Absorption and metabolism of caffeic acid and chlorogenic acid in the small intestine of rats

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The absorption and metabolism in the small intestine of chlorogenic acid (5-O-caffeoylquinic acid), the main phenolic acid in the human diet, and of caffeic acid were studied in rats in order to determine whether chlorogenic acid is directly absorbed or hydrolysed in the small intestine. Chlorogenic and caffeic acids were perfused into a segment of ileum plus jejunum during 45 min (50 μM, 0.75 ml/min) using an in situ intestinal perfusion rat model with cannulation of the biliary duct, and were quantified together with their metabolites in perfusion effluent, bile and plasma. The net absorption (influent flux minus effluent flux of phenolic acids and their metabolites) accounted for 19.5% and 8% of the perfused caffeic and chlorogenic acids, respectively. A minor fraction of the perfused caffeic acid was metabolized in the intestinal wall and secreted back into the gut lumen in the form of ferulic acid (0.5% of the perfused flux). Part of the chlorogenic acid (1.2% of the perfused flux) was recovered in the gut effluent as caffeic acid, showing the presence of trace esterase activity in the gut mucosa. No chlorogenic acid was detected in either plasma or bile, and only low amounts of phenolic acids (less than 0.4%) were secreted in the bile. The present results show that chlorogenic acid is absorbed and hydrolysed in the small intestine. In contrast to numerous flavonoids, absorbed phenolic acids are poorly excreted in the bile or gut lumen. Their bioavailability therefore appears to be governed largely by their uptake into the gut mucosa.

Caffeic acid: Chlorogenic acid: Intestinal perfusion: Absorption: Rat

Hydroxycinnamic acids are one of the major classes of phenolic compounds. They are present in a large variety of fruits and vegetables (Clifford, 1999; Manach et al. 2004). The major representative of hydroxycinnamic acids in food is caffeic acid. It occurs largely conjugated with quinic acid, as in chlorogenic acid (5-O-caffeoylquinic acid; Fig. 1). Coffee, one of the most widely consumed beverages in the world, is the major dietary source of chlorogenic acid. Regular coffee-drinkers commonly ingest 0.5–1 g chlorogenic acid/d, corresponding to about 250–500 mg caffeic acid/d (Clifford, 1999).

Caffeic acid and chlorogenic acid have antioxidant properties, illustrated by their ability to scavenge various free radicals when tested in vitro (Ohnishi et al. 1994; Nardini et al. 1995; Abu-Amsha Caccetta et al. 1996; Rice-Evans et al. 1996; Foley et al. 1999; Laranjinha & Cadenas, 1999). In vivo, when ingested with the diet, caffeic acid and chlorogenic acid increase the plasma antioxidant capacity, the concentrations of endogenous antioxidants such as vitamin E and the ex vivo resistance of lipoproteins to oxidation (Nardini et al. 1997; Natella et al. 2002; Lafay et al. 2005). Chlorogenic acid is also able to reverse the pro-oxidant effects of drugs such as paraquat (Tsuchiya et al. 1996). Chlorogenic acid and caffeic acid have been reported to prevent different cancers and CVD in several experimental studies on animal models (Mori et al. 1986; Tanaka et al. 1990, 1993; Zhou et al. 1993; Suzuki et al. 2002).

The biological properties of hydroxycinnamic acids depend on their absorption in the gut and on their metabolism. The absorption of caffeic acid in the small intestine has been well characterized in both experimental animals and in man (Teuchy & Van Sumere, 1971; Camarasa et al. 1988; Azuma et al. 2000; Olthof et al. 2001; Simonetti et al. 2001; Scalbert et al. 2002), but the bioavailability of chlorogenic acid is more controversial.

In some studies, chlorogenic acid has been detected in urine with a recovery varying from 0.3% to 2.3%, suggesting absorption without structural modification (Cremin et al. 2001; Olthof et al. 2001; De Maria & Moreira, 2004; Ito et al. 2005). Other authors have failed to detect chlorogenic acid in the plasma of both rats and man after its ingestion as a pure compound or in coffee (Choudhury et al. 1999; Azuma et al. 2000; Nardini et al. 2002). Caffeic acid and its O-methylated metabolites are commonly found in plasma and urine after ingestion of chlorogenic acid in rats and man, showing that chlorogenic acid is hydrolysed in the body (Azuma et al. 2000; Rechner et al. 2001a; Nardini et al. 2002; Wittermer et al. 2005).

