		H	78	2		H
	13	0		H	46	1		H	79	2		H
30	14	0		H	47	1		H	80	2		H
	15	0	Metyl	Me	48	1	Metyl	Me	81	2	Metyl	Me
	16	0	Etyl	Me	49	1	Etyl	Me	82	2	Etyl	Me
35	17	0	Klorprofam	Me	50	1	Klorprofam	Me	83	2	Klorprofam	Me
	18	0	t-Butyl	Me	51	1	t-Butyl	Me	84	2	t-Butyl	Me
	19	0	Propyl	Me	52	1	Propyl	Me	85	2	Propyl	Me
40	20	0	Benzyl	Me	53	1	Benzyl	Me	86	2	Benzyl	Me
	21	0	Vinyl	Me	54	1	Vinyl	Me	87	2	Vinyl	Me
	22	0	Allyl	Me	55	1	Allyl	Me	88	2	Allyl	Me
	23	0	CF <sub>3</sub>	Me	56	1	CF <sub>3</sub>	Me	89	2	CF <sub>3</sub>	Me
45	24	0		Me	57	1		Me	90	2		Me
	25	0		Me	58	1		Me	91	2		Me
	26	0		Me	59	1		Me	92	2		Me
50	27	0			60	1			93	2		
	28	0			61	1			94	2		
55	29	0			62	1			95	2		
60	30	0			63	1			96	2		

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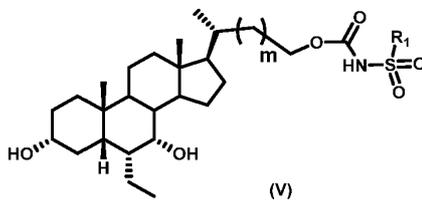
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Eksempel	m	R <sub>a</sub>	R <sub>b</sub>	Eksempel	m	R <sub>a</sub>	R <sub>b</sub>	Eksempel	m	R <sub>a</sub>	R <sub>b</sub>
31	0			64	1			97	2		
32	0			65	1			98	2		
33	0			66	1			99	2		

6. Forbindelsen ifølge patentkrav 1, representert ved formel V

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eller et farmasøytisk akseptabelt salt derav.

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7. Forbindelse ifølge krav 6, som er valgt fra forbindelser med formel V, eller et farmasøytisk akseptabelt salt derav hvori, R<sub>1</sub> og m er avgrenset for hver forbindelse i tabell 2:

**Tabell 2**

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Eksempel	m	R <sub>1</sub>	Eksempel	m	R <sub>1</sub>	Eksempel	m	R <sub>1</sub>
100	0	Metyl	127	1	Metyl	154	2	Metyl
101	0	Etyl	128	1	Etyl	155	2	Etyl
102	0	Klorprofam	129	1	Klorprofam	156	2	Klorprofam
103	0	Butyl	130	1	Butyl	157	2	Butyl
104	0	t-Butyl	131	1	t-Butyl	158	2	t-Butyl
105	0	Propyl	132	1	Propyl	159	2	Propyl
106	0	Benzyl	133	1	Benzyl	160	2	Benzyl
107	0	Vinyl	134	1	Vinyl	161	2	Vinyl
108	0	Allyl	135	1	Allyl	162	2	Allyl
109	0	CF <sub>3</sub>	136	1	CF <sub>3</sub>	163	2	CF <sub>3</sub>
110	0		137	1		164	2	
111	0		138	1		165	2	

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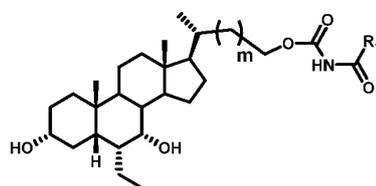
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Eksempel	m	R <sub>1</sub>	Eksempel	m	R <sub>1</sub>	Eksempel	m	R <sub>1</sub>
112	0		139	1		166	2	
113	0		140	1		167	2	
114	0		141	1		168	2	
115	0		142	1		169	2	
116	0		143	1		170	2	
117	0		144	1		171	2	
118	0	NH <sub>2</sub>	145	1	NH <sub>2</sub>	172	2	NH <sub>2</sub>
119	0		146	1		173	2	
120	0		147	1		174	2	
121	0		148	1		175	2	
122	0		149	1		176	2	
123	0		150	1		177	2	
124	0		151	1		178	2	
125	0		152	1		179	2	
126	0	F	153	1	F	180	2	F

8. En forbindelsen ifølge patentkrav 1, valgt fra forbindelser med formel VI:

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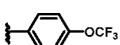
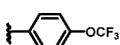
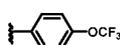
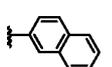
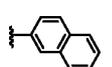
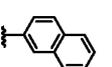
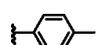
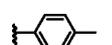
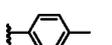
(VI)

hvor R<sub>1</sub> og m for hver forbindelse er angitt i tabell 3, eller et farmasøytisk akseptabelt salt derav:

## EP 3 277 286 B1

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Tabell 3

Forbindelse	m	R <sub>1</sub>	Forbindelse	m	R <sub>1</sub>	Forbindelse	m	R <sub>1</sub>
181	0	Metyl	208	1	Metyl	235	2	Metyl
182	0	Etyl	209	1	Etyl	236	2	Etyl
183	0	Klorprofam	210	1	Klorprofam	237	2	Klorprofam
184	0	Butyl	211	1	Butyl	238	2	Butyl
185	0	t-Butyl	212	1	t-Butyl	239	2	t-Butyl
186	0	Propyl	213	1	Propyl	240	2	Propyl
187	0	Benzyl	214	1	Benzyl	241	2	Benzyl
188	0	Vinyl	215	1	Vinyl	242	2	Vinyl
189	0	Allyl	216	1	Allyl	243	2	Allyl
190	0	CF <sub>3</sub>	217	1	CF <sub>3</sub>	244	2	CF <sub>3</sub>
191	0		218	1		245	2	
192	0		219	1		246	2	
193	0		220	1		247	2	
194	0		221	1		248	2	
195	0		222	1		249	2	
196	0		223	1		250	2	
197	0		224	1		251	2	
198	0		225	1		252	2	
199	0	NH <sub>2</sub>	226	1	NH <sub>2</sub>	253	2	NH <sub>2</sub>
200	0		227	1		254	2	
201	0		228	1		255	2	
202	0		229	1		256	2	
203	0		230	1		257	2	
204	0		231	1		258	2	
205	0		232	1		259	2	
206	0		233	1		260	2	
207	0	F	234	1	F	261	2	F

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- 5 9. En forbindelse ifølge patentkrav 1, valgt fra forbindelsene angitt nedenfor, eller et farmasøytisk akseptabelt salt derav:

