Digestion

D'Eustachio, P., Jassal, B., Nichols, BL.

European Bioinformatics Institute, New York University Langone Medical Center, Ontario Institute for Cancer Research, Oregon Health and Science University.

The contents of this document may be freely copied and distributed in any media, provided the authors, plus the institutions, are credited, as stated under the terms of Creative Commons Attribution 4.0 International (CC BY 4.0) License. For more information see our license.

This is just an excerpt of a full-length report for this pathway. To access the complete report, please download it at the Reactome Textbook.

16/12/2022

https://reactome.org
Introduction

Reactome is open-source, open access, manually curated and peer-reviewed pathway database. Pathway annotations are authored by expert biologists, in collaboration with Reactome editorial staff and cross-referenced to many bioinformatics databases. A system of evidence tracking ensures that all assertions are backed up by the primary literature. Reactome is used by clinicians, geneticists, genomics researchers, and molecular biologists to interpret the results of high-throughput experimental studies, by bioinformaticians seeking to develop novel algorithms for mining knowledge from genomic studies, and by systems biologists building predictive models of normal and disease variant pathways.

The development of Reactome is supported by grants from the US National Institutes of Health (P41 HG003751), University of Toronto (CFREF Medicine by Design), European Union (EU STRP, EMI-CD), and the European Molecular Biology Laboratory (EBI Industry program).

Literature references


Reactome database release: 83

This document contains 3 pathways and 3 reactions (see Table of Contents)
Digestion

Stable identifier: R-HSA-8935690

Dietary carbohydrates, fats, and proteins must be broken down to their constituent monosaccharides, fatty acids and sterols, and amino acids, respectively, before they can be absorbed in the intestine.

Dietary lipids such as long-chain triacylglycerols and cholesterol esters are hydrolyzed in the stomach and small intestine to yield long-chain fatty acids, monoacylglycerols, glycerol and cholesterol through the action of a variety of lipases, and are then absorbed into enterocytes.

Carbohydrates include starch (amylose and amyllopectin) and disaccharides such as sucrose, lactose, maltose and, in small amounts, trehalose. The digestion of starch begins with the action of amylase enzymes secreted in the saliva and small intestine, which convert it to maltotriose, maltose, limit dextrans, and some glucose. Digestion of the limit dextrans and disaccharides, both dietary and starch-derived, to monosaccharides - glucose, galactose, and fructose - is accomplished by enzymes located on the luminal surfaces of enterocytes lining the microvilli of the small intestine.

Dietary protein is hydrolyzed to dipeptides and amino acids by the action of pepsin in the stomach and an array of intestinal hydrolases. All of these enzymes are released in inactive (proenzyme) forms and activated by proteolytic cleavage within the gastrointestinal lumen (Van Beers et al. 1995; Yamada 2015).

Literature references


<table>
<thead>
<tr>
<th>Date</th>
<th>Action</th>
<th>Author/Editor</th>
</tr>
</thead>
<tbody>
<tr>
<td>2016-08-12</td>
<td>Reviewed</td>
<td>D'Eustachio, P.</td>
</tr>
<tr>
<td>2016-08-15</td>
<td>Authored, Edited</td>
<td>Jassal, B.</td>
</tr>
</tbody>
</table>
Dietary lipids such as long-chain triacylglycerols and cholesterol esters are digested in the stomach and small intestine to yield long-chain fatty acids, monoacylglycerols, glycerol and cholesterol through the action of a variety of lipases, and are then absorbed into enterocytes.

**Literature references**


Carbohydrate is a major component of the human diet, and includes starch (amylose and amylopectin) and disaccharides such as sucrose, lactose, maltose and, in small amounts, trehalose. The digestion of starch begins with the action of amylase enzymes secreted in the saliva and small intestine, which convert it to maltotriose, maltose, limit dextrins, and some glucose. Digestion of the limit dextrins and disaccharides, both dietary and starch-derived, to monosaccharides - glucose, galactose, and fructose - is accomplished by enzymes located on the luminal surfaces of enterocytes lining the microvilli of the small intestine (Van Beers et al. 1995).