Such a reaction could either take place in the gut mucosa or arise from catalysis by the gut microflora. No esterase

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activity able to hydrolyse chlorogenic acid could be detected in human tissues (intestinal mucosa, liver) or biological fluids (plasma, gastric juice, duodenal fluid) in rats or man (Plumb et al. 1999; Azuma et al. 2000; Andreasen et al. 2001; Olthof et al. 2001; Rechner et al. 2001b). On the other hand, microflora in the large intestine possess esterase activity towards chlorogenic acid (Couteau et al. 2001; Rechner et al. 2004). These results suggest that caffeic acid found in plasma originates from the hydrolysis of chlorogenic acid in the colon. This would, however, be inconsistent with the rapid detection (30 min after administration of chlorogenic acid) of caffeic and ferulic acids in the plasma of rats (Azuma et al. 2000).

The aim of the present work was therefore to examine whether chlorogenic acid is absorbed or hydrolysed in the small intestine. The absorption and metabolism of chlorogenic acid in the small intestine were explored using an in situ intestinal perfusion model (segment of ileum plus jejunum) and compared with those of caffeic acid. Such a model has been used in different laboratories to study the intestinal absorption and metabolism of polyphenols (Crespy et al. 1999; Andlauer et al. 2000; Sesink et al. 2003; Arts et al. 2004; Talavera et al. 2004). Chlorogenic and caffeic acids were analysed together with their main metabolites in the perfusion effluent, bile and plasma.

Materials and methods

Chemicals

β-Glucuronidase/sulfatase from Helix pomatia (type H2) and from Escherichia coli (type X-A), chlorogenic acid, caffeic acid and ferulic acid were purchased from Sigma (L’Isle d’Abeau Chesnes, France). Isoferulic acid was purchased from Extrasthèse (Genay, France).

Animals and diets

Eight Wistar rats, born at the Institut National de la Recherche Agronomique and weighing approximately 150 g, were housed, two per cage, in temperature-controlled rooms (22°C), with a dark period from 20.00 hours to 08.00 hours and with access to food from 16.00 hours to 08.00 hours. They were fed a standard semi-purified diet (composition, per kg of diet: 755 g wheat starch, 150 g casein, 50 g peanut oil, 35 g mineral AIN-93M mixture, 10 g vitamin AIN-93 mixture supplemented with 1.36 g choline) for 2 weeks. Animals were maintained and handled according to the recommendations of the National Institute for Agricultural Research Ethics Committee, in accordance with Decree 87-848.

In situ gut perfusion

Rats were anaesthetized with sodium pentobarbital (40 mg/kg body weight) 24 h after the end of the last meal by intraperitoneal injection, maintained at 37°C on a heat plate and kept alive during the perfusion period. After cannulation of the biliary duct, a perfusion of a jejunal and ileal segment of intestine (from the duodeno-jejunal flexure to 5 cm distal to the ileocecal valve) was prepared by installing cannulas at each extremity. This segment was continuously perfused in situ with a buffer containing KH2PO4 (5 mmol/l), K2HPO4 (2.5 mmol/l), NaHCO3 (5 mmol/l), NaCl (50 mmol/l), KCl (50 mmol/l), CaCl2 (2 mmol/l), MgCl2 (2 mmol/l), glucose (10 mmol/l) and taurocholic acid (1 mmol/l; pH adjusted at 6.7) and supplemented with 50 μM-caffeic acid (n 4) or 50 μM-chlorogenic acid (n 4). A flow rate of about 0.75 ml/min and a temperature of 37°C were maintained during the 45 min of the perfusion (Crespy et al. 1999; Andlauer et al. 2000; Sesink et al. 2003; Arts et al. 2004; Talavera et al. 2004). The average flow rate was determined for each experiment over the 45 min of perfusion. These quantities of chlorogenic acid perfused correspond to an intake in man of about 500 mg chlorogenic acid (calculated for rats and human subjects consuming, respectively, 20 g and 500 g DM/d). This amount of chlorogenic acid corresponds to three or four cups of coffee (Scalbert & Williamson, 2000).