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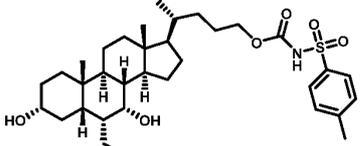
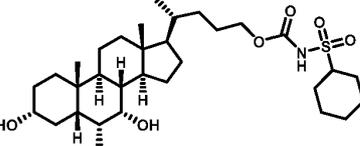
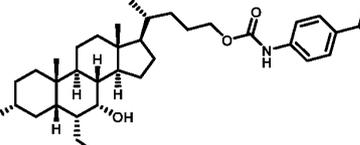
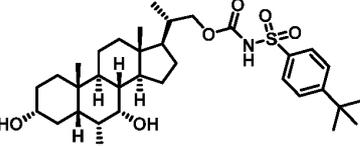
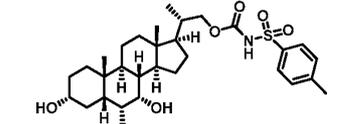
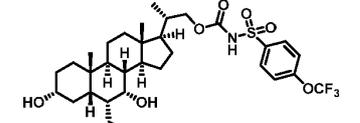
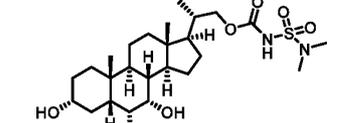
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Eksempel #	Struktur
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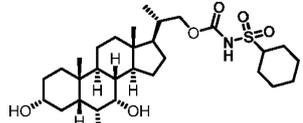
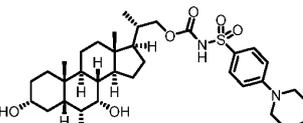
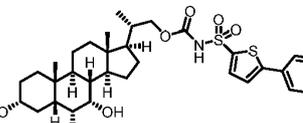
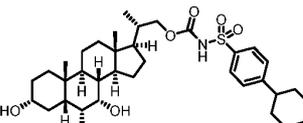
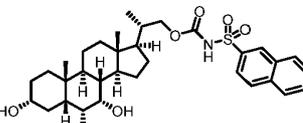
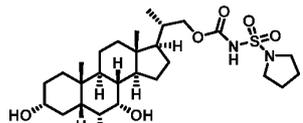
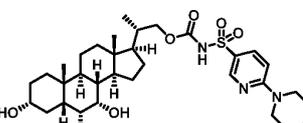
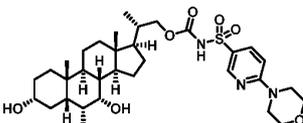
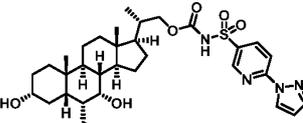
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Eksempel #	Struktur
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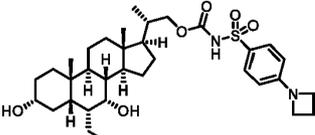
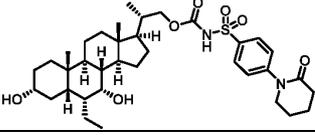
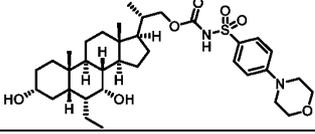
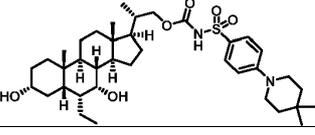
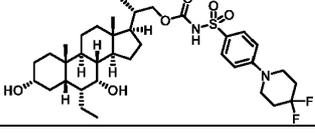
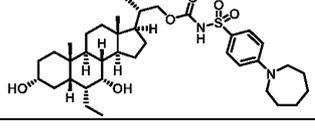
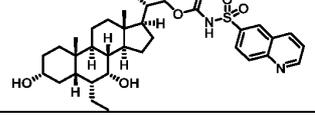
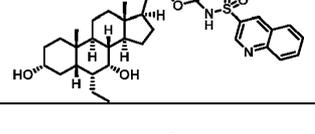
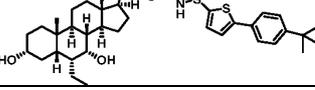
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Eksempel #	Struktur
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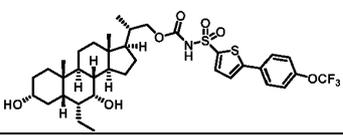
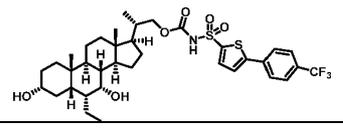
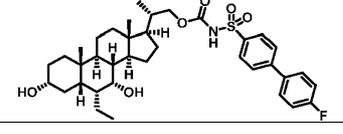
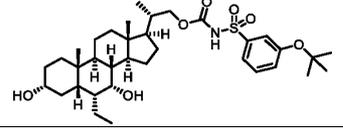
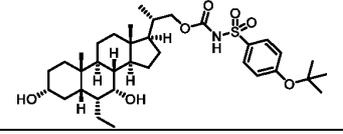
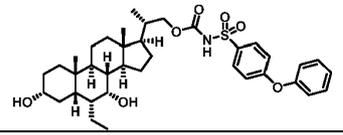
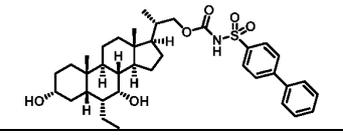
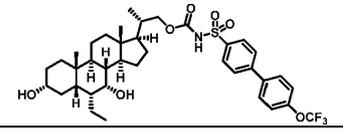
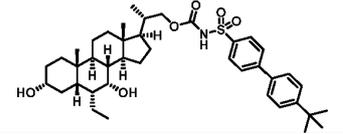
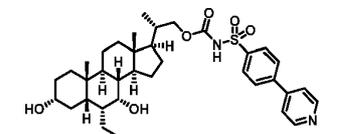
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Eksempel #	Struktur
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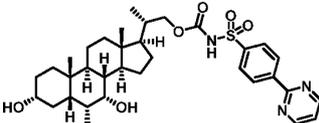
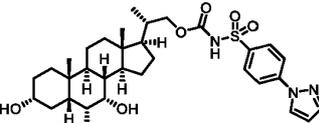
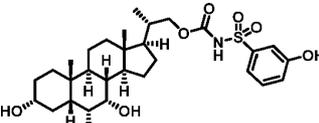
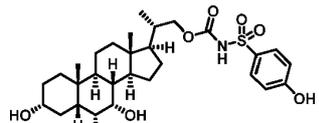
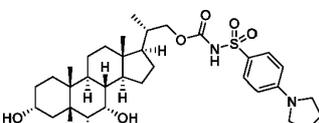
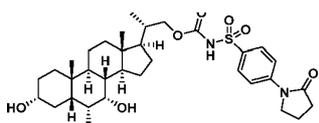
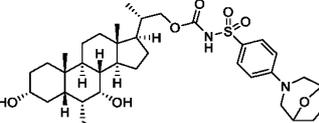
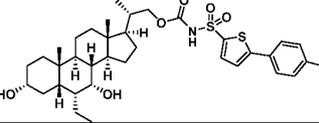
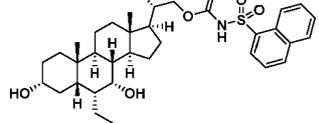
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Eksempel #	Struktur
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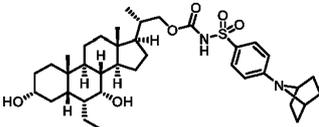
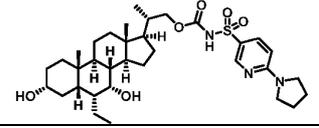
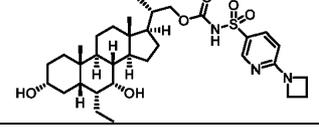
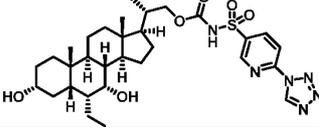
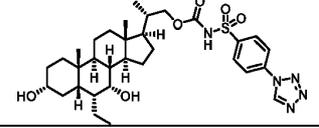
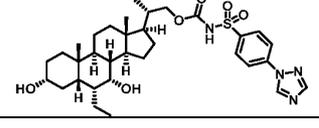
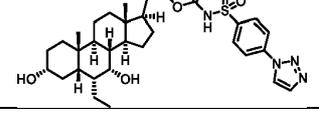
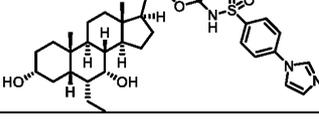
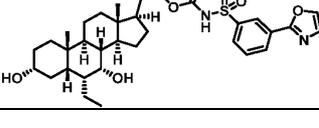
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Eksempel #	Struktur
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44-b	
44-c	
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44-f	
44-g	
44-h	
44-i	