**Literature references**


**Editions**

<table>
<thead>
<tr>
<th>Date</th>
<th>Action</th>
<th>Author/Reviewer</th>
</tr>
</thead>
<tbody>
<tr>
<td>2006-11-03</td>
<td>Authored</td>
<td>D'Eustachio, P.</td>
</tr>
<tr>
<td>2007-01-16</td>
<td>Reviewed</td>
<td>Nichols, BL.</td>
</tr>
<tr>
<td>2007-01-18</td>
<td>Revised</td>
<td>D'Eustachio, P.</td>
</tr>
</tbody>
</table>
Alkaline phosphatases (ALPs) are ubiquitous membrane-bound glycoproteins that catalyse the hydrolysis of phosphate monoesters in alkaline conditions (Sharma et al. 2014). To date, little is known about the physiological function of ALPs in most tissues. In humans, four isozymes exist, named from their tissue localisations. One isozyme, intestinal-type alkaline phosphatase (ALPI, IAP), possesses alkaline phosphatase activity but has no specific physiological substrate defined for it yet. It may be involved in the hydrolysis of pro-drugs in the intestine (Lowe et al. 1990).

**Literature references**


**Editions**

<table>
<thead>
<tr>
<th>Date</th>
<th>Action</th>
<th>Author</th>
</tr>
</thead>
<tbody>
<tr>
<td>2016-07-13</td>
<td>Authored, Edited</td>
<td>Jassal, B.</td>
</tr>
<tr>
<td>2016-08-12</td>
<td>Reviewed</td>
<td>D'Eustachio, P.</td>
</tr>
</tbody>
</table>
GUCY2C trimer binds GUCA2A,B

**Location:** Digestion

**Stable identifier:** R-HSA-8936214

**Type:** binding

**Compartments:** plasma membrane, extracellular region

Heat-stable enterotoxin receptor (GUCY2C, STAR) is the receptor for the endogenous peptides guanylin (GUCA2A) and uroguanylin (GUCA2B) and E.coli heat-stable enterotoxin. GUCY2C is an integral membrane protein composed of an extracellular ligand-binding domain, an intracellular domain and a guanylyl cyclase catalytic domain and functions in trimeric form (Vijayachandra et al. 2000). Once activated by its ligands, GUCY2C mediates fluid-ion homeostasis, intestinal inflammation, and cell proliferation in a cGMP-dependent manner (Arshad et al. 2013). In the intestine, salt and fluid secretion is stimulated by E.coli heat-stable enterotoxins through activation of GUCY2C. The endogenous peptides GUCA2A and GUCA2B have structural similarity to these bacterial enterotoxins and function as mediators of Cl- and water secretion in the intestine (Hamra et al. 1993, Forte et al. 1993, Basu et al. 2010).

**Literature references**


**Editions**

<table>
<thead>
<tr>
<th>Date</th>
<th>Author/Reviewer</th>
<th>Last Updated</th>
</tr>
</thead>
<tbody>
<tr>
<td>2016-08-17</td>
<td>Authored, Edited</td>
<td>Jassal, B.</td>
</tr>
<tr>
<td>2016-10-11</td>
<td>Reviewed</td>
<td>D'Eustachio, P.</td>
</tr>
</tbody>
</table>
Quercetin is an abundant flavonoid found in edible vegetables, grains and fruits and is used as an ingredient in supplements, beverages, or foods. Pirin (PIR) is a highly conserved nuclear protein (Wendler et al. 1997) which possesses quercetinase activity, transforming quercetin to 2-protocatechuoylphloroglucinol carboxylic acid (2PCPGCA) and carbon monoxide (CO) (Adams & Jia 2005). Quercetin supplements have been promoted for the treatment of a wide spectrum of diseases including cancer but there is insufficient evidence to draw any conclusive proof of its beneficial effects to date (Miles et al. 2014).

**Literature references**


# Table of Contents

- Introduction 1
- Digestion 2
  - Digestion of dietary lipid 4
  - Digestion of dietary carbohydrate 5
  - ALPI dimer hydrolyses phosphate monoesters 6
  - GUCY2C trimer binds GUCA2A,B 7
  - PIR oxygenates quercetin 8

Table of Contents 9