

**Table S1.** Two-way ANOVA of DPPH Inhibition Activity

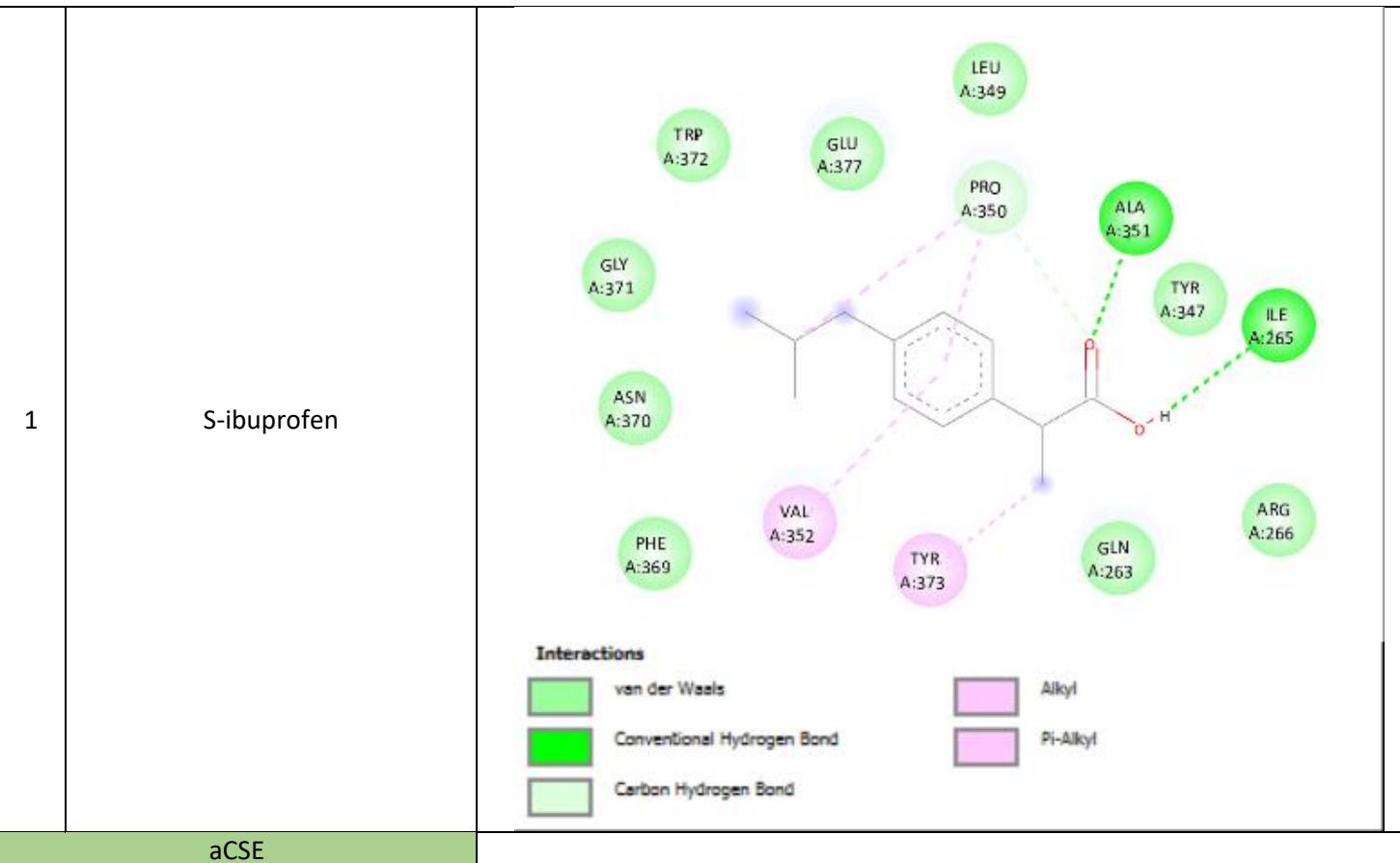
| DPPH; Dunnett's multiple comparisons test | Mean Diff. | 95.00% CI of diff. | Below threshold? | Summary | Adjusted P Value |
|---|------------|--------------------|------------------|---------|------------------|
| <b>Row 1 or 25 µg/mL</b>                  |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE                   | -0.5600    | -2.860 to 1.740    | No               | ns      | 0.7966           |
| Trolox/Control vs. aCSE                   | 2.233      | -0.06640 to 4.533  | No               | ns      | 0.0578           |
| <b>Row 2 or 50 µg/mL</b>                  |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE                   | -0.3667    | -2.666 to 1.933    | No               | ns      | 0.9055           |
| Trolox/Control vs. aCSE                   | 2.733      | 0.4336 to 5.033    | Yes              | *       | 0.0184           |
| <b>Row 3 or 100 µg/mL</b>                 |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE                   | 0.6000     | -1.700 to 2.900    | No               | ns      | 0.7711           |
| Trolox/Control vs. aCSE                   | 6.467      | 4.167 to 8.766     | Yes              | ****    | <0.0001          |
| <b>Row 4 or 150 µg/mL</b>                 |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE                   | 3.577      | 1.277 to 5.876     | Yes              | **      | 0.0022           |
| Trolox/Control vs. aCSE                   | 5.133      | 2.834 to 7.433     | Yes              | ****    | <0.0001          |
| <b>Row 5 or 200 µg/mL</b>                 |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE                   | 0.06667    | -2.233 to 2.366    | No               | ns      | 0.9967           |
| Trolox/Control vs. aCSE                   | 2.840      | 0.5403 to 5.140    | Yes              | *       | 0.0142           |

**Table S2.** Two-way ANOVA of ABTS Inhibition Activity

| Dunnett's multiple comparisons test | Mean Diff. | 95.00% CI of diff. | Below threshold? | Summary | Adjusted P Value |
|-------------------------------------|------------|--------------------|------------------|---------|------------------|
| <b>Row 1 or 25 µg/mL</b>            |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE             | 3.143      | 0.6042 to 5.683    | Yes              | *       | 0.0140           |
| Trolox/Control vs. aCSE             | 5.973      | 3.434 to 8.513     | Yes              | ****    | <0.0001          |
| <b>Row 2 or 50 µg/mL</b>            |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE             | 1.970      | -0.5692 to 4.509   | No               | ns      | 0.1439           |
| Trolox/Control vs. aCSE             | 8.847      | 6.307 to 11.39     | Yes              | ****    | <0.0001          |
| <b>Row 3 or 100 µg/mL</b>           |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE             | 4.877      | 2.337 to 7.416     | Yes              | ***     | 0.0002           |
| Trolox/Control vs. aCSE             | 6.973      | 4.434 to 9.513     | Yes              | ****    | <0.0001          |
| <b>Row 4 or 150 µg/mL</b>           |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE             | 0.1367     | -2.403 to 2.676    | No               | ns      | 0.9886           |
| Trolox/Control vs. aCSE             | 3.773      | 1.234 to 6.313     | Yes              | **      | 0.0032           |
| <b>Row 5 or 200 µg/mL</b>           |            |                    |                  |         |                  |
| Trolox/Control vs. rCSE             | 2.403      | -0.1358 to 4.943   | No               | ns      | 0.0653           |
| Trolox/Control vs. aCSE             | 4.400      | 1.861 to 6.939     | Yes              | ***     | 0.0007           |

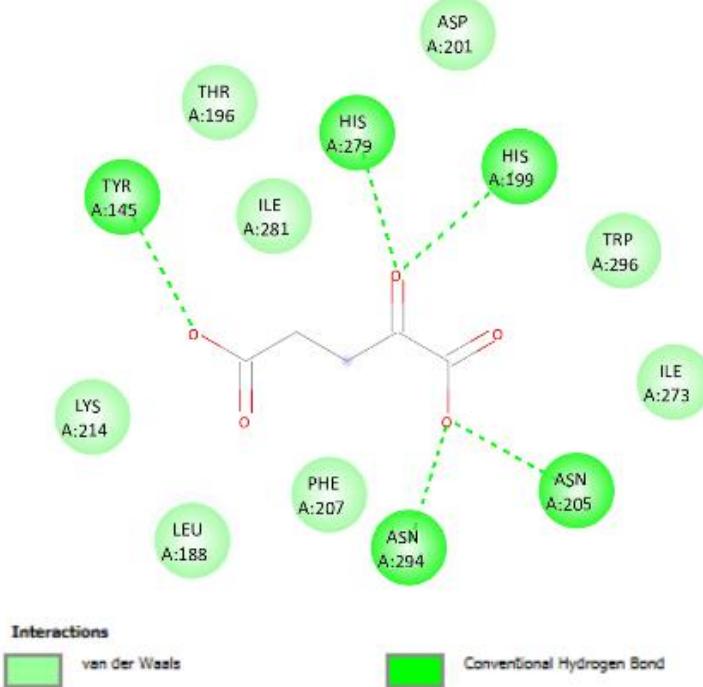
**Table S3.** The fully visualization from molecular docking simulation of amino acid interactions throughout the observed CSE substance

| <b>iNOS      3E7G</b> |               |   |
|-----------------------|---------------|---|
| No.                   | Substance     | Visualization   |
|                       | Native Ligand |   |
| 1                     | 3E7G          | <p><b>Interactions</b></p> <ul style="list-style-type: none"> <li>[Green Box] van der Waals</li> <li>[Green Box] Conventional Hydrogen Bond</li> <li>[Light Green Box] Carbon Hydrogen Bond</li> <li>[Purple Box] Alkyl</li> <li>[Pink Box] Pi-Alkyl</li> </ul> |
|                       | Control       |   |