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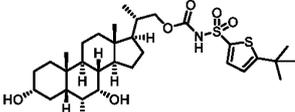
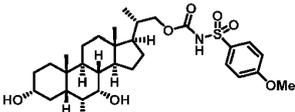
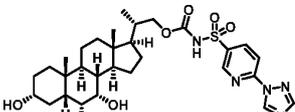
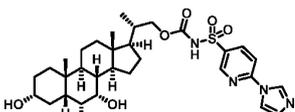
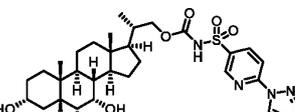
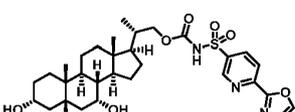
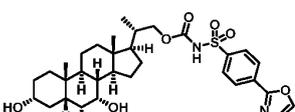
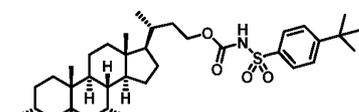
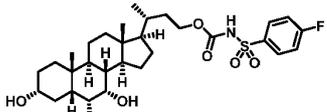
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Eksempel #	Struktur
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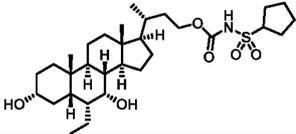
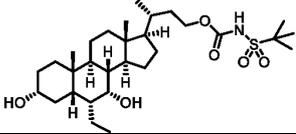
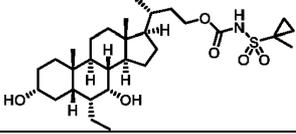
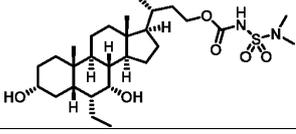
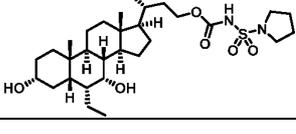
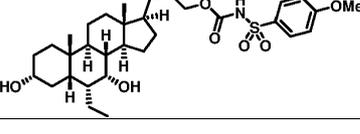
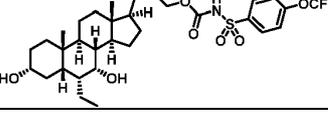
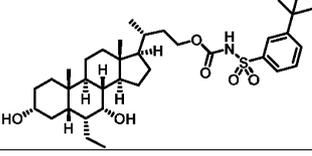
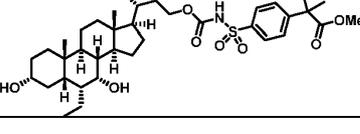
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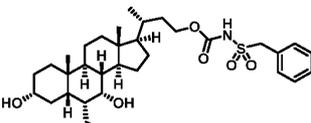
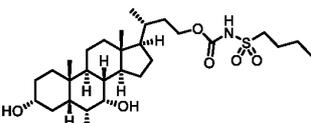
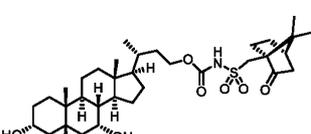
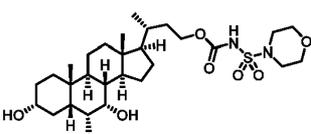
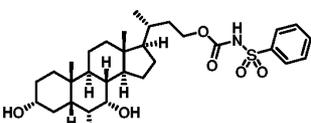
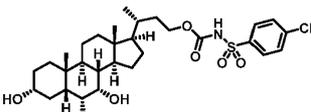
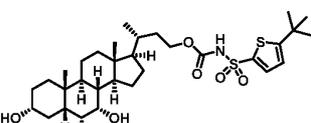
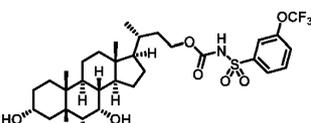
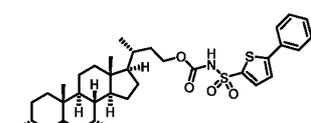
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Eksempel #	Struktur
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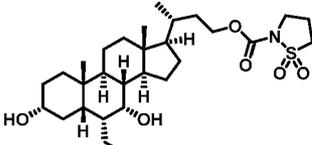
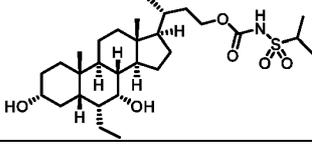
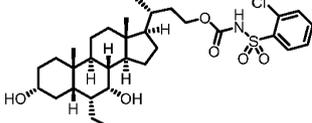
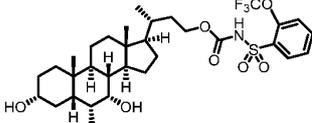
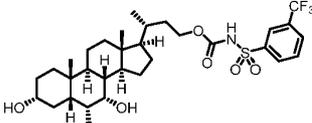
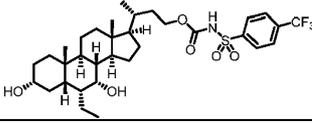
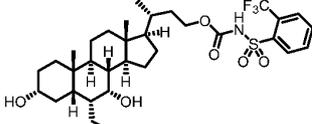
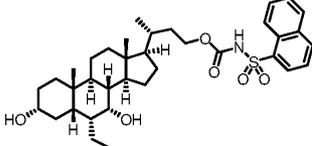
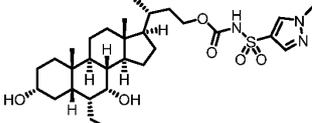
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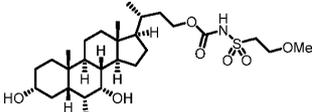
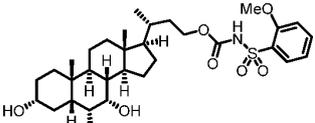
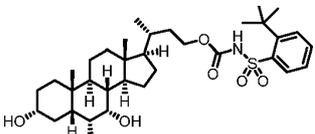
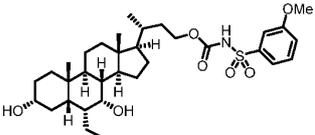
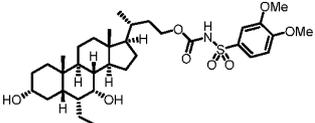
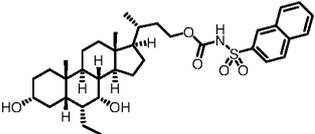
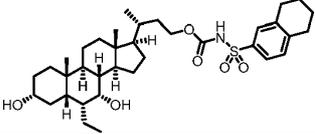
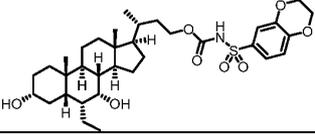
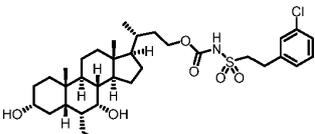
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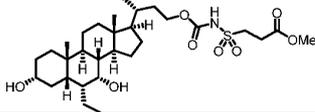
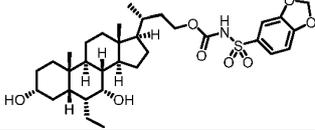
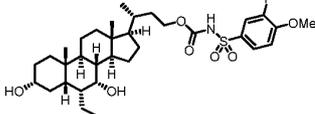
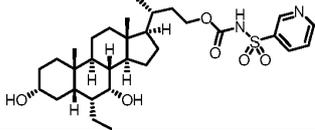
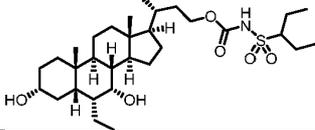
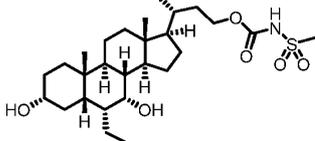
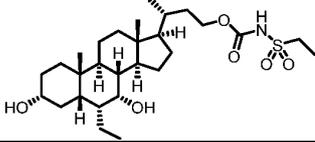
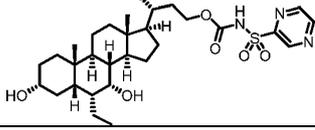
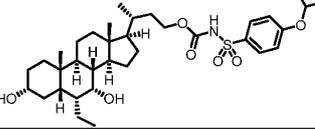
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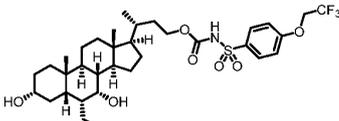
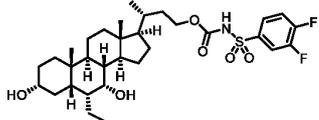
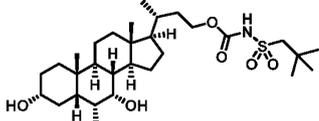
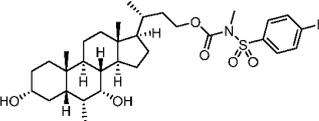
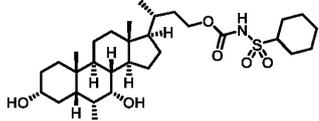
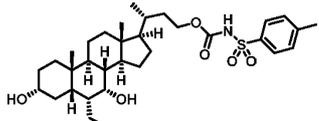
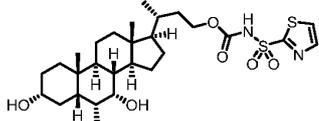
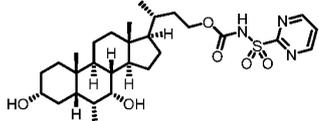
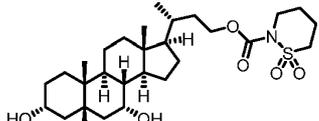
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Eksempel #	Struktur
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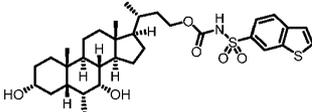
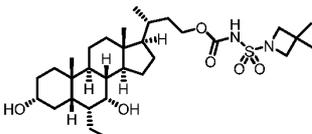
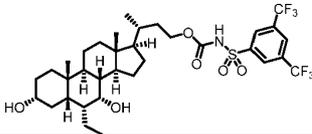
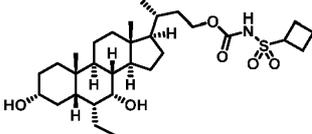
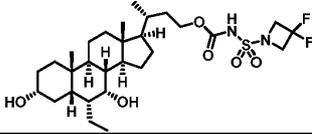
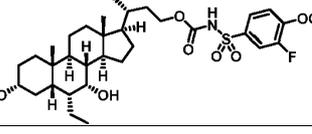
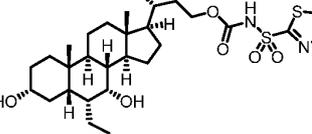
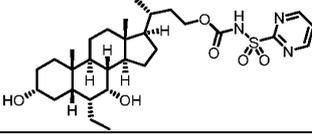
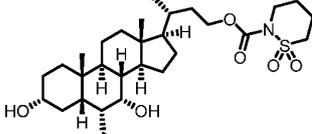
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Eksempel #	Struktur
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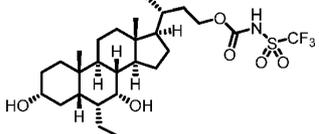
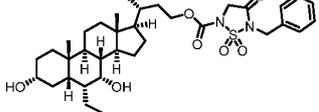
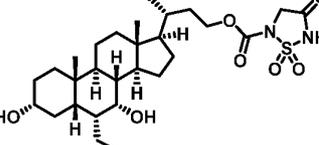
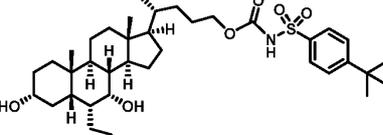
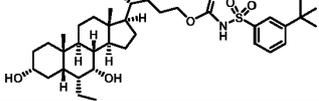
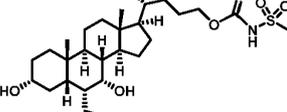
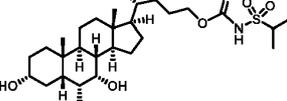
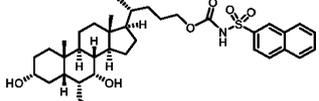
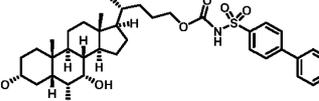
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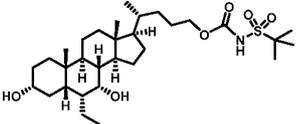
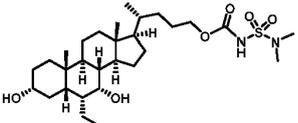
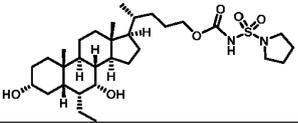
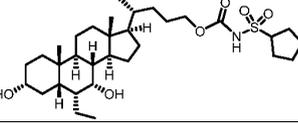
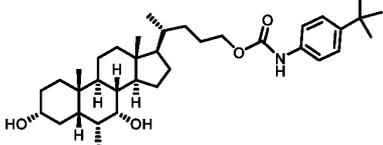
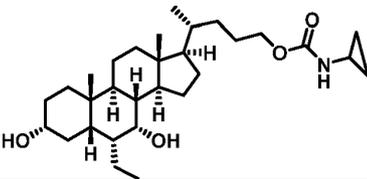
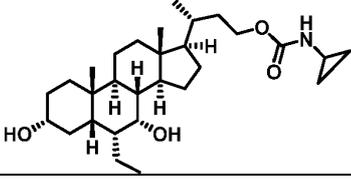
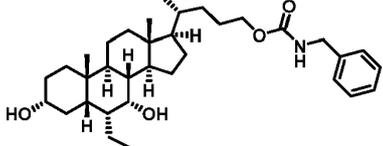
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Eksempel #	Struktur
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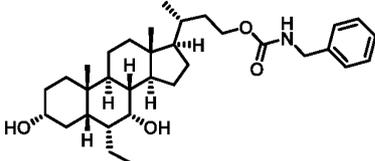
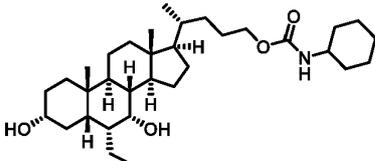
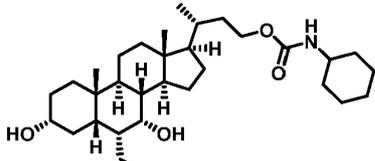
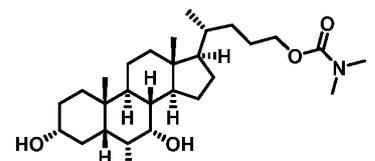
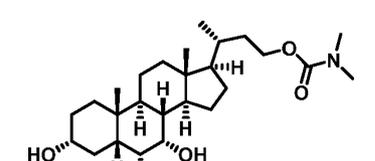
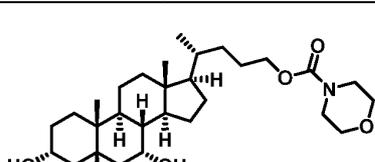
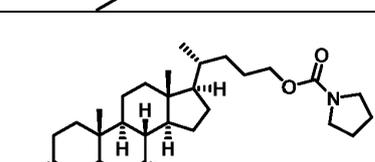
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Eksempel #	Struktur
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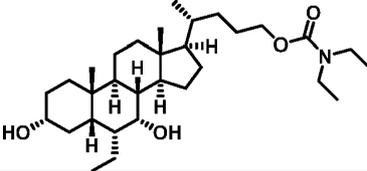
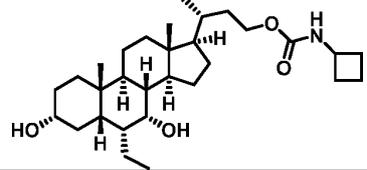
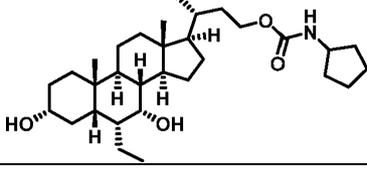
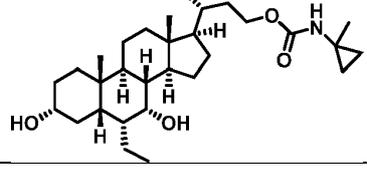
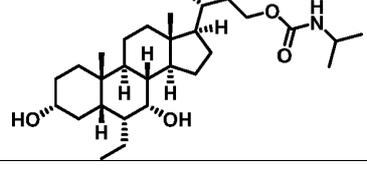
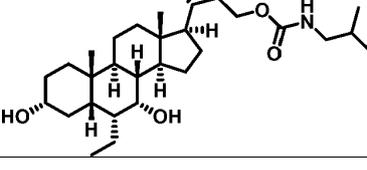
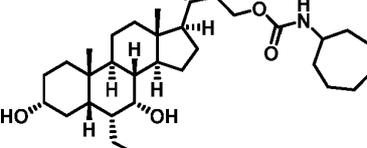
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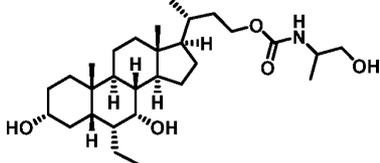
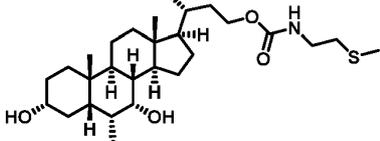
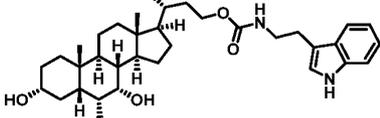
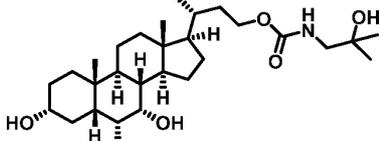
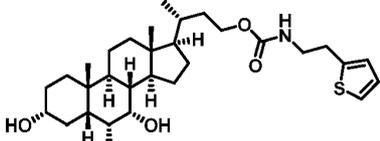
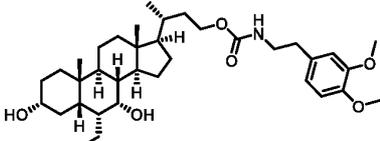
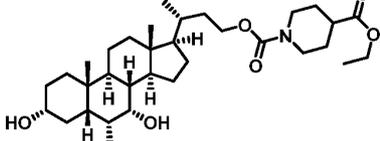
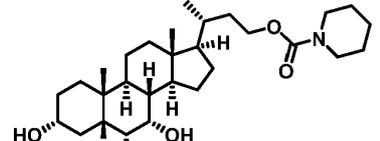
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Eksempel #	Struktur
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Eksempel #	Struktur
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## EP 3 277 286 B1