Both caffeic and chlorogenic acids were quantified in the buffer at the end of the experiment and found to be stable in the buffer throughout the perfusion period. The effluent at the exit of ileum and the bile were collected throughout the 45 min of the perfusion period for analysis of phenolic acids. At the end of the experiment, blood samples were withdrawn from the mesenteric vein into heparinized tubes. Perfusate, bile and plasma samples were acidified with 10 mM-acetic acid and stored at −20°C.
Sample treatment. Perfusate, bile and plasma samples were spiked at 5 μmol/l with sinapic acid or vanillic acid as internal standard. Samples collected after caffeic acid perfusion were acidified to pH 4.9 with 0.1 volume of 0.5 M-sodium acetate containing 2 g ascorbic acid/l (pH 3). Perfusate and bile samples were incubated for 45 min at 37°C, and plasma samples for 4 h at the same temperature, in the presence of 10^8 units β-glucuronidase and 2.5 × 10^6 units sulfatase from *H. pomatia* per litre.

The procedure was modified for samples collected after chlorogenic acid perfusion as we noticed that chlorogenic acid was partially hydrolysed by *H. pomatia* enzymes (Ito et al. 2005). Samples were then adjusted to pH 6.8 with 0.1 volume of 0.5 M-sodium acetate containing 2 g ascorbic acid/l (pH 4.9). Perfusate and bile samples were incubated for 45 min at 37°C, plasma samples for 4 h at the same temperature, in absence or in the presence of 2 × 10^6 units β-glucuronidase from *E. coli* per litre. No caffeic acid was formed in these conditions, as confirmed by the addition of pure chlorogenic acid to a control plasma containing the enzyme from *E. coli*. Sulfatase activity was also present in the enzyme fraction as observed using a rat plasma sample containing known sulfate esters of quercetin (data not shown).

Samples were extracted by adding 2.85 volumes of methanol–H2O–concentrated HCl (70:28:2, by vol.) and centrifuged for 4 min at 12,000g. The resulting supernatant was analysed as described below.

Determination of the intestinal and biliary fluxes. All the calculated fluxes were expressed in nmol/min. The fluxes in the effluent were calculated taking into account the intestinal absorption of H2O as previously described (Crespy et al. 1999). For a given phenolic acid, the net transfer into the enterocyte was evaluated by the difference between the perfused flux and the flux of perfused molecules recovered in the non-hydrolysed effluent at the end of the perfusion. The secretion of conjugates back into the gut lumen was determined as the difference between the flux of perfused molecules measured in the non-hydrolysed effluent and the flux of total phenolic acids in the hydrolysed effluent. The net absorption was calculated as the difference between the net transfer into the enterocyte and the intestinal secretion of metabolites (*O*-methylated conjugated forms). The biliary secretion of conjugated forms was determined as the product between the biliary flow rate (μl/min) and the concentrations of the phenolic acids measured in the bile after enzymatic hydrolysis (μmol/l).

Chromatographic conditions. The conjugated metabolites of phenolic acids were analysed by HPLC coupled to an eight-electrode CoulArray Model 5600 detector (Eurosep, Cergy, France) with potentials set at 25, 100, 250, 360, 400, 475 and 550 mV. The system was fitted with a 5 μm C-18 Hypersil BDS analytical column (150 × 4.6 mm; Life Sciences International, Cergy, France). Mobile phase A was 5 % acetonitrile in 30 mM-NaH2PO4 at pH 3, and mobile phase B was 50 % acetonitrile in 30 mM-NaH2PO4 at pH 3. The separation was performed at 35°C. The flow rate was 0.8 ml/min, and the following gradient was applied: linear gradient from 0 % to 22 % B from 0 to 21 min; linear gradient from 22 % to 29 % B from 21 to 41 min; 100 % B from 41 to 45 min. The identification of phenolic acids was based on the comparison with appropriate standards (Fig. 2). In perfused buffer, detection limits were 0.06 μmol/l for chlorogenic, caffeic and isoferulic acids, and 0.08 μmol/l for ferulic acid. In bile and plasma, detection limits were 0.06 μmol/l for caffeic and isoferulic acids, and 0.08 μmol/l for ferulic acid.

Data analysis

Data were treated with the Instat statistical program (Instat, San Diego, CA). Values are given as means with their standard errors. An unpaired *t* test or Mann–Whitney test (non-parametric test) was applied to determine differences.
between values. Differences with a value of $P<0.05$ were considered significant.