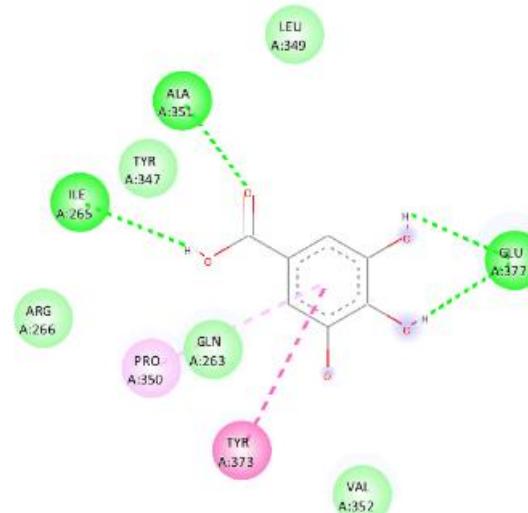
1

Epicatechin



2

Gallic acid

**Interactions**

van der Waals

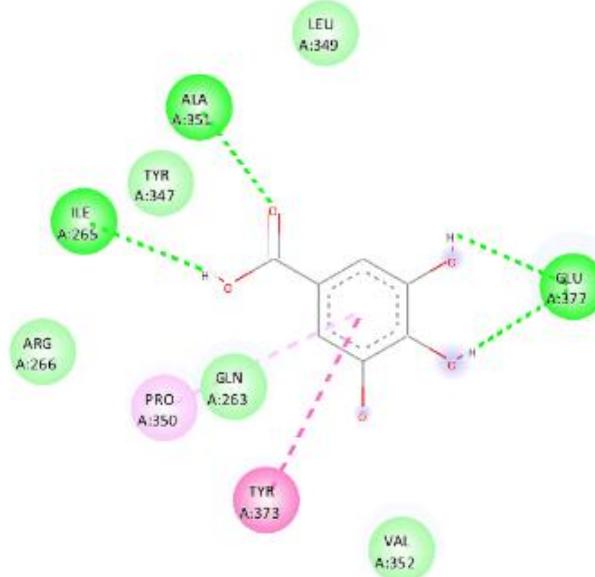
Conventional Hydrogen Bond

Pi-Pi T-shaped

Pi-Alkyl

3

## 4-Hydroxycinnamic acid



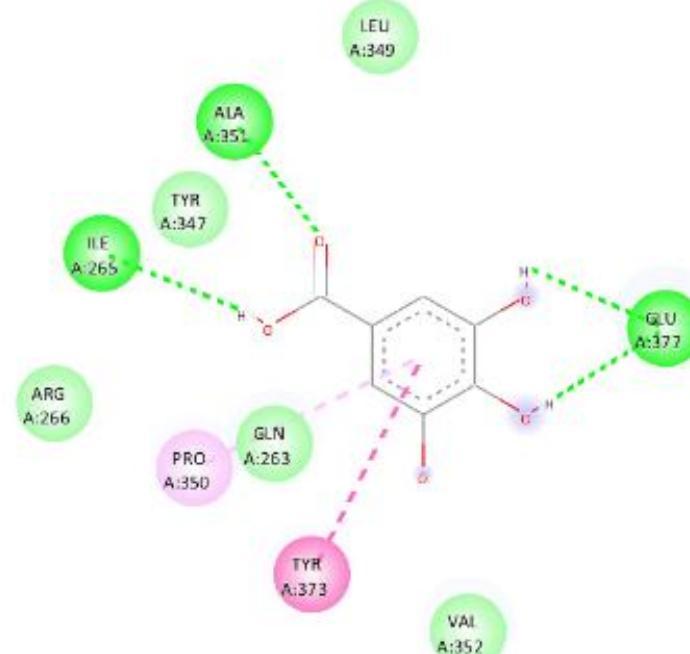
## Interactions

- van der Waals
- Conventional Hydrogen Bond

- Pi-Pi T-shaped
- Pi-Alkyl

4

## Kaempferol



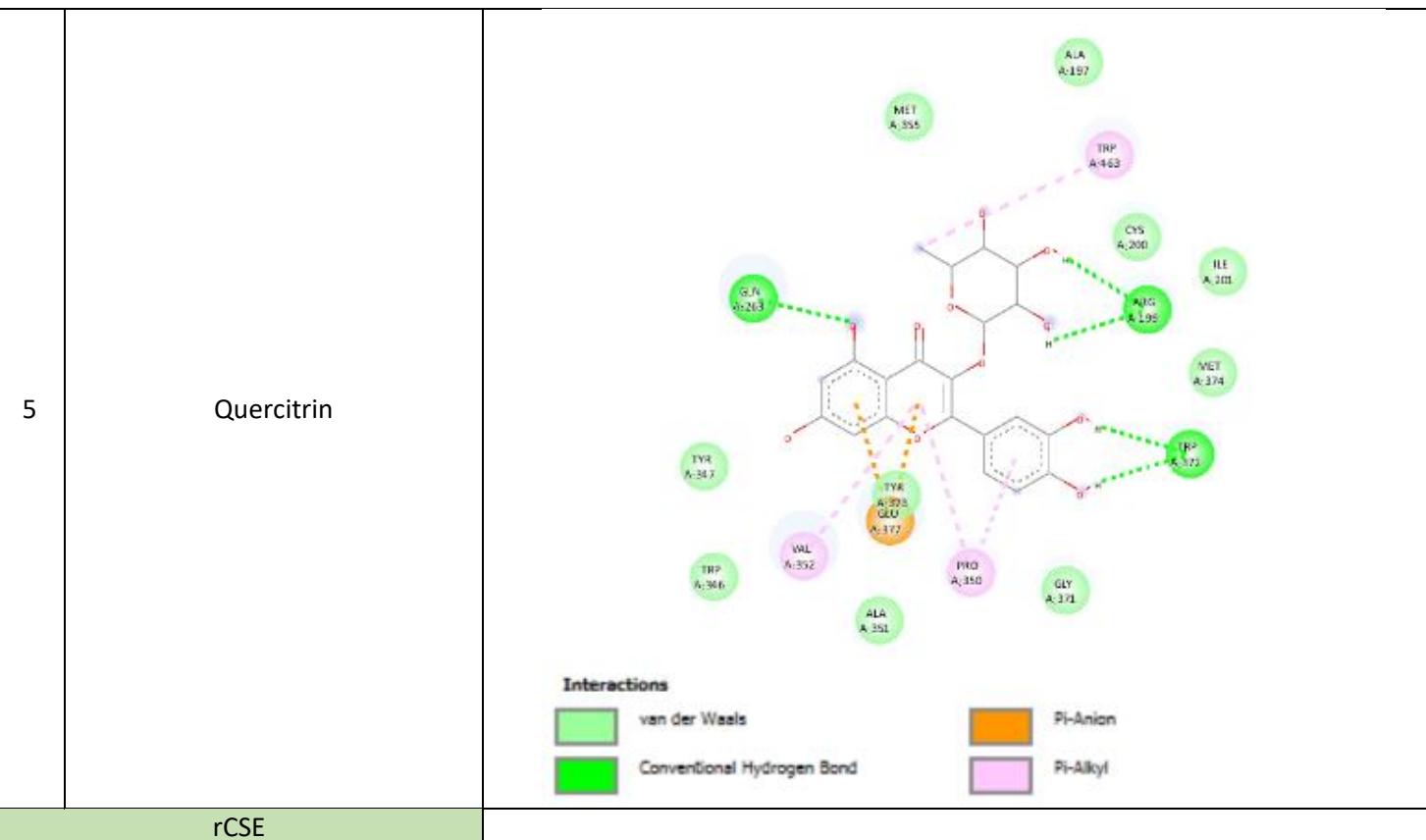
## Interactions



van der Waals

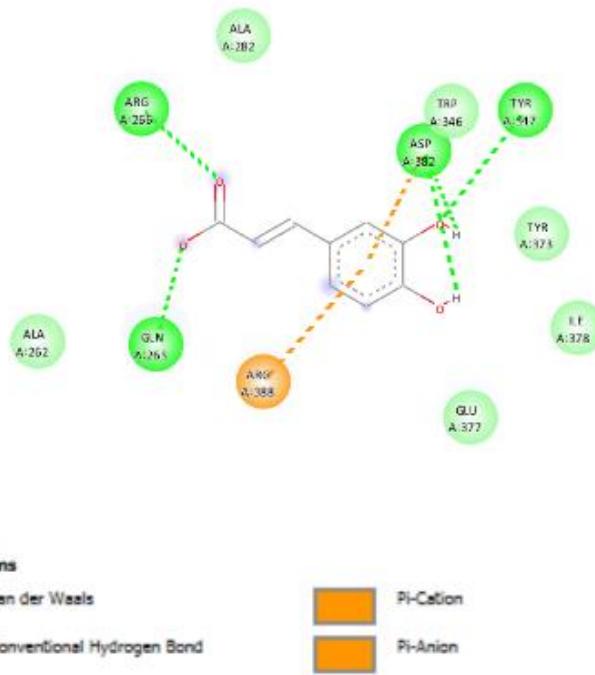
Conventional Hydrogen Bond  
Pi-Alkyl

Pi-Pi T-shaped



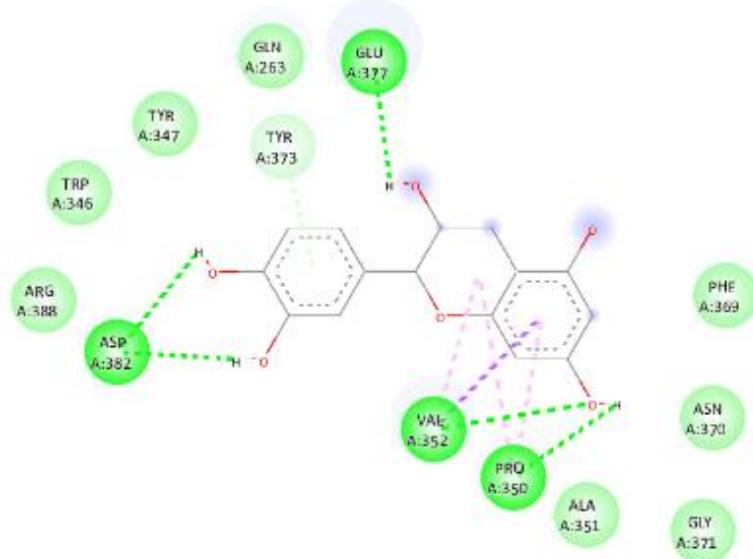
1

Caffeic acid



2

## (+)-Catechin

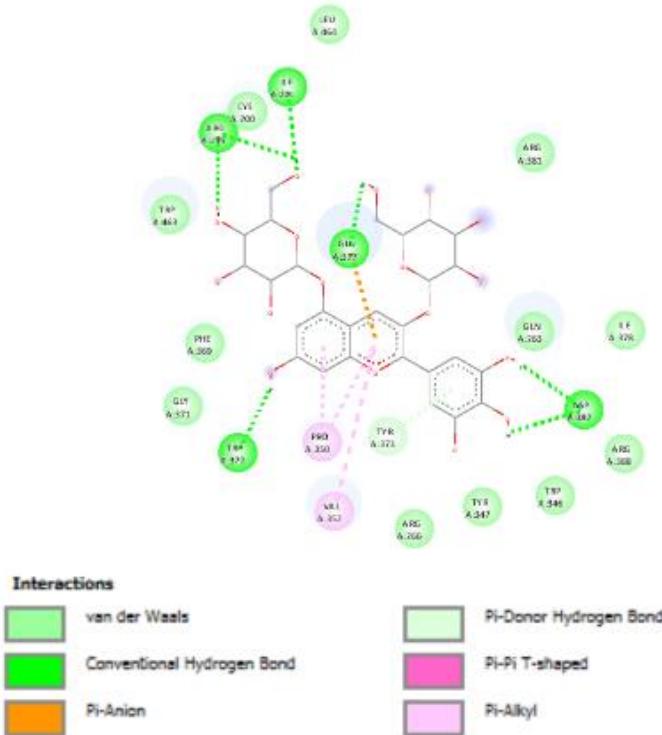


## Interactions

|  |                            |
|--|----------------------------|
|  | van der Waals              |
|  | Conventional Hydrogen Bond |
|  | Pi-Donor Hydrogen Bond     |
|  | Pi-Sigma                   |
|  | Pi-Pi T-shaped             |
|  | Alkyl                      |
|  | Pi-Alkyl                   |

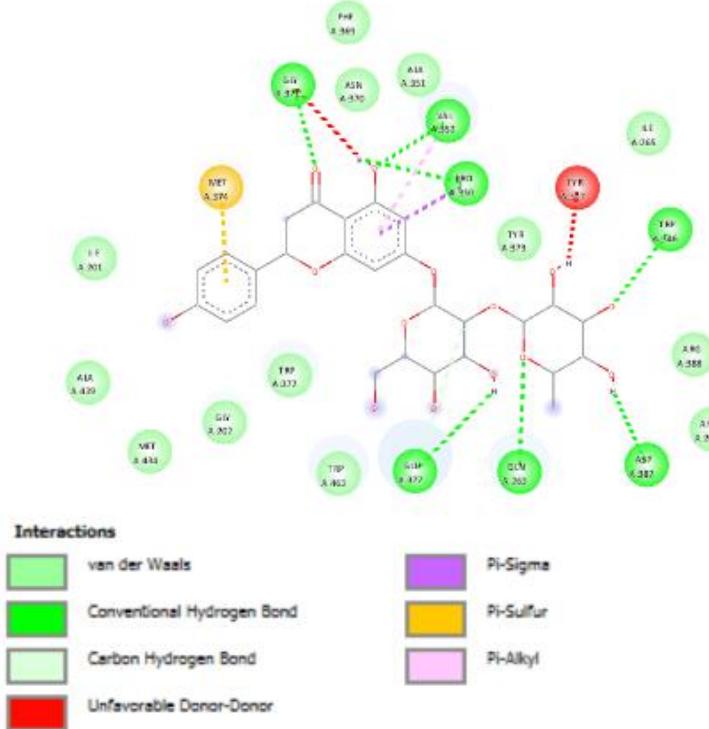
3

(2R,3S,4S,5R,6S)-2-(hydroxymethyl)-6-[7-hydroxy-3-[(2S,3R,4S,5S,6R)-3,4,5-trihydroxy-6-(hydroxymethyl)oxan-2-yl]oxy-2-(3,4,5-trihydroxyphenyl)chromenylium-5-yl]oxyoxane-3,4,5-triol



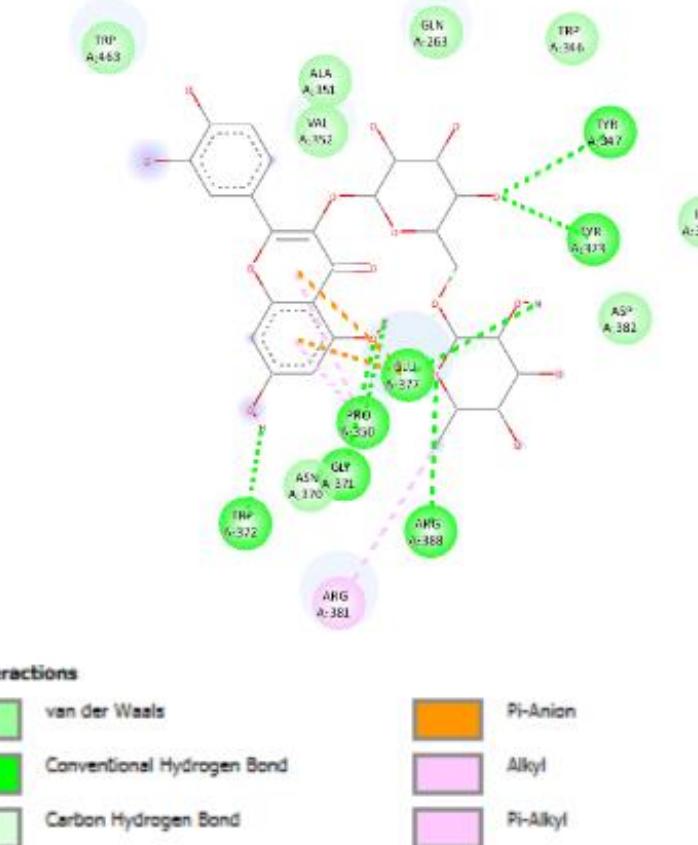
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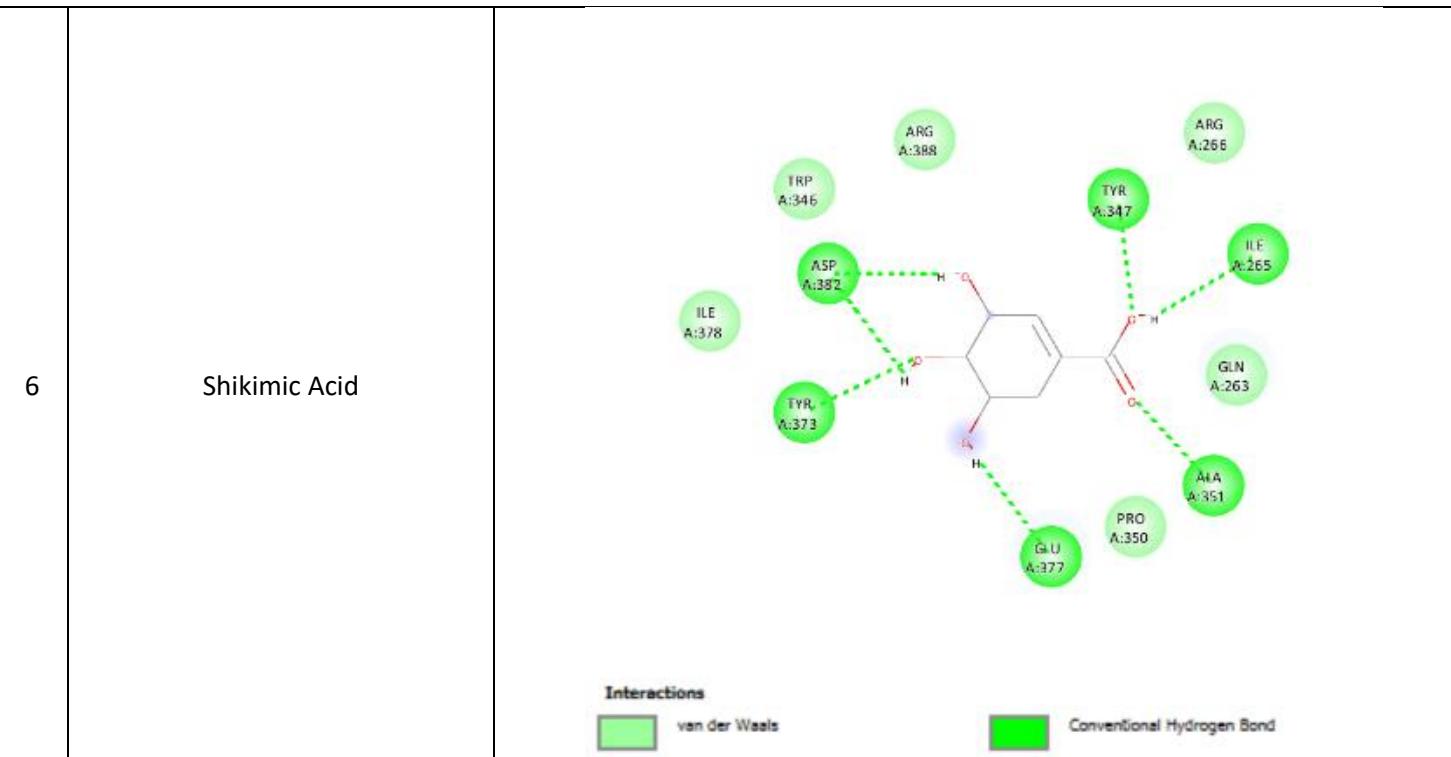
## Naringin