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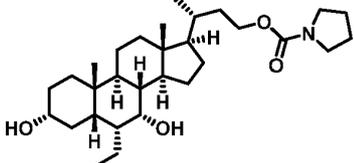
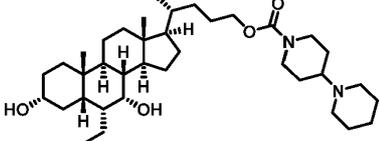
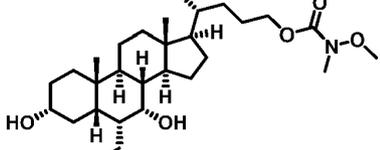
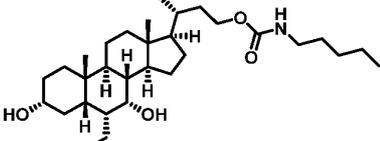
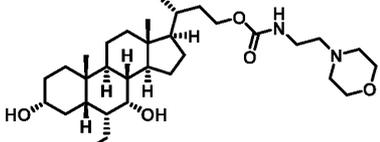
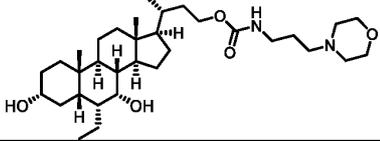
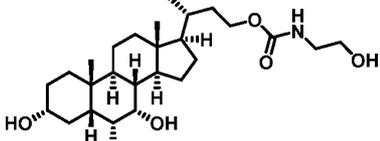
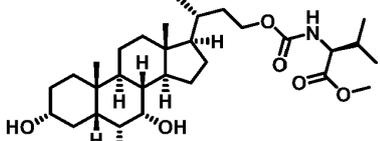
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Eksempel #	Struktur
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## EP 3 277 286 B1