**Results**

**Intestinal metabolism of caffeic acid**

When caffeic acid was perfused at 44.6 (SE 1.3) nmol/min into the upper intestinal tract, both caffeic and ferulic acids were recovered in the effluent at the end of perfusion. Their respective fluxes in the non-hydrolysed effluent were 35.7 (SE 1.1) and 0.17 (SE 0.01) nmol/min (Table 1). These two fluxes correspond to respectively 80% and 0.4% of the perfused flux of caffeic acid (Fig. 3). The net transfer of caffeic acid into the enterocyte was 8.8 (SE 0.7) nmol/min, corresponding to 20% of the perfused flux.

When the effluent was treated with a β-glucuronidase/sulfatase, the flux of caffeic acid did not change significantly – 36.7 (SE 1.2) v. 35.7 (SE 1.1) nmol/min in non-hydrolysed effluent – indicating the absence of intestinal secretion of conjugates for caffeic acid. Conversely, the flux of ferulic acid in the effluent significantly increased after enzymatic hydrolysis (0.17 (SE 0.01) v. 0.24 (SE 0.01) nmol/min in non-hydrolysed and hydrolysed effluent; $P<0.05$), showing a release of ferulic acid conjugates. As the biliary duct had been cannulated before starting perfusion, the ferulic acid and conjugated derivatives found in the effluent at the end of perfusion had an intestinal origin and represented 0.5% of the perfused flux. Thus, the recovery of caffeic acid and its metabolites accounted for 80.5% of the perfused flux. The net absorption of caffeic acid is therefore determined by the difference between the perfused flux of caffeic acid and the flux of total phenolic acids in the effluent, accounting for 19.5% of the perfused flux.

**Intestinal metabolism of chlorogenic acid**

When chlorogenic acid was perfused at 40.4 (SE 0.8) nmol/min into the upper intestinal tract, both chlorogenic and caffeic acids were recovered in the effluent at the end of perfusion. Their respective fluxes in the non-hydrolysed effluent were 36.7 (SE 1.1) nmol/min and 0.23 (SE 0.01) nmol/min (Table 1). These phenolic acids fluxes correspond to 90.8% and 0.6% of the perfused flux of chlorogenic acid respectively (Fig. 3). The net transfer of chlorogenic acid into the enterocyte was 3.7 (SE 0.7) nmol/min, corresponding to 9.2% of the perfused flux.

When the effluent was treated with deconjugating enzymes, the flux of chlorogenic acid was not changed – 36.7 (SE 0.3) v. 36.7 (SE 1.1) nmol/min in non-hydrolysed effluent – indicating the absence of intestinal secretion of conjugates of chlorogenic acid. After enzymatic hydrolysis, the flux of caffeic acid significantly increased (0.48 (SE 0.02) nmol/min; $P<0.05$ v. caffeic acid in non-hydrolysed effluent), indicating the presence of caffeic acid conjugates released in the lumen. Free caffeic acid and conjugated derivatives found in the effluent resulted from intestinal activity, and their secretion into the lumen represented 1.2% of perfused flux. Thus, net absorption of chlorogenic acid was 3.2 (SE 0.8) nmol/min and accounted for 8% of the perfused flux.

**Metabolites excreted in the bile after perfusion of hydroxycinnamic acids**

Phenolic acids were analysed in the bile after enzymatic deconjugation. Three phenolic acids – caffeic, ferulic and isoflavonoid acid – were found in the bile after caffeic acid perfusion (Table 2). The respective biliary fluxes were 109 (SE 23) pmol/min, 35.0 (SE 5.0) pmol/min and 26.3 (SE 1.5) pmol/min. Caffeic acid was also found in low concentration in the bile after chlorogenic acid perfusion (Table 2).

**Metabolites recovered in the plasma after perfusion of hydroxycinnamic acids**

Phenolic acids were analysed after enzymatic deconjugation in the plasma mesenteric vein. Samples collected after the perfusion of caffeic acid showed the presence of ferulic and isoflavonoid acids (Table 3). Low amounts of caffeic acid were also detected in the mesenteric vein of two rats. Analysis of the plasma samples collected after chlorogenic acid perfusion showed the presence of isoflavonoid acid in mesenteric plasma.

**Discussion**

Epidemiological and clinical studies have shown that the consumption of coffee, a major dietary source of chlorogenic acids, reduces the risk of non-insulin-dependent diabetes, decreases hyperglycaemia and increases plasma antioxidant capacity (Natella et al. 2002; van Dam & Feskens, 2002; Yamaji et al. 2004). Coffee polyphenols, particularly chlorogenic acid, could contribute to these effects (Scalbert