5

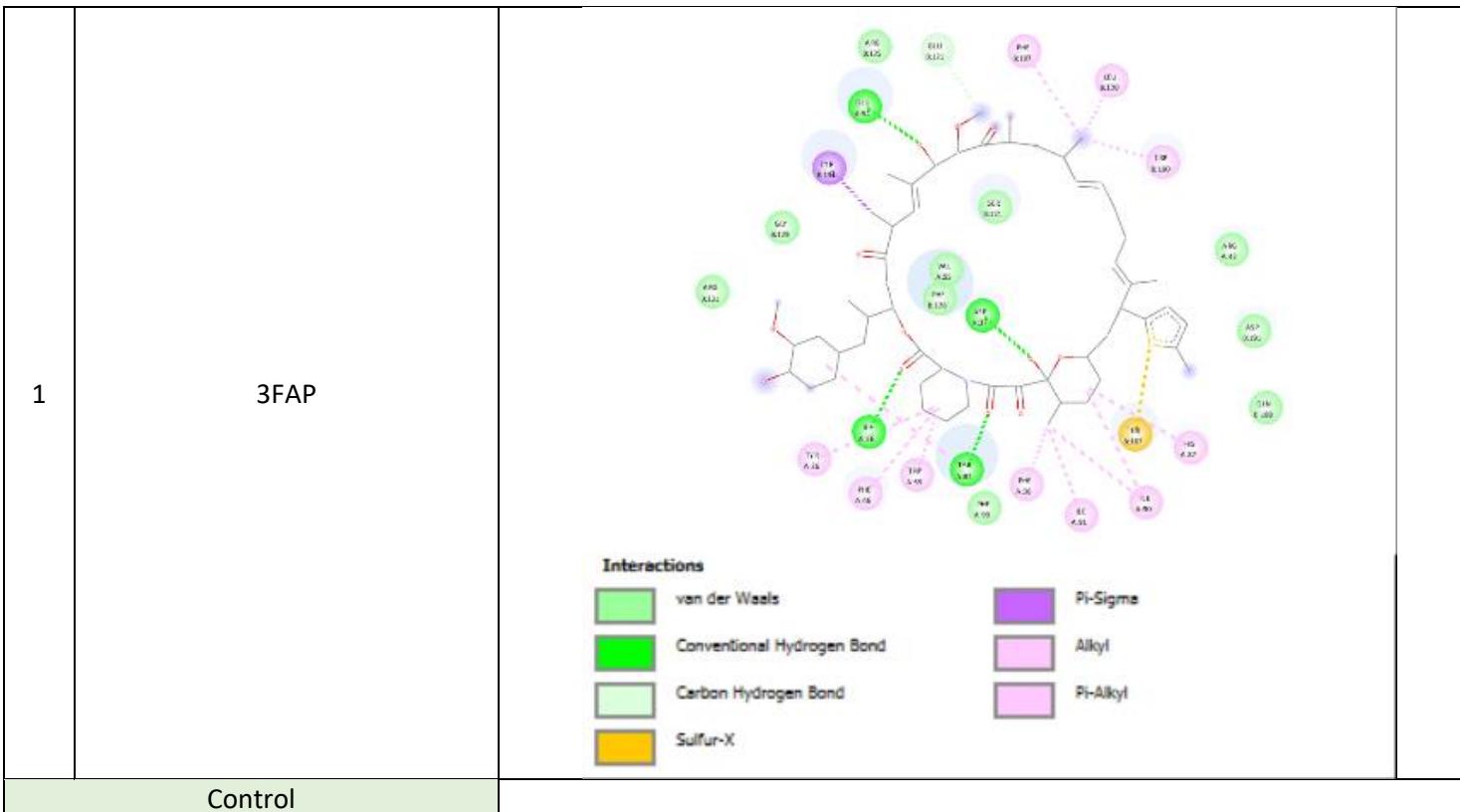
## Rutin

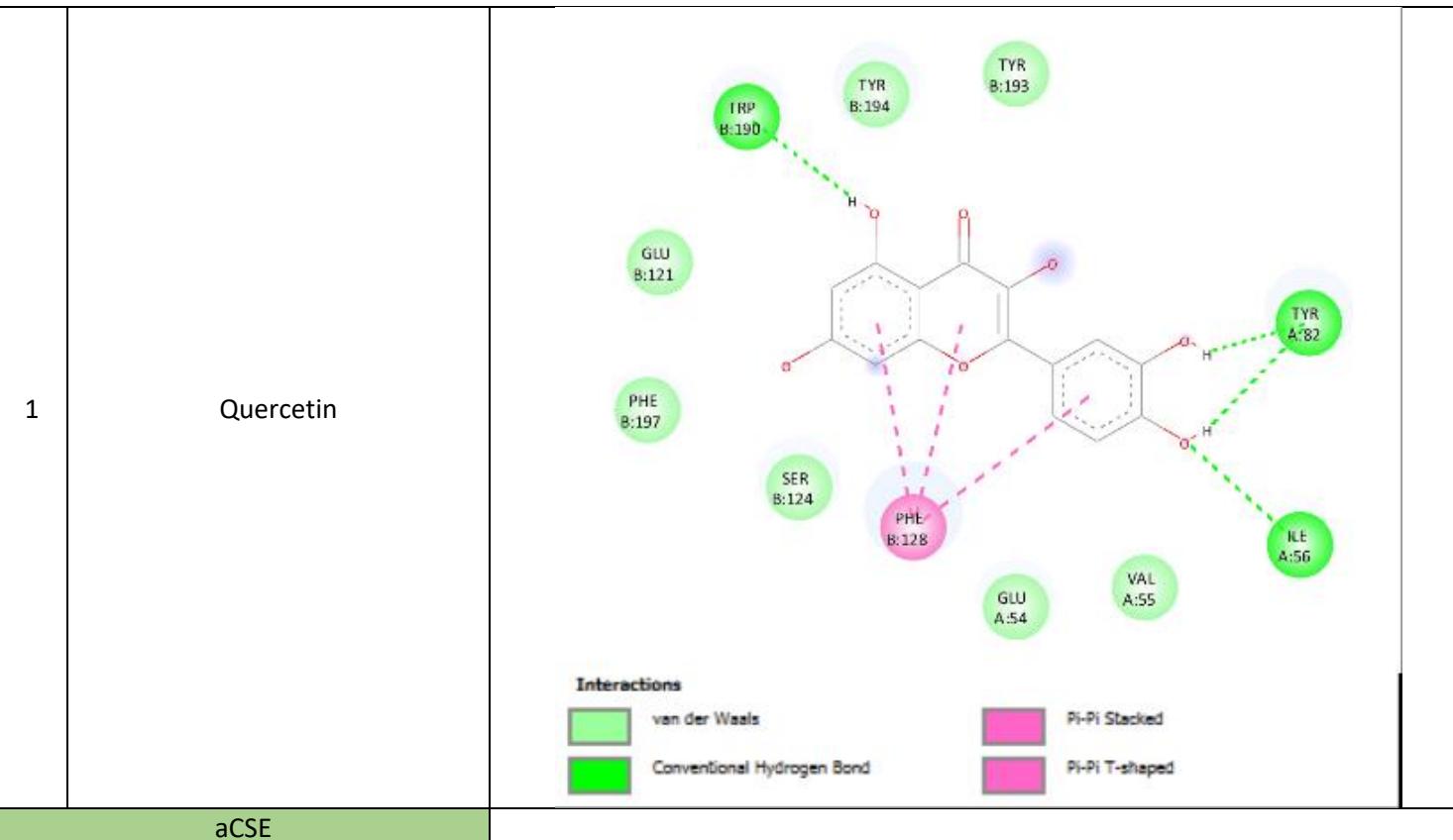




## mTOR

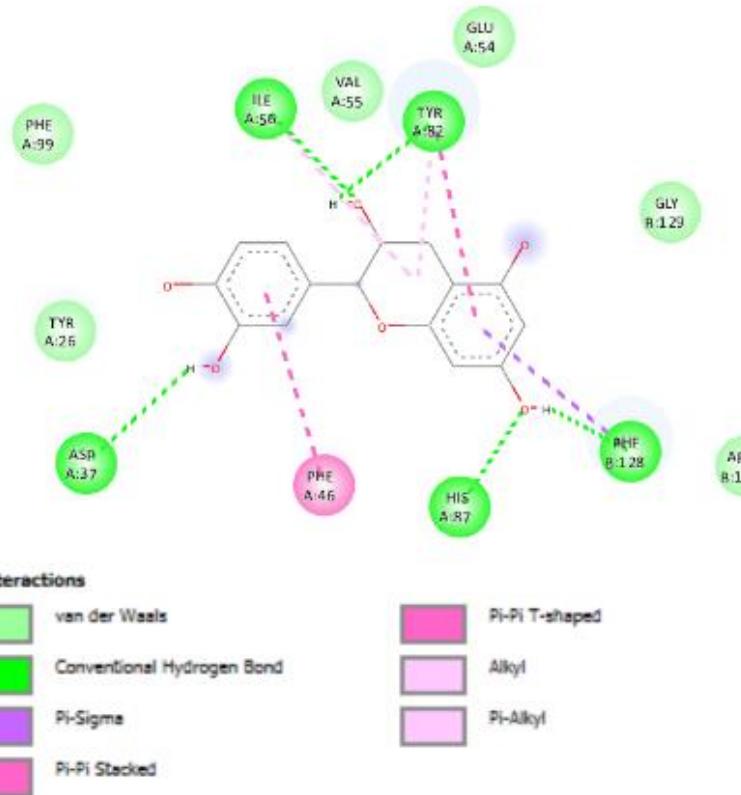
| No. | Substance     | Visualization |
|-----|---------------|---------------|
|     | Native Ligand |               |





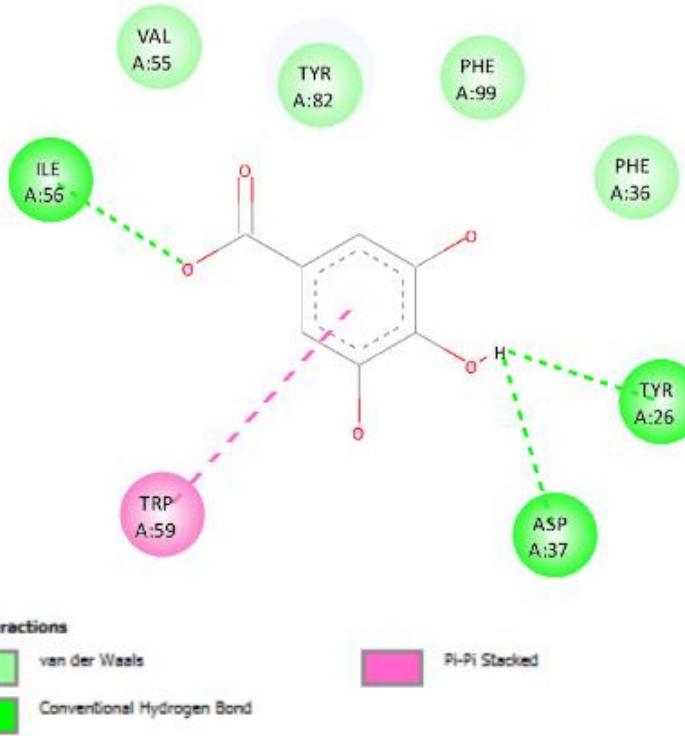
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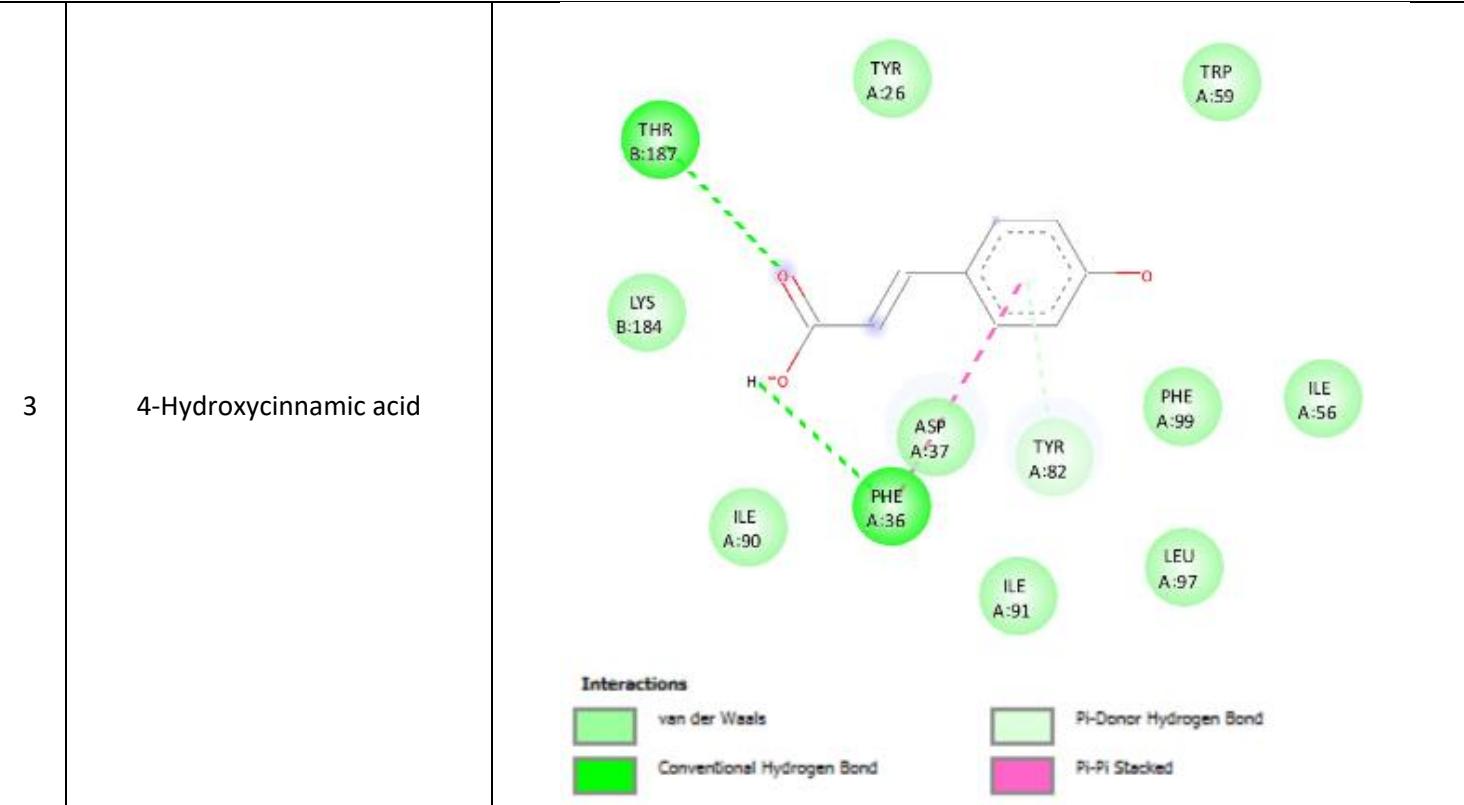
## Epicatechin



2

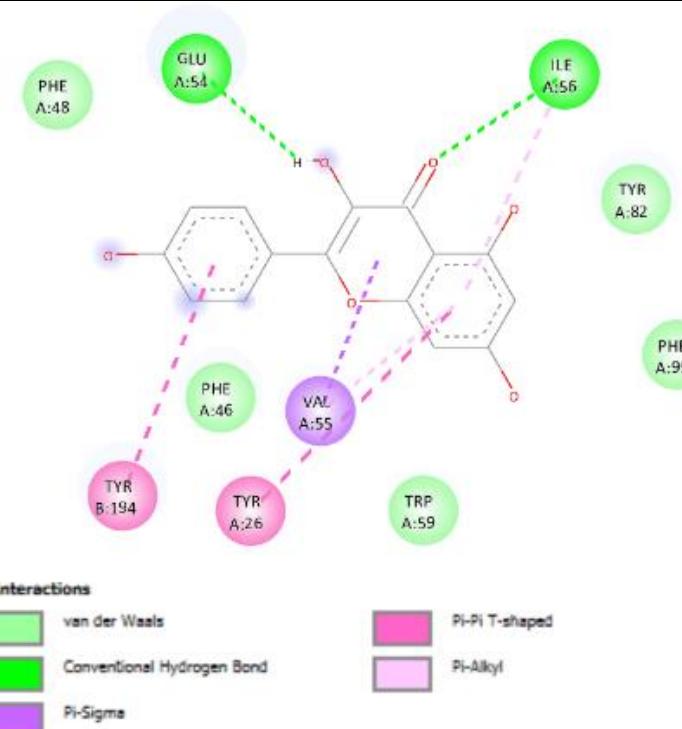
Gallic acid

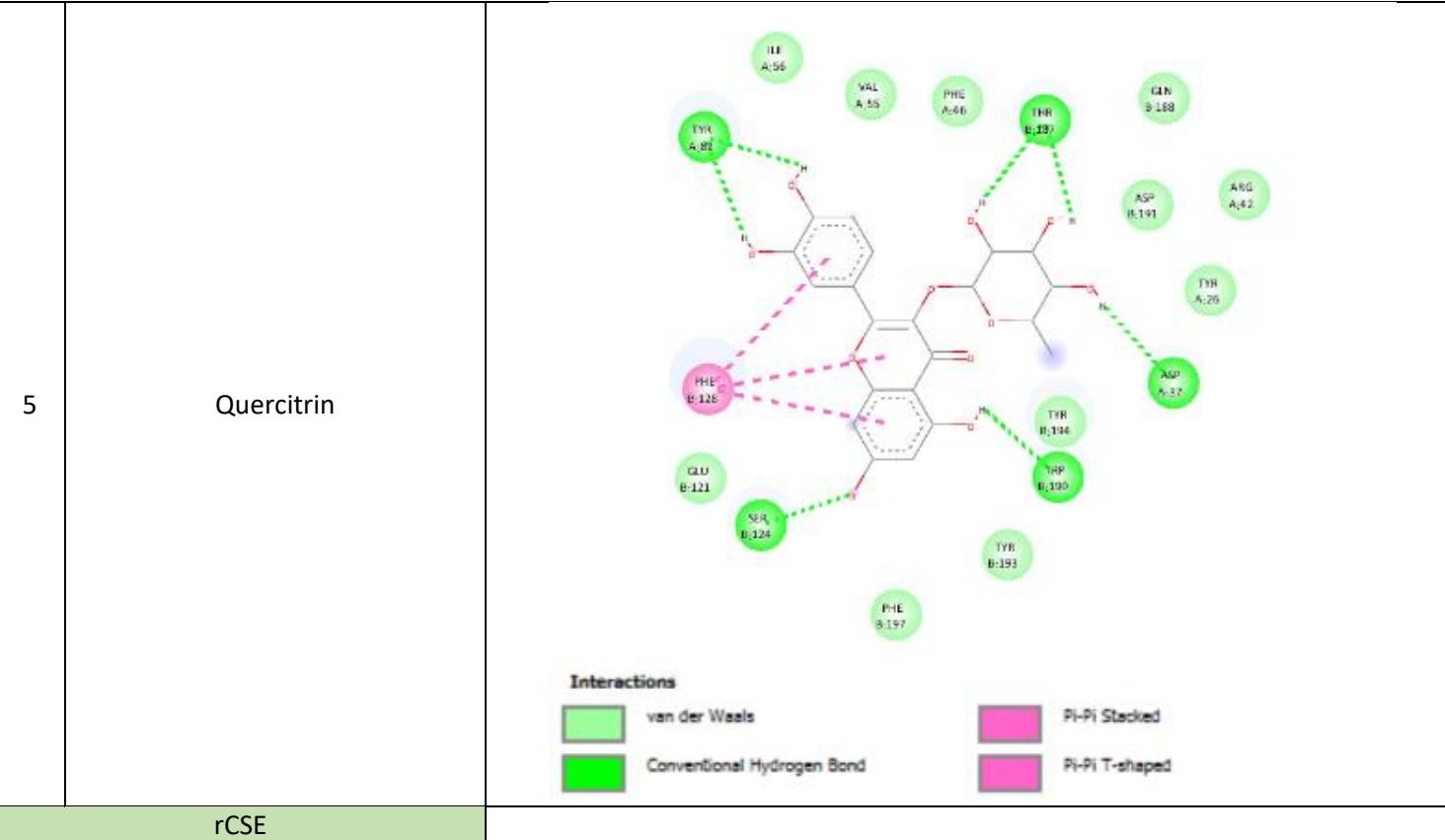




4

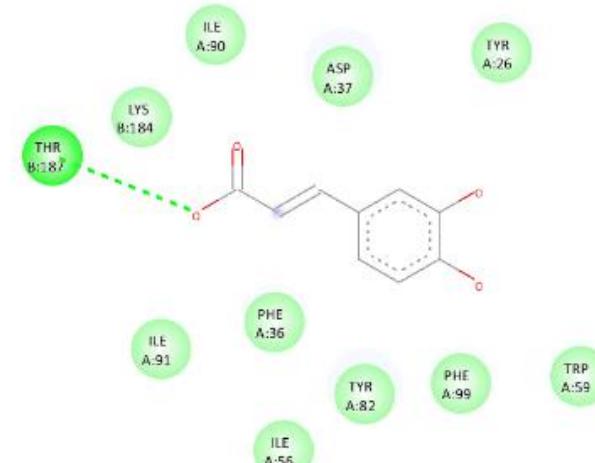
## Kaempferol





1

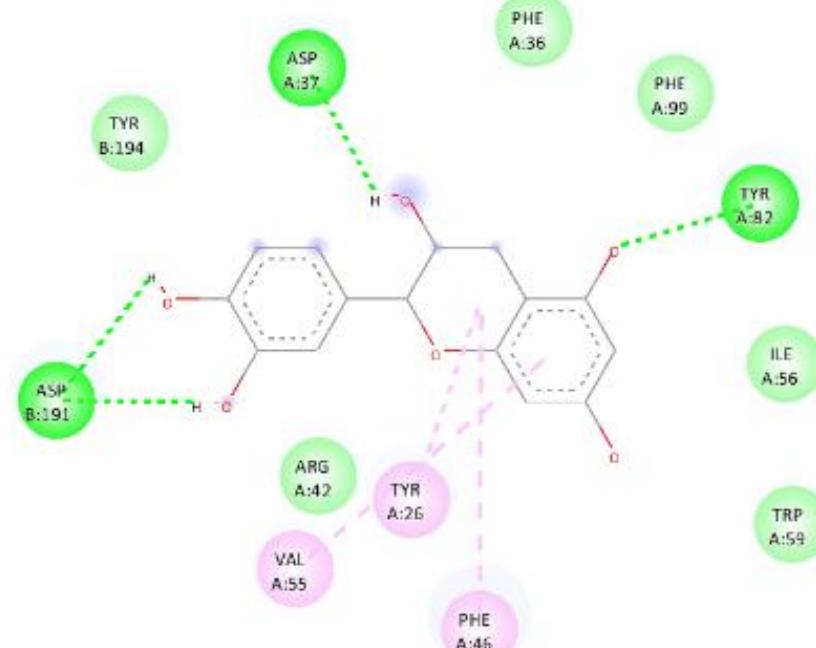
Caffeic acid



Interactions  
van der Waals      Conventional Hydrogen Bond

2

## (+)-Catechin

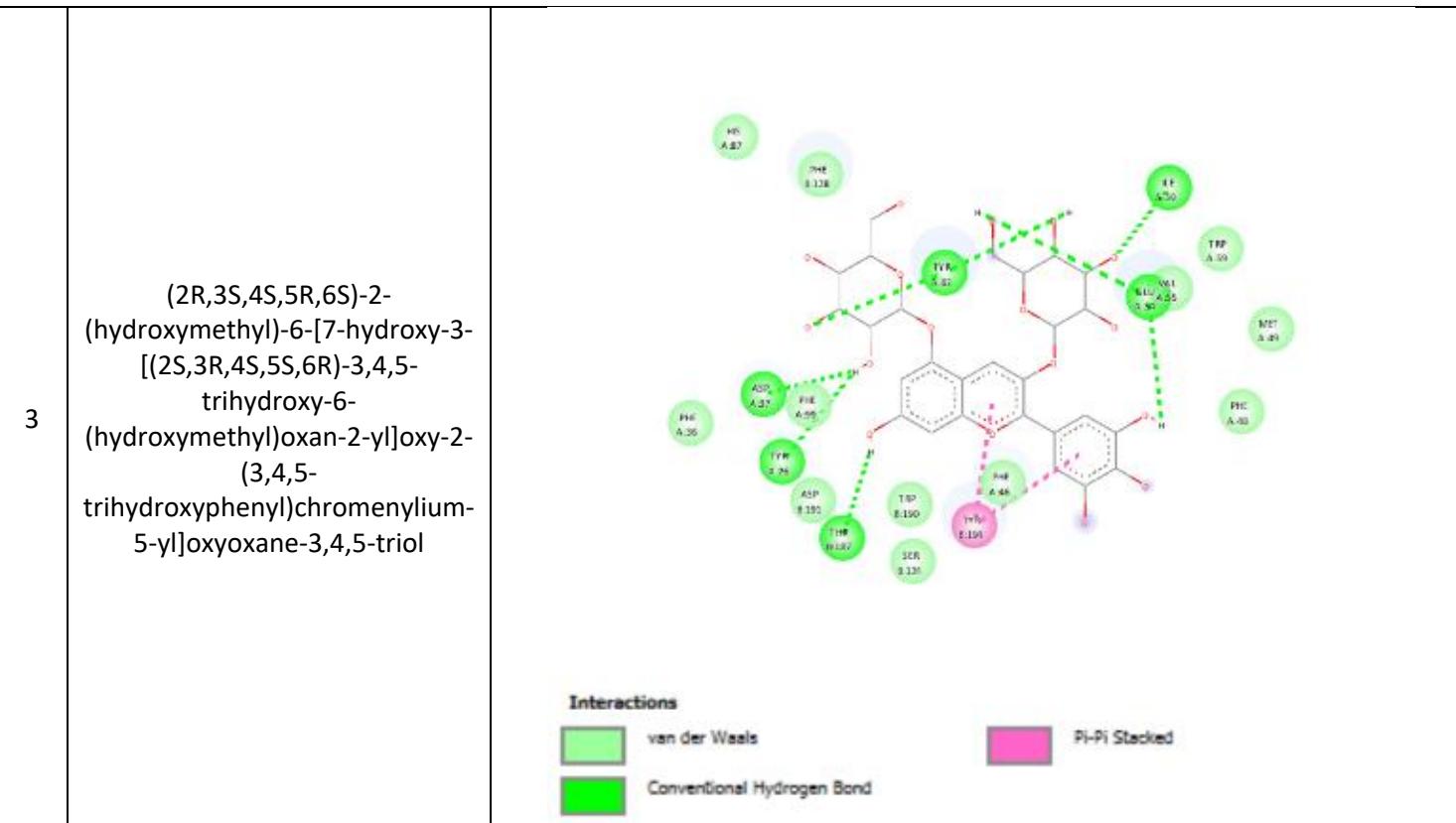


## Interactions

van der Waals

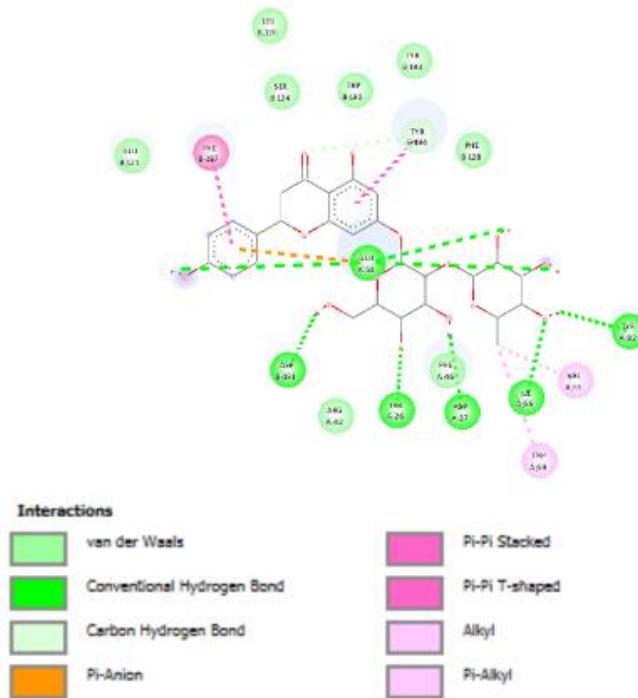
Conventional Hydrogen Bond

Pi-Alkyl



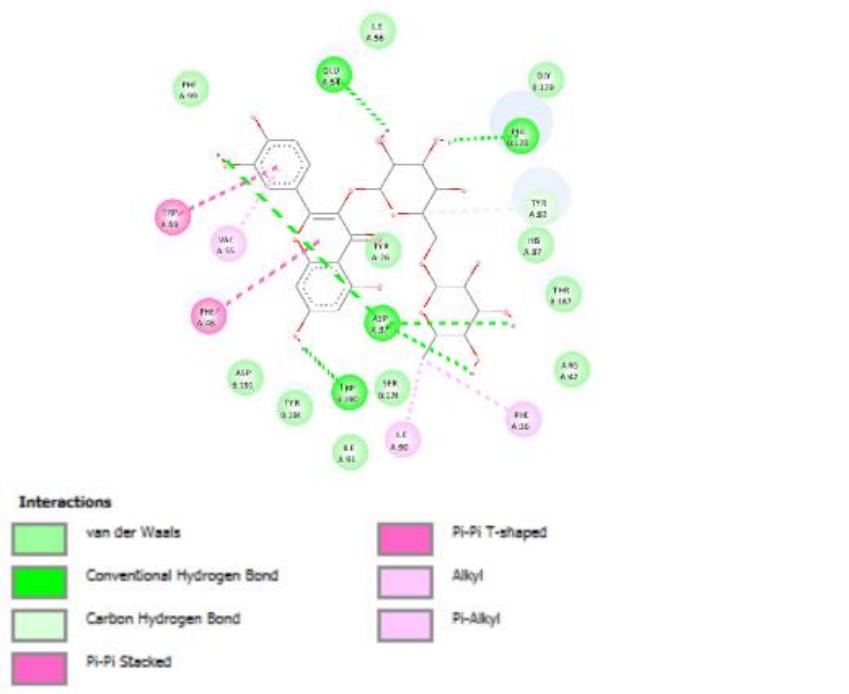
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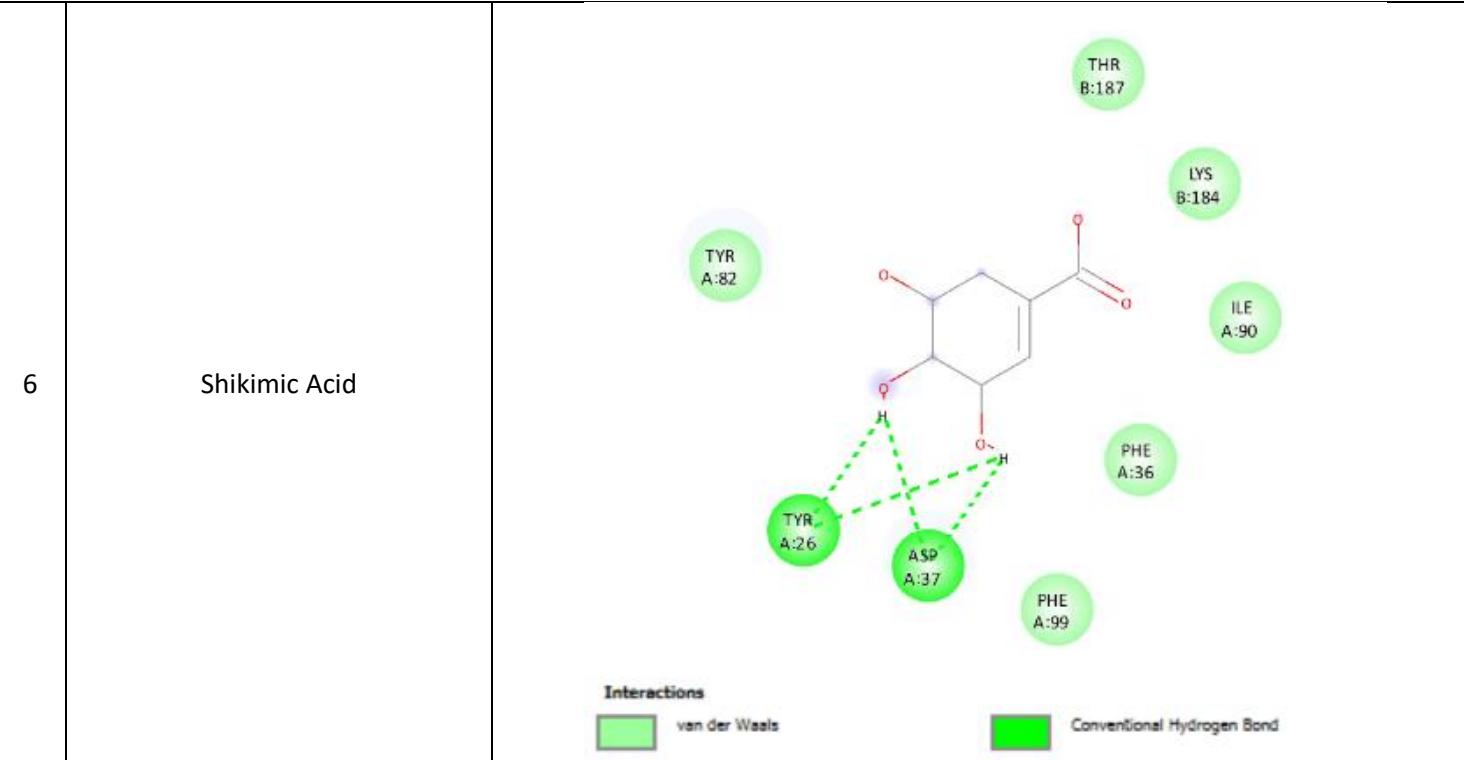
## Naringin



5

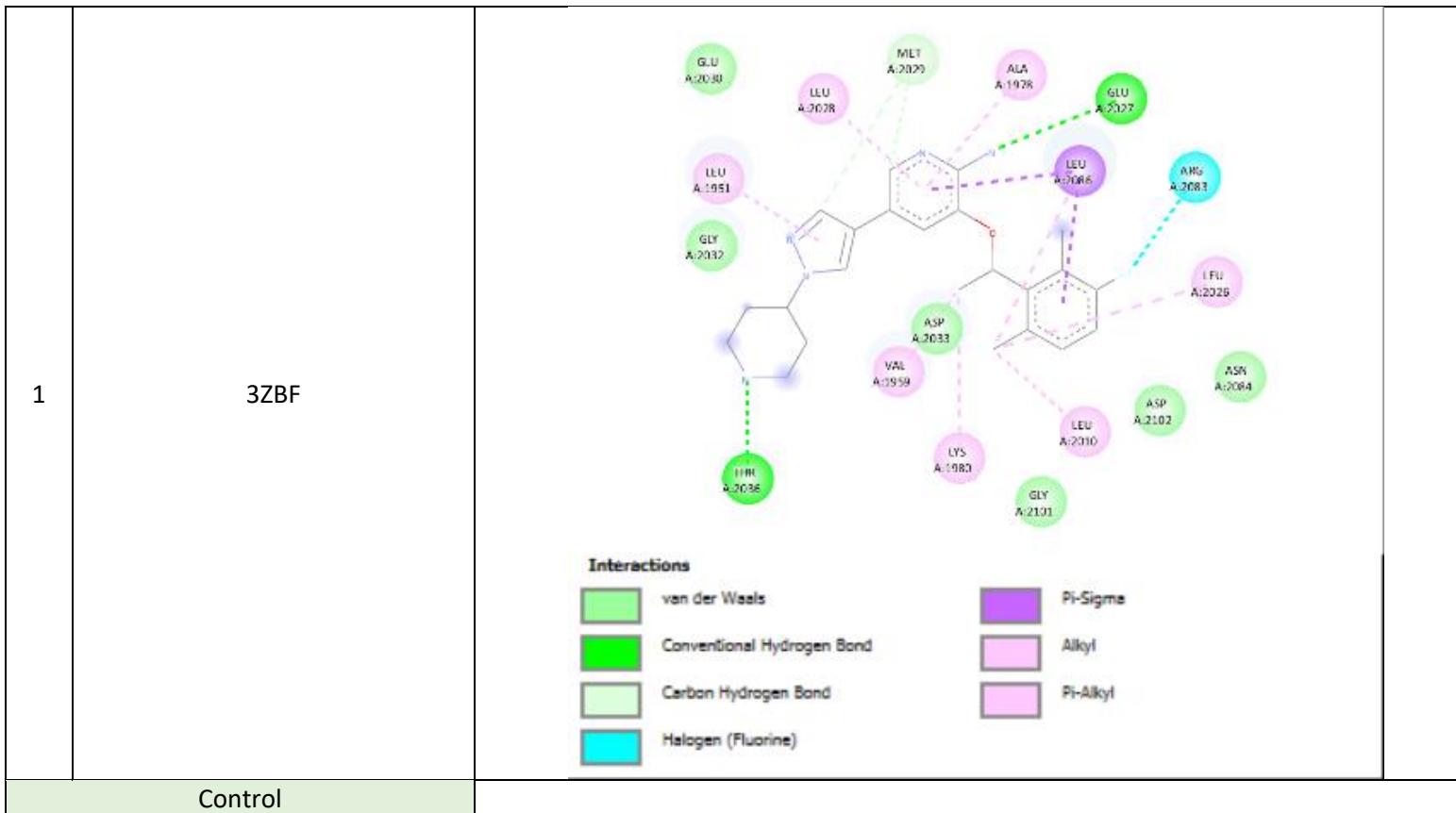
## Rutin

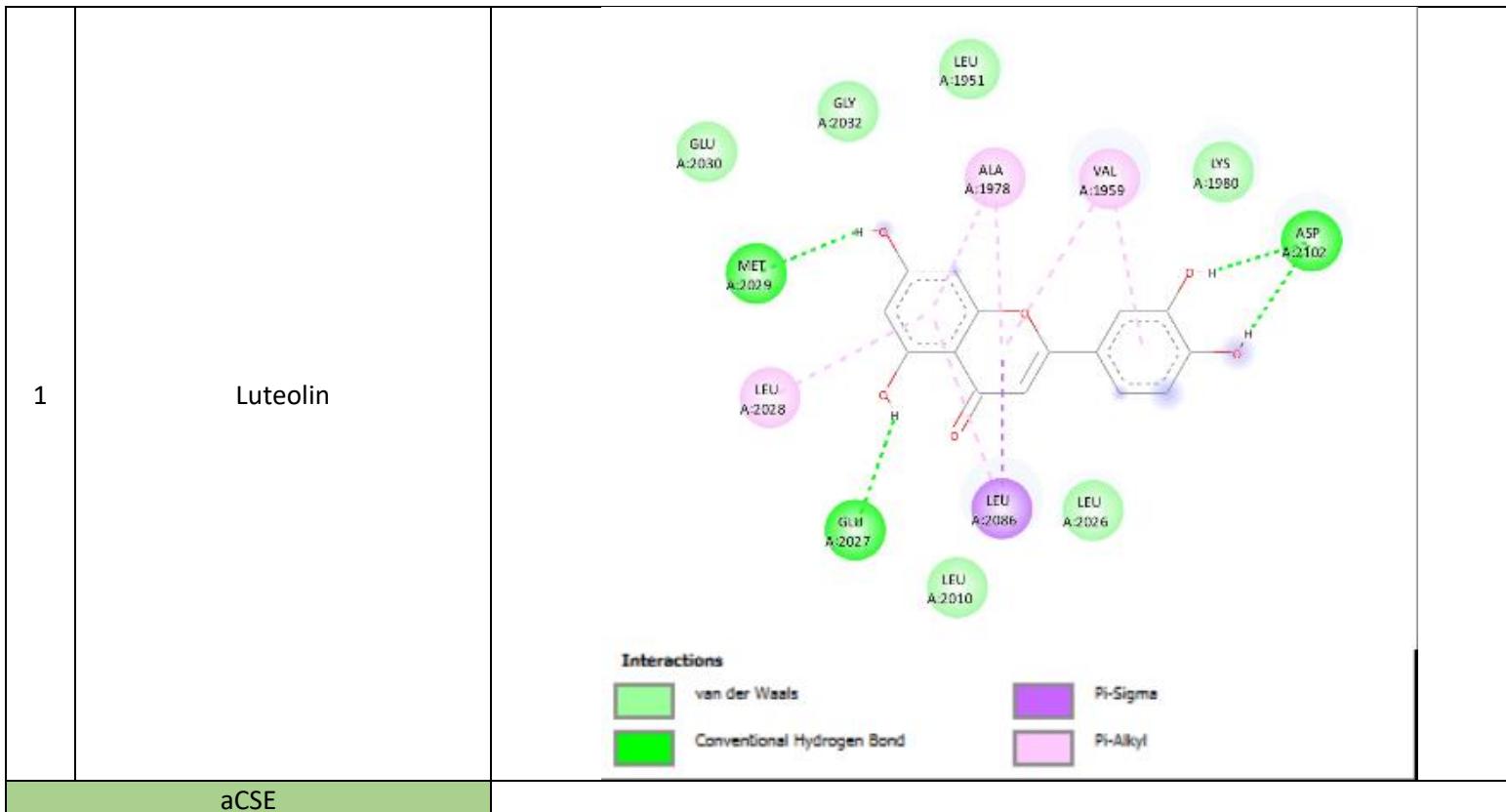




## ROS1 Kinase

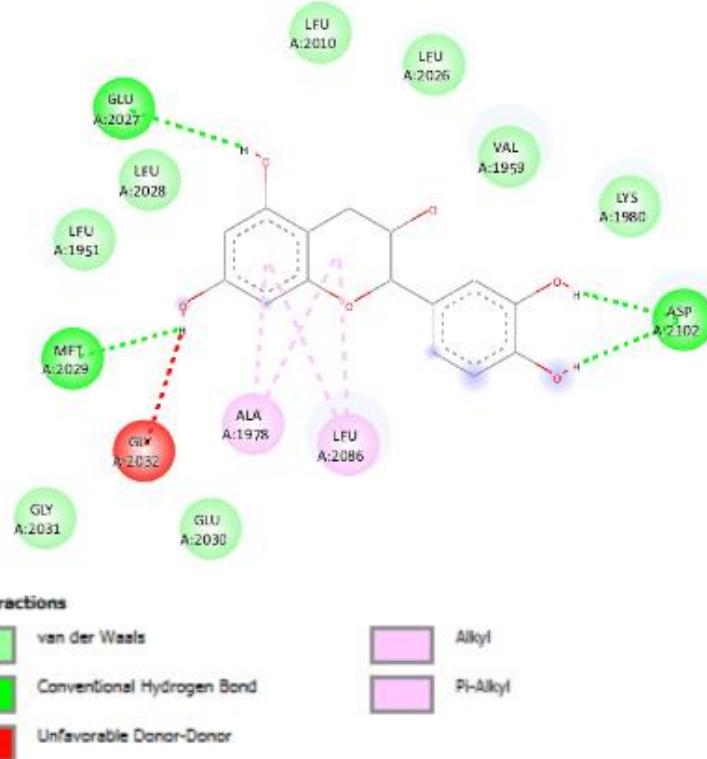
| No. | Substance     | Visualization |
|-----|---------------|---------------|
|     | Native Ligand |               |





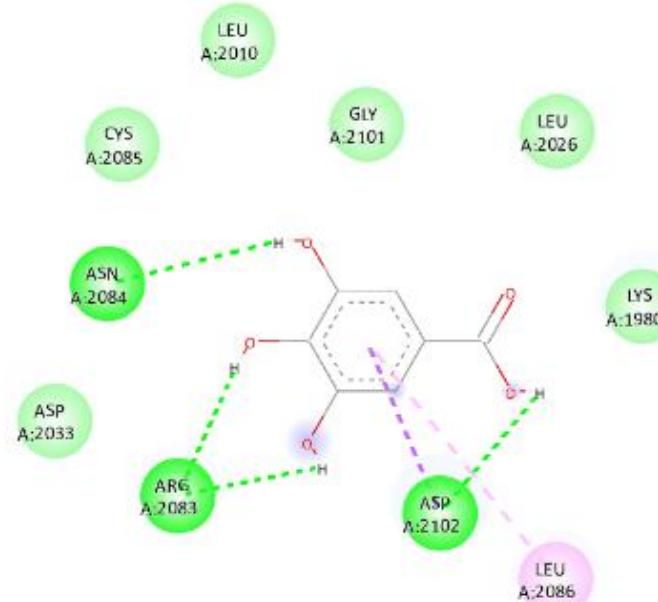
1

## Epicatechin



2

Gallic acid

**Interactions**

van der Waals

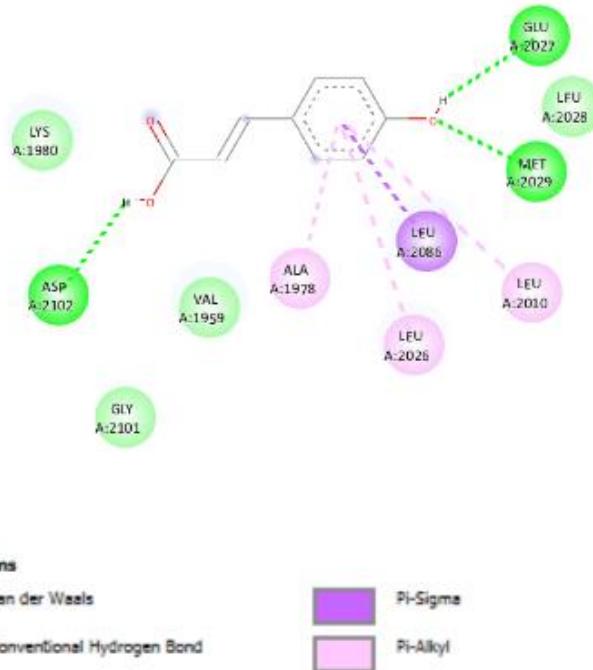
Conventional Hydrogen Bond

Pi-Sigma

Pi-Alkyl

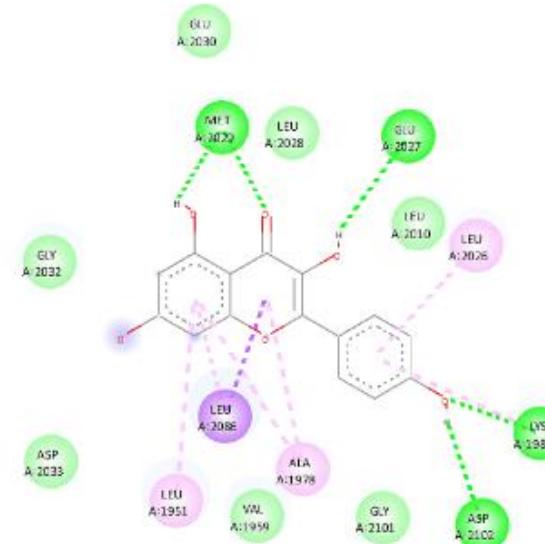
3

## 4-Hydroxycinnamic acid



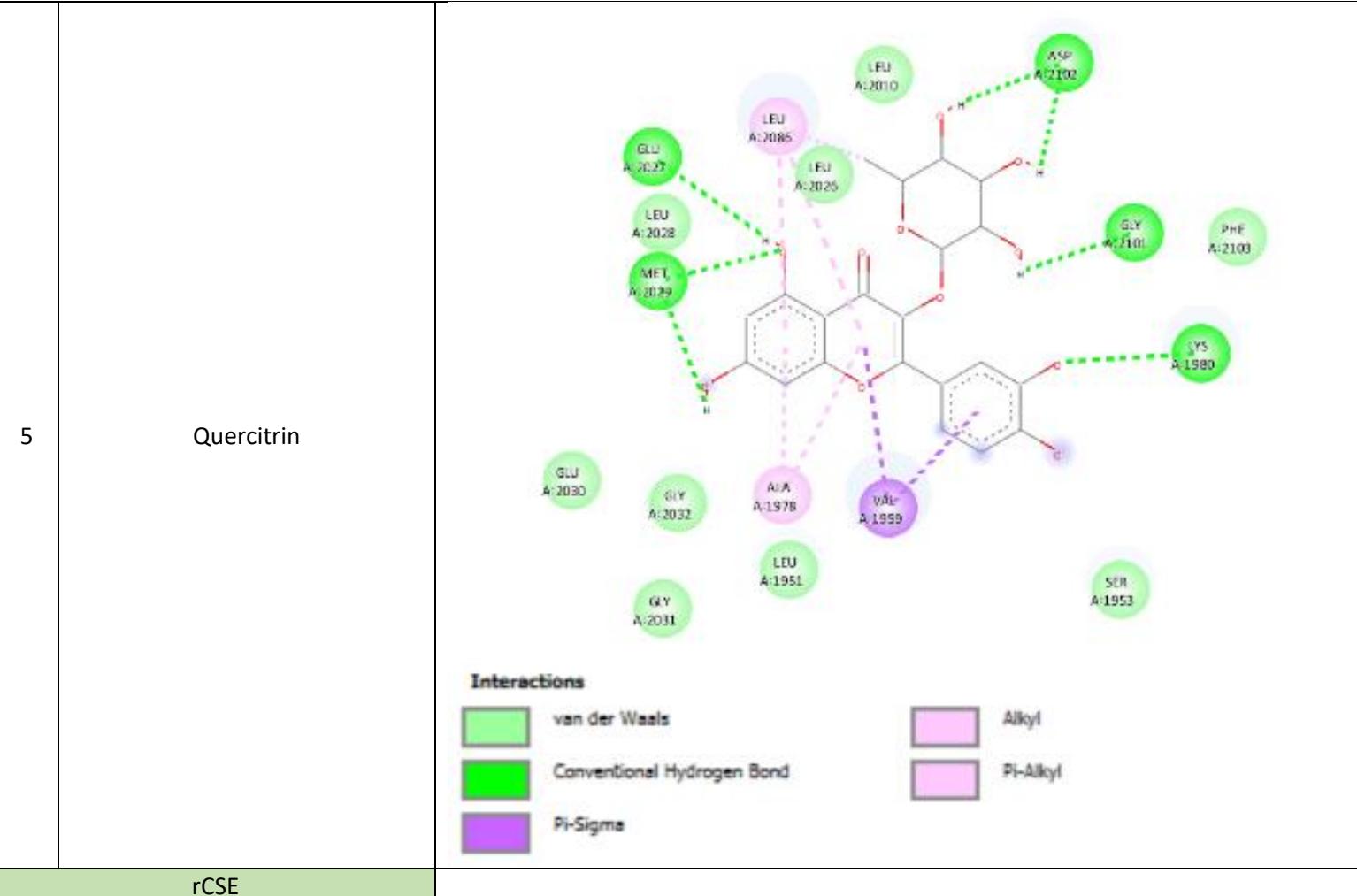
4

## Kaempferol



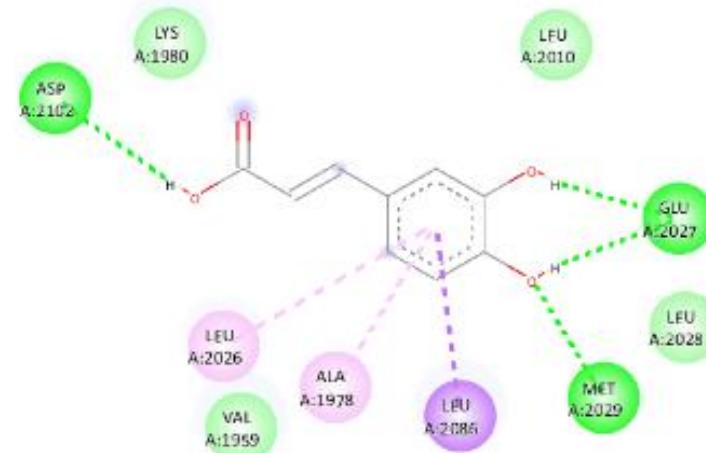
## Interactions:

- |                            |          |
|----------------------------|----------|
| van der Waals              | Pi-Sigma |
| Conventional Hydrogen Bond | Pi-Alkyl |
| Unfavorable Donor-Donor    |          |



1

Caffeic acid

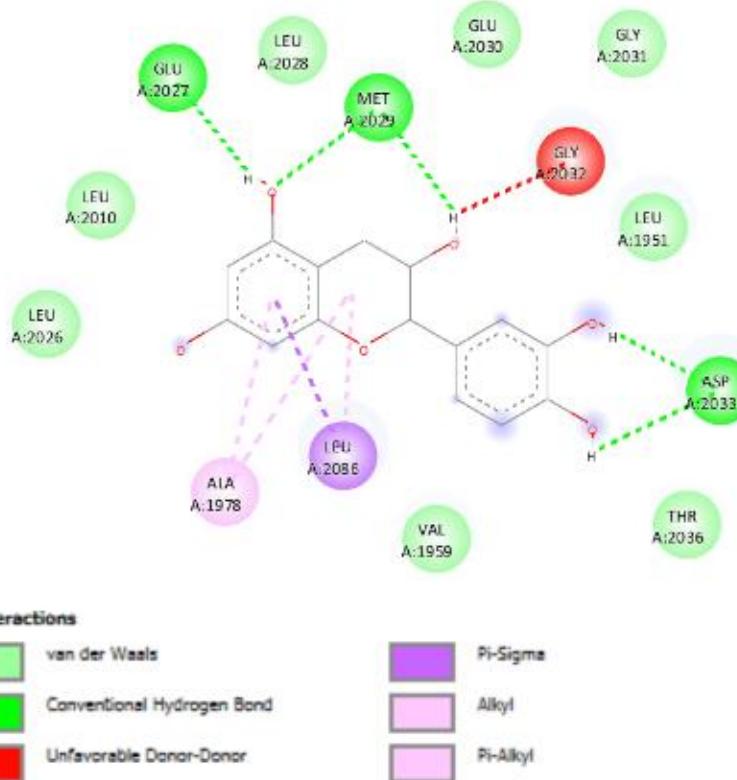
**Interactions**

- van der Waals
- Conventional Hydrogen Bond

- Pi-Sigma
- Pi-Alkyl

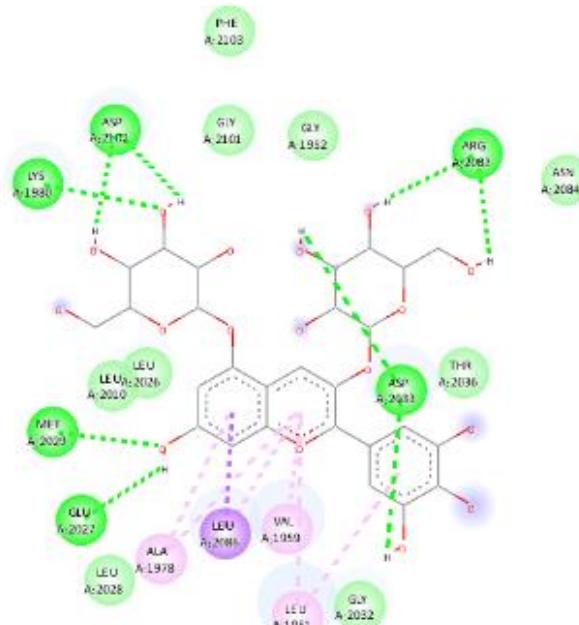
2

## (+)-Catechin



3

(2R,3S,4S,5R,6S)-2-(hydroxymethyl)-6-[7-hydroxy-3-[(2S,3R,4S,5S,6R)-3,4,5-trihydroxy-6-(hydroxymethyl)oxan-2-yl]oxy-2-(3,4,5-trihydroxyphenyl)chromenylium-5-yl]oxyoxane-3,4,5-triol



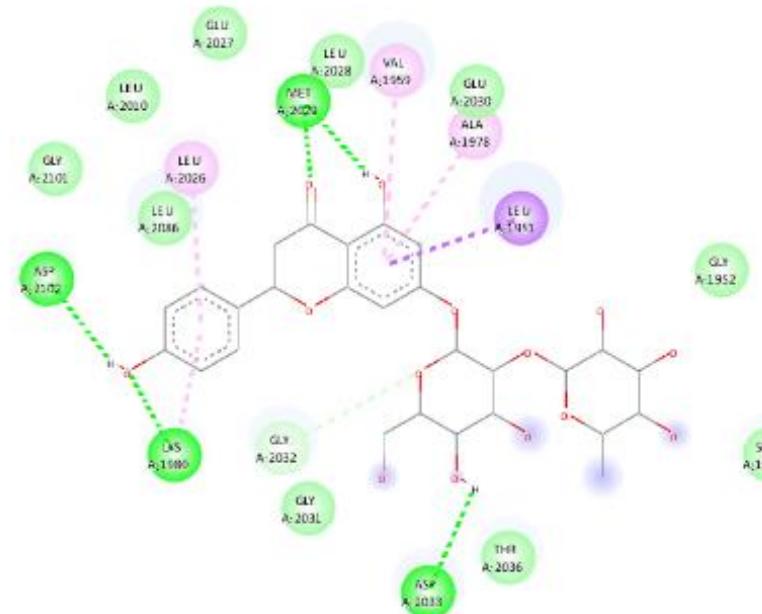
#### Interactions

- van der Waals
- Conventional Hydrogen Bond

- Pi-Sigma
- Pi-Alkyl

4

## Naringin

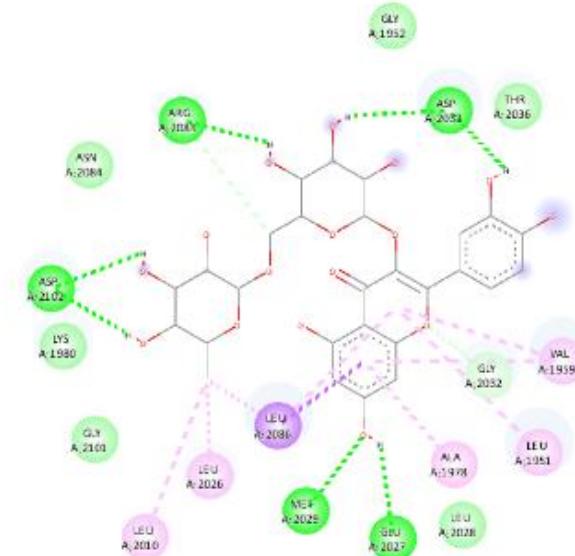


## Interactions

|  |                            |
|--|----------------------------|
|  | van der Waals              |
|  | Conventional Hydrogen Bond |
|  | Carbon Hydrogen Bond       |
|  | Unfavorable Donor-Donor    |
|  | Pi-Sigma                   |
|  | Pi-Alkyl                   |

5

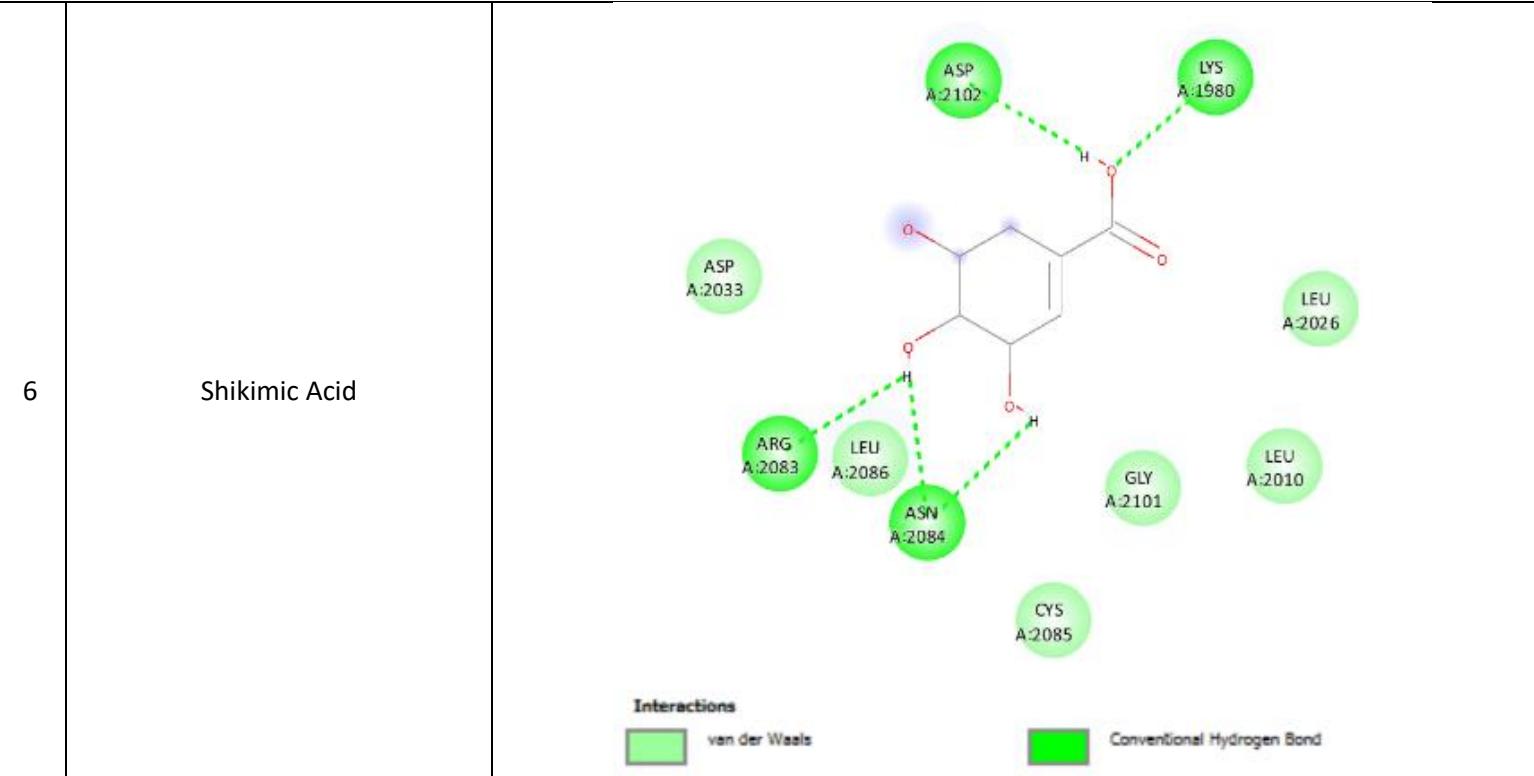
## Rutin



## Interactions

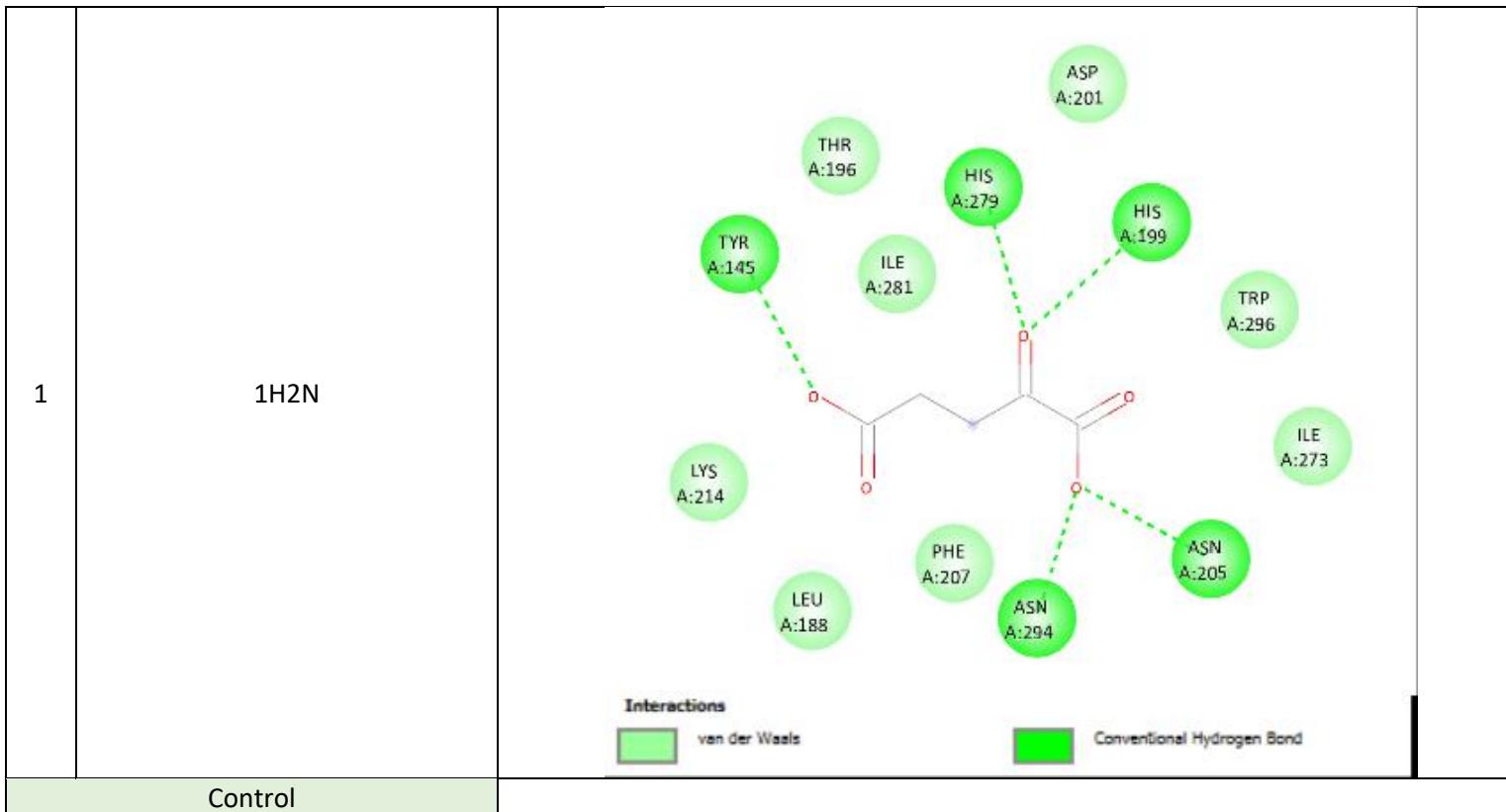
- van der Waals
- Conventional Hydrogen Bond
- Carbon Hydrogen Bond

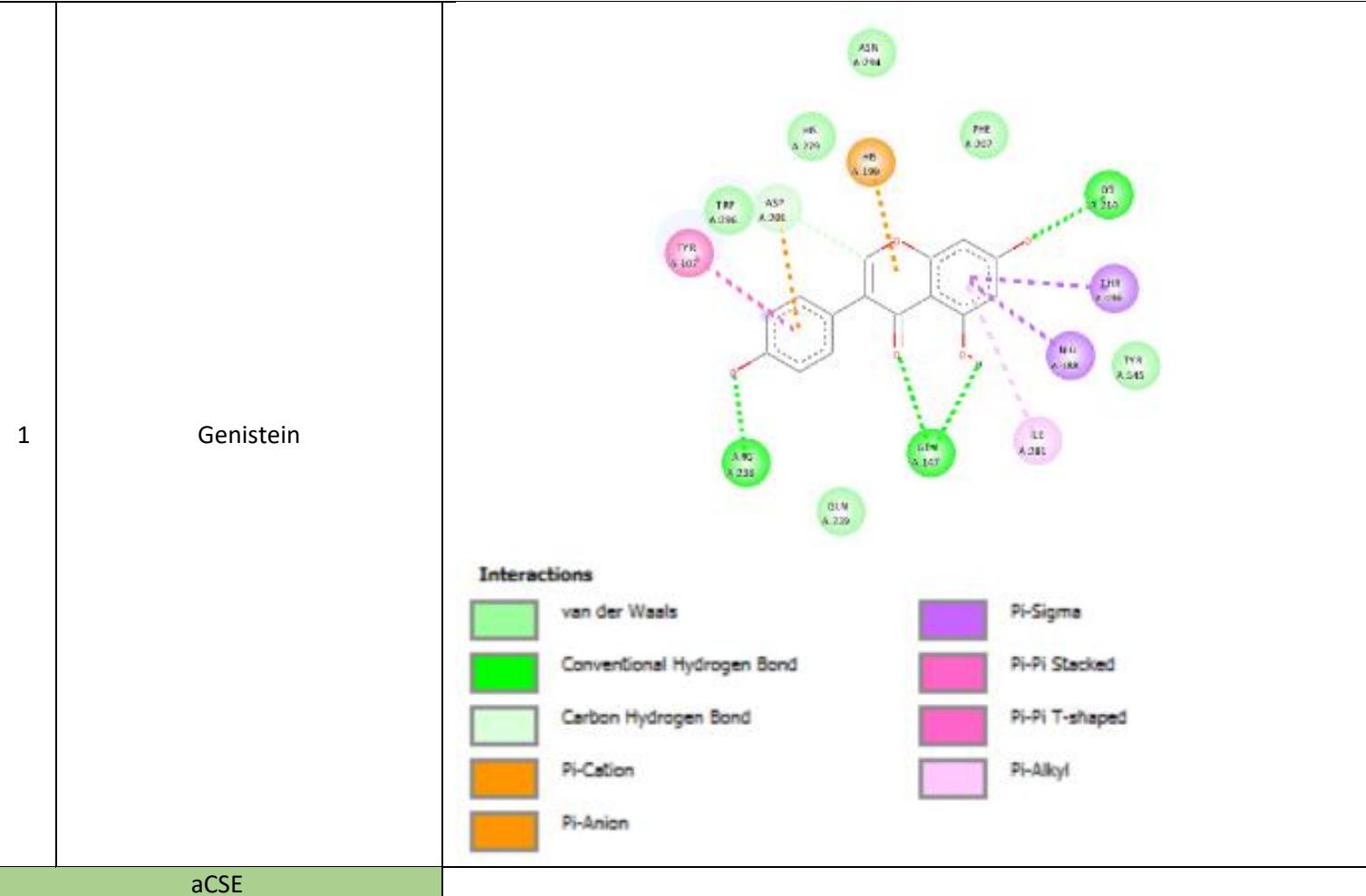
- Pi-Sigma
- Alkyl
- Pi-Alkyl



## HIF-1alpha

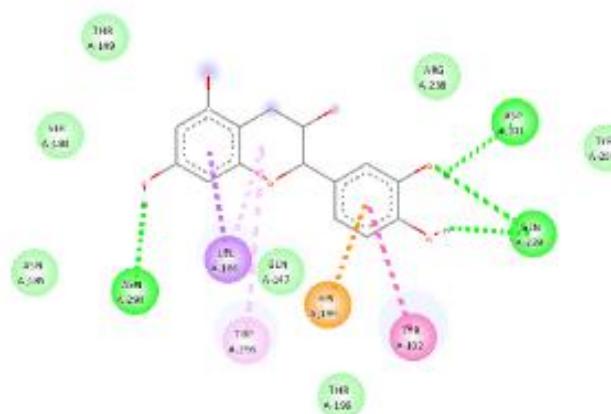
| No. | Substance     | Visualization |
|-----|---------------|---------------|
|     | Native Ligand |               |





1

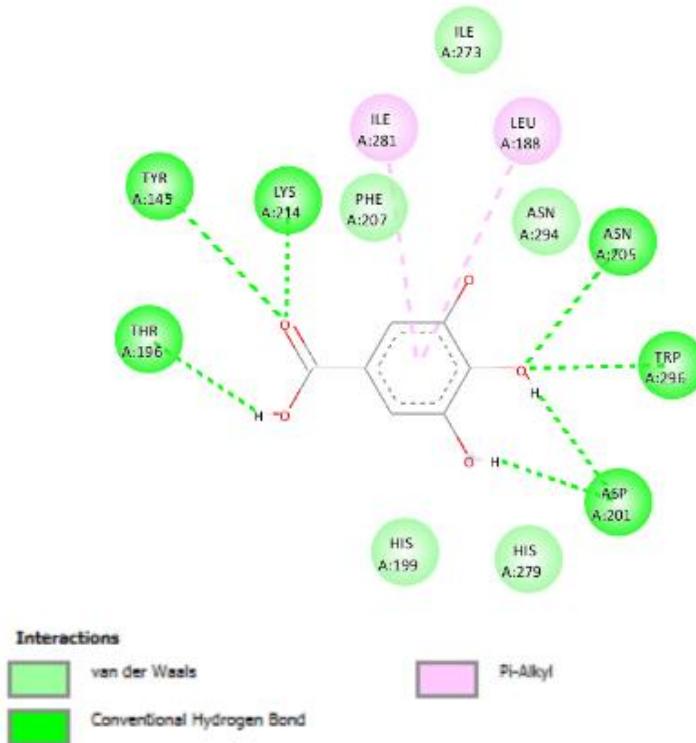
Epicatechin

**Interactions**

|  |                            |  |               |
|--|----------------------------|--|---------------|
|  | van der Waals              |  | Pi-Pi Stacked |
|  | Conventional Hydrogen Bond |  | Alkyl         |
|  | Pi-Cation                  |  | Pi-Alkyl      |
|  | Pi-Sigma                   |  |               |

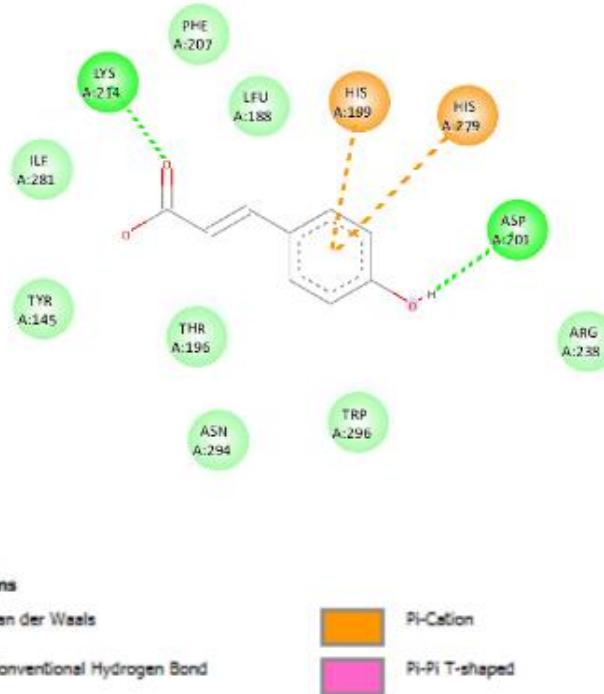
2

Gallic acid



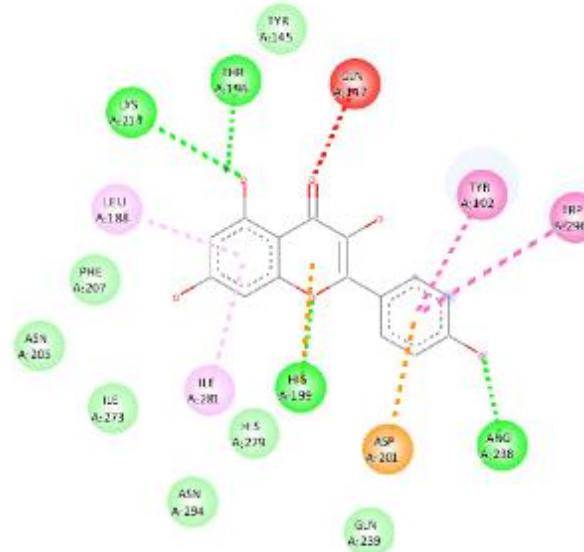
3

## 4-Hydroxycinnamic acid



4

## Kaempferol



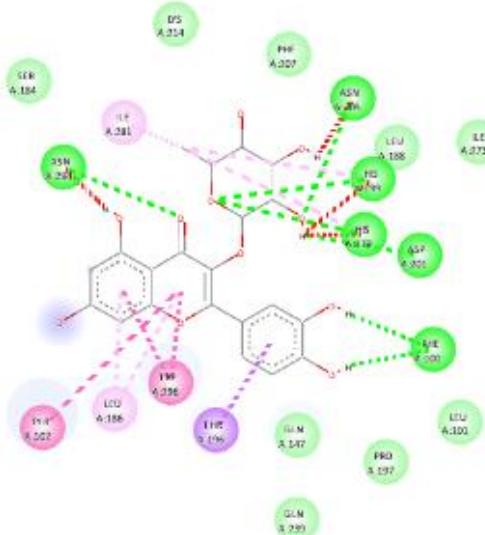
## Interactions

|  |                               |  |                |
|--|-------------------------------|--|----------------|
|  | van der Waals                 |  | Pi-Anion       |
|  | Conventional Hydrogen Bond    |  | Pi-Pi Stacked  |
|  | Unfavorable Acceptor-Acceptor |  | Pi-Pi T-shaped |
|  | Pi-Cation                     |  | Pi-Alkyl       |

5

Quercitrin

rCSE

**Interactions**

van der Waals

Conventional Hydrogen Bond

Unfavorable Donor-Donor

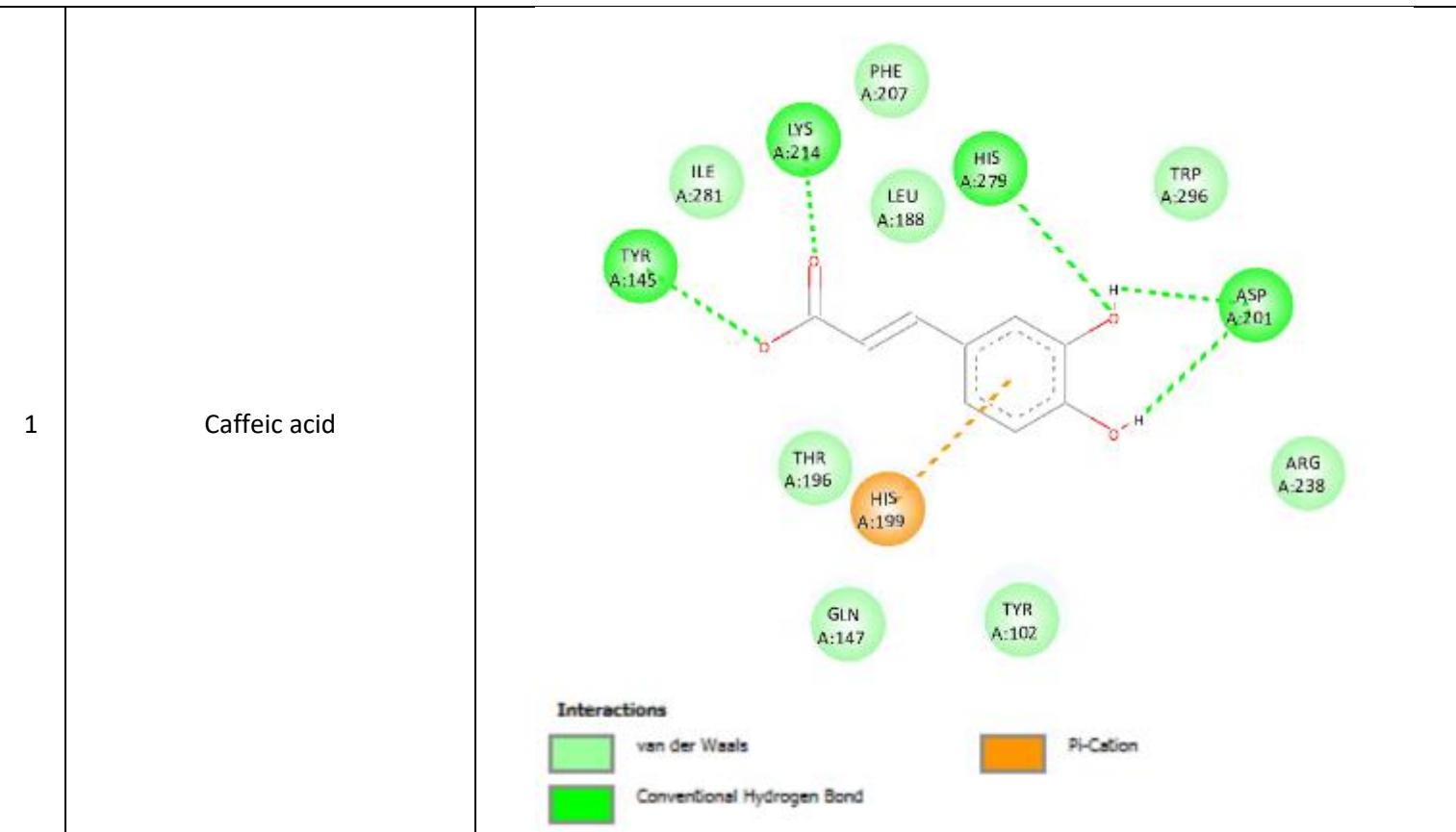
Pi-Sigma

Pi-Pi Stacked

Pi-Pi T-shaped

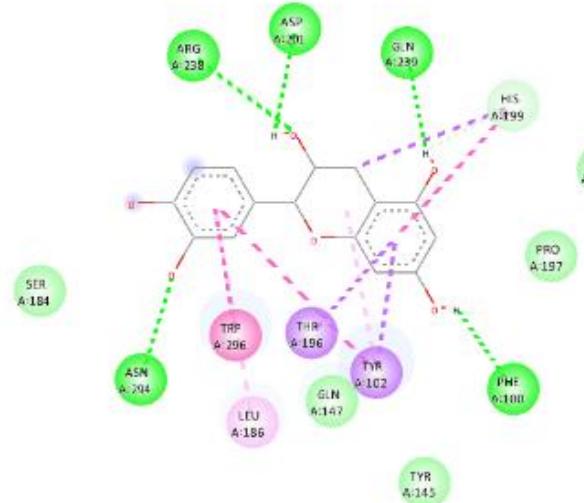
Alkyl

Pi-Alkyl



2

## (+)-Catechin

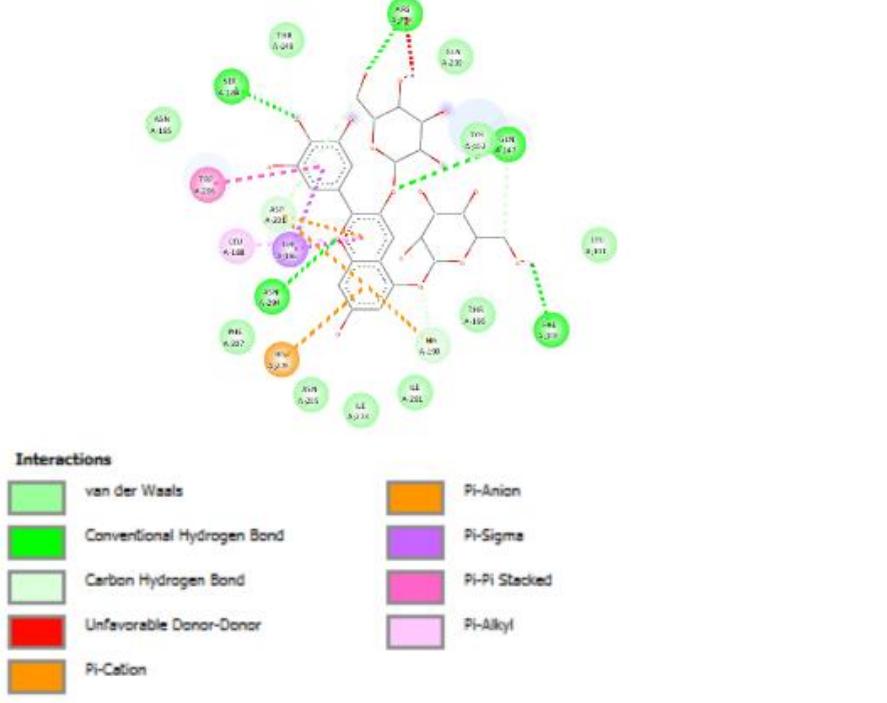


## Interactions

- |                            |                |
|----------------------------|----------------|
| van der Waals              | Pi-Pi Stacked  |
| Conventional Hydrogen Bond | Pi-Pi T-shaped |
| Pi-Donor Hydrogen Bond     | Pi-Alkyl       |
| Pi-Sigma                   |                |

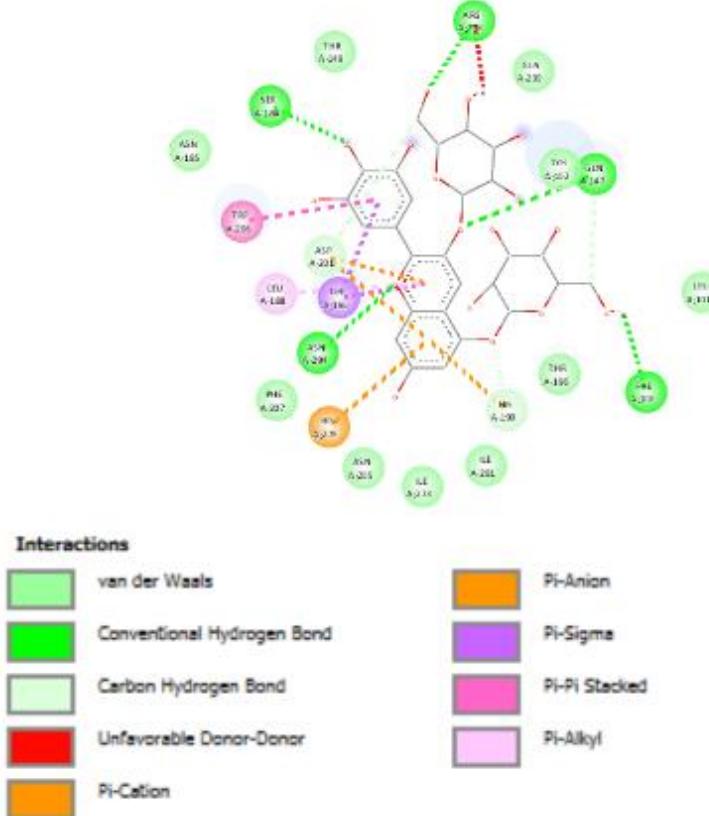
3

(2R,3S,4S,5R,6S)-2-(hydroxymethyl)-6-[7-hydroxy-3-[(2S,3R,4S,5S,6R)-3,4,5-trihydroxy-6-(hydroxymethyl)oxan-2-yl]oxy-2-(3,4,5-trihydroxyphenyl)chromenylium-5-yl]oxyoxane-3,4,5-triol



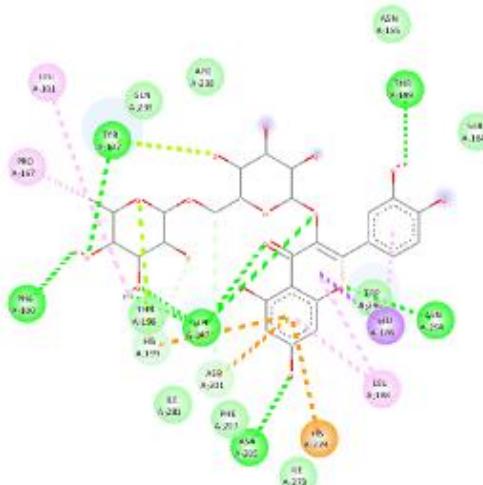
4

## Naringin



5

Rutin

**Interactions**

- |                            |              |
|----------------------------|--------------|
| van der Waals              | Pi-Sigma     |
| Conventional Hydrogen Bond | Pi-Lone Pair |
| Carbon Hydrogen Bond       | Alkyl        |
| Pi-Cation                  | Pi-Alkyl     |
| Pi-Anion                   |              |

6

## Shikimic Acid