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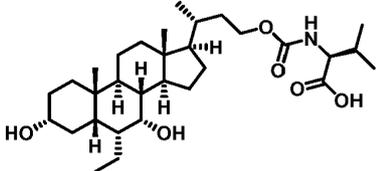
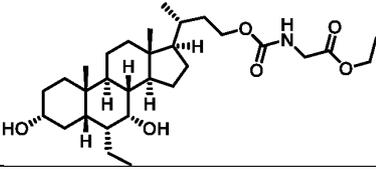
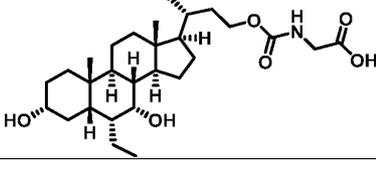
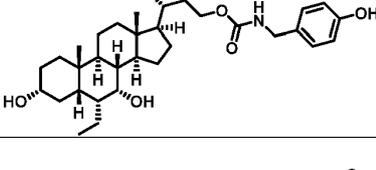
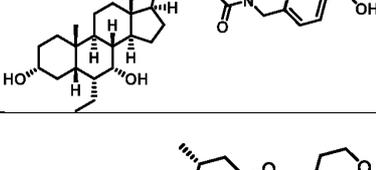
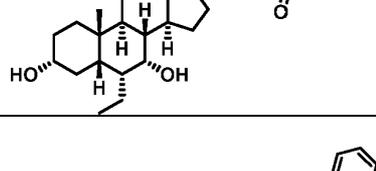
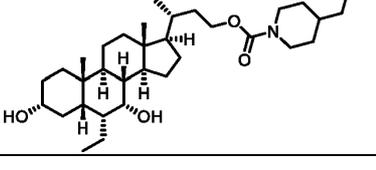
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Eksempel #	Struktur
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## EP 3 277 286 B1

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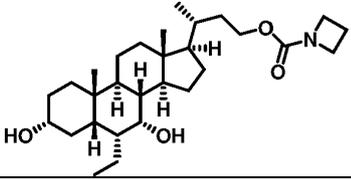
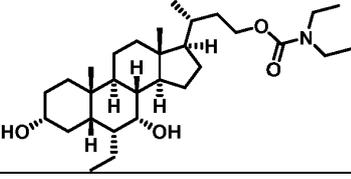
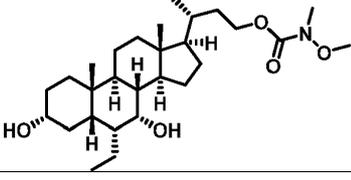
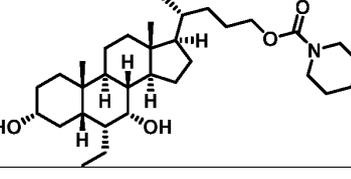
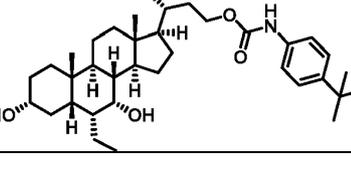
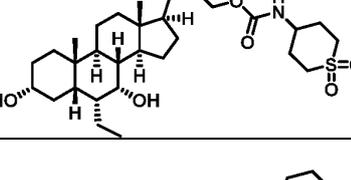
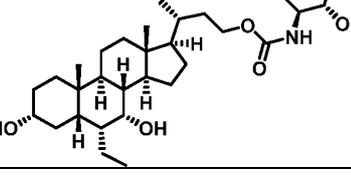
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Eksempel #	Struktur
162	 <p>Chemical structure of a steroid derivative (Example 162). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a pyrrolidine ring via an ester linkage.</p>
163	 <p>Chemical structure of a steroid derivative (Example 163). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a diethylamide group via an ester linkage.</p>
164	 <p>Chemical structure of a steroid derivative (Example 164). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a dimethylamide group via an ester linkage.</p>
165	 <p>Chemical structure of a steroid derivative (Example 165). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a piperidine ring via an ester linkage.</p>
166	 <p>Chemical structure of a steroid derivative (Example 166). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a 4-tert-butylphenylamide group via an ester linkage.</p>
167	 <p>Chemical structure of a steroid derivative (Example 167). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a 1,2,4-thiazolidine-5-sulfonamide group via an ester linkage.</p>
168	 <p>Chemical structure of a steroid derivative (Example 168). The steroid core has hydroxyl groups at C-3 and C-14, and a methyl group at C-10. At C-3, there is a side chain consisting of a propyl group linked to a cyclohexanone-3-ylamide group via an ester linkage.</p>

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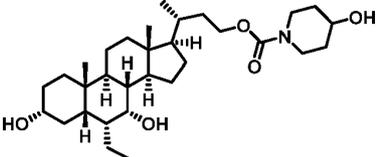
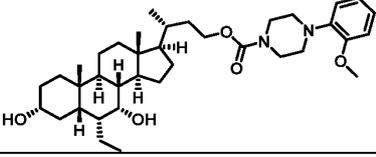
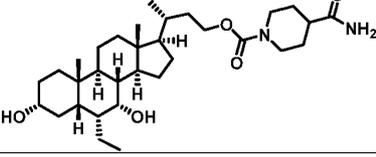
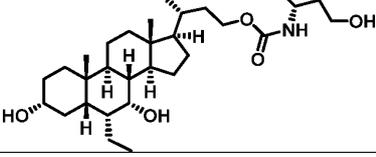
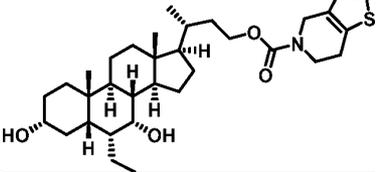
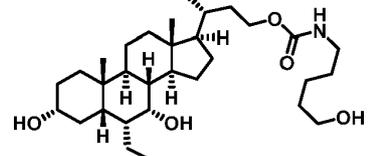
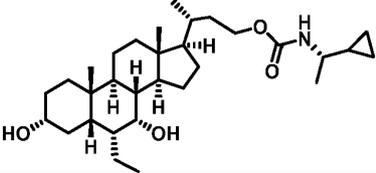
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Eksempel #	Struktur
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## EP 3 277 286 B1

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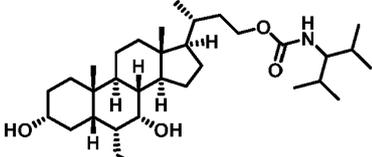
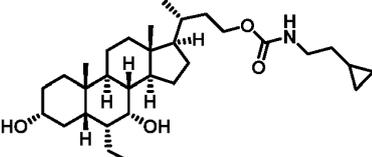
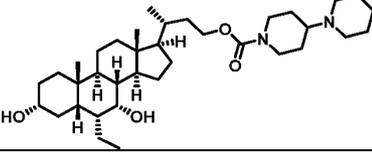
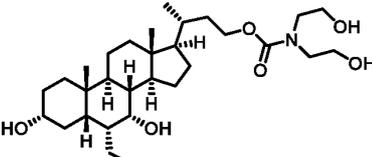
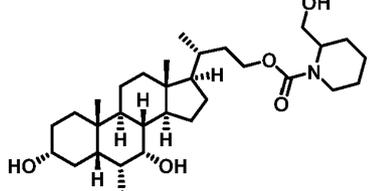
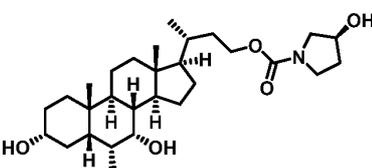
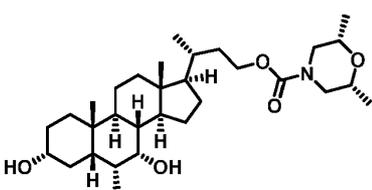
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Eksempel #	Struktur
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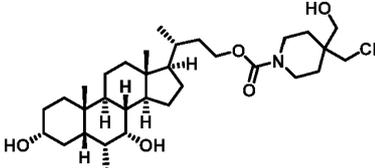
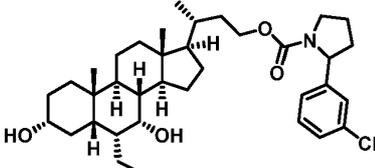
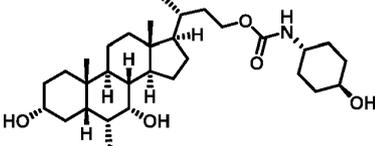
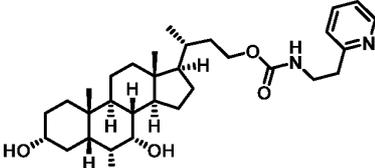
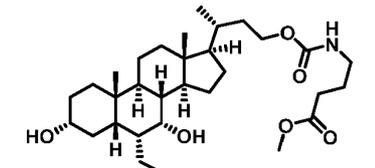
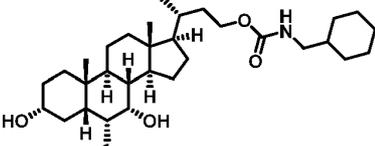
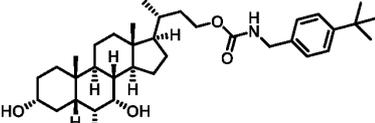
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Eksempel #	Struktur
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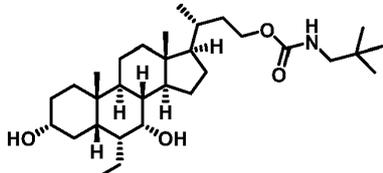
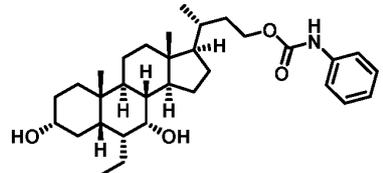
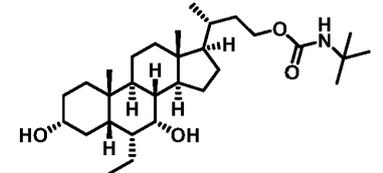
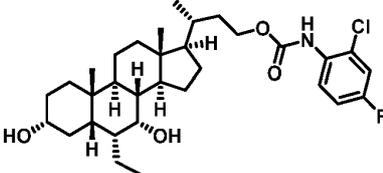
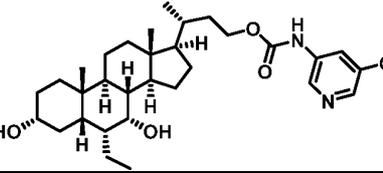
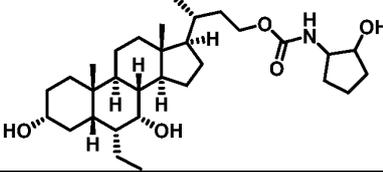
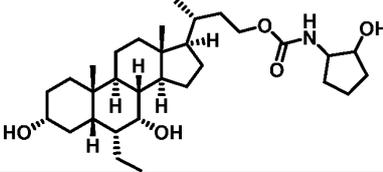
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Eksempel #	Struktur
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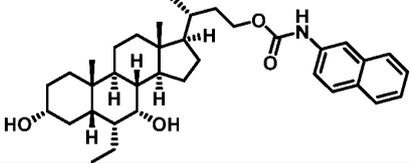
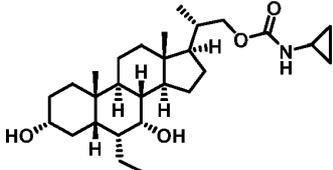
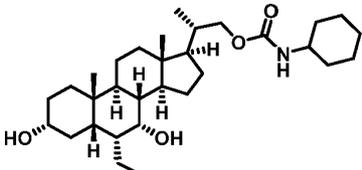
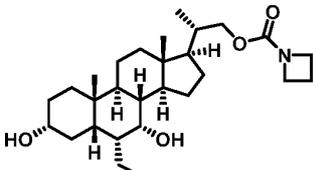
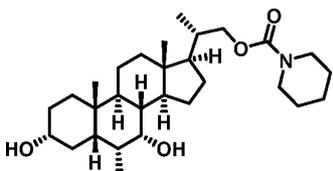
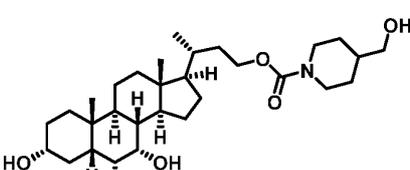
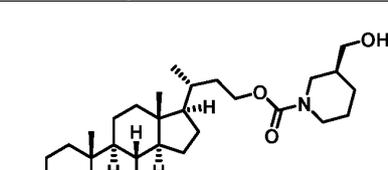
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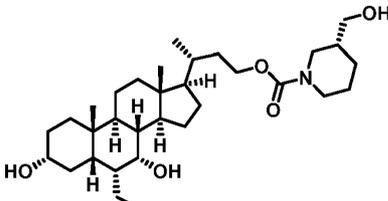
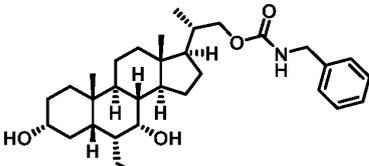
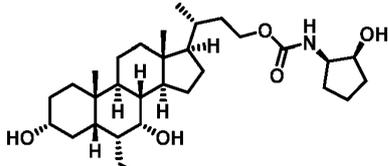
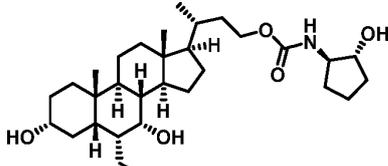
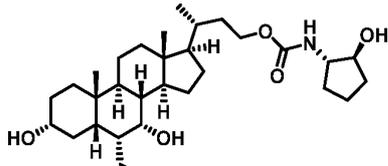
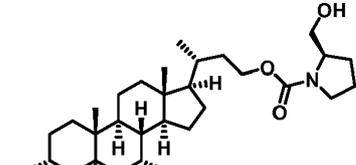
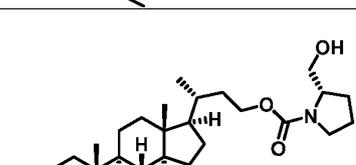
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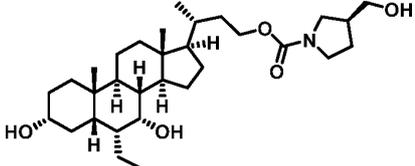
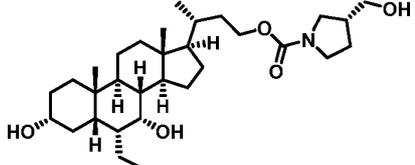
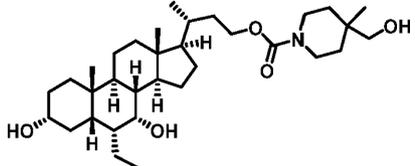
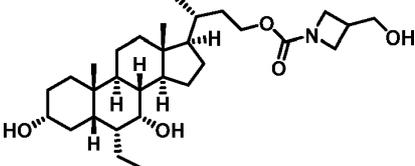
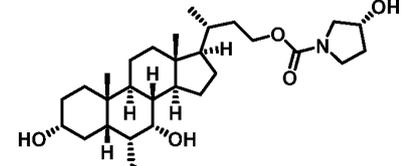
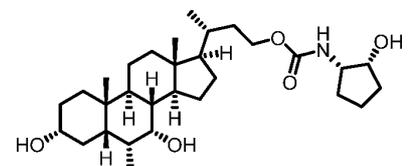
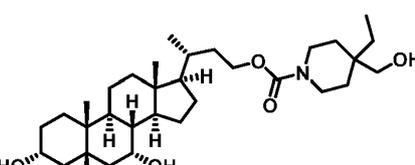
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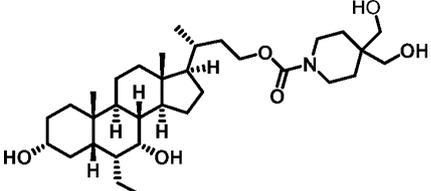
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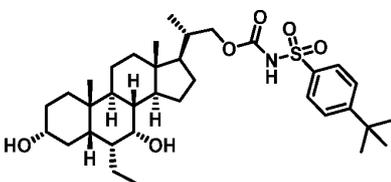
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10. Forbindelsen ifølge patentkrav 9, som har strukturen

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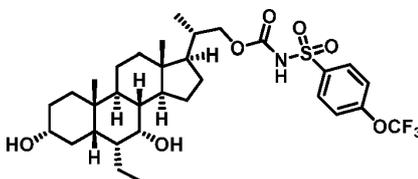


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eller et farmasøytisk akseptabelt salt derav.

11. Forbindelsen ifølge patentkrav 9, som har strukturen

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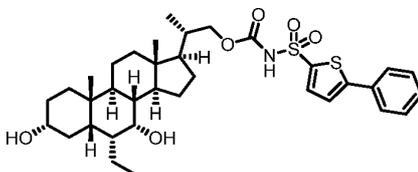


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eller et farmasøytisk akseptabelt salt derav.

12. Forbindelsen ifølge patentkrav 9, som har strukturen

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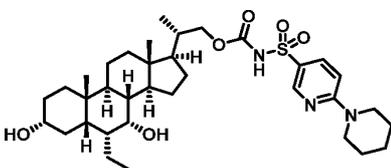


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eller et farmasøytisk akseptabelt salt derav.

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13. Forbindelsen ifølge patentkrav 9, som har strukturen

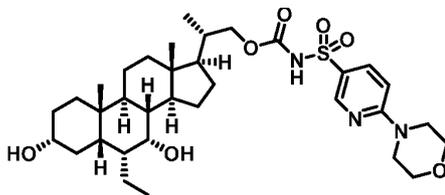


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5 eller et farmasøytisk akseptabelt salt derav.

14. Forbindelsen ifølge patentkrav 9, som har strukturen

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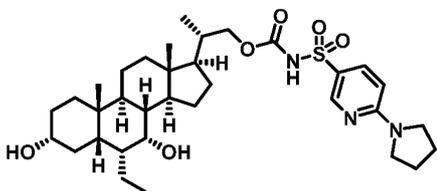


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eller et farmasøytisk akseptabelt salt derav.

15. Forbindelsen ifølge patentkrav 9, som har strukturen

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eller et farmasøytisk akseptabelt salt derav.

30 16. Forbindelsen ifølge hvilket som helst av patentkravene 1–15 for anvendelse for å forhindre eller behandle en FXR-mediert sykdom eller tilstand.

35 17. Forbindelsen for anvendelse ifølge patentkrav 16, hvor den FXR-medierte sykdommen eller tilstanden er valgt fra gruppen bestående av kronisk leversykdom, gastrointestinal sykdom, nyresykdom, kardiovaskulær sykdom og metabolsk sykdom.

40 18. Forbindelsen for anvendelse ifølge patentkrav 17, hvori den FXR-medierte sykdommen eller tilstanden er en kronisk leversykdom valgt fra gruppen bestående av primær biliær kolangitt, cerebrotendinøs xantomatose, primær skleroserende kolangitt, medikamentindusert kolestase, intrahepatisk svangerskapskolestase, parenteral ernæringsassosiert kolestase, bakteriell gjengroing eller sepsisassosiert kolestase, autoimmun hepatitt, kronisk viral hepatitt, alkoholisk leversykdom, ikke-alkoholisk fettleversykdom, ikke-alkoholisk steatohepatitt, levertransplantasjon assosiert transplantat versus vertssykdom, levende donortransplantasjon leverregenerering, medfødt leverfibrose, koledokolitase, granulomatøs leversykdom, intra- eller ekstrahepatisk malignitet, Sjögrens syndrom, sarkoidose, Wilsons sykdom, Gauchers sykdom, hemokromatose og alfa 1-antitrypsinmangel.

50 19. Forbindelsen for anvendelse ifølge patentkrav 17, hvor den FXR-medierte sykdommen eller tilstanden er en nyresykdom valgt fra gruppen bestående av diabetisk nefropati, fokal segmentell glomerulosklerose, hypertensiv nefrosklerose, kronisk glomerulonefritt, kronisk transplantasjonsglomerulopati, kronisk interstitiell nefritt og polycystisk nyresykdom;

55 20. Forbindelsen for anvendelse ifølge patentkrav 17, hvor den FXR-medierte sykdommen eller tilstanden er en kardiovaskulær sykdom valgt fra gruppen bestående av (a) aterosklerose, arteriosklerose, dyslipidemi, hyperkolesterolemi og hypertriglyseridemi; eller (b) en metabolsk sykdom valgt fra gruppen bestående av insulinresistens, diabetes type 1 og type 2 og fedme.

60 21. En farmasøytisk sammensetning som omfatter forbindelsen ifølge hvilket som helst av patentkravene 1–15 og en farmasøytisk akseptabel bærer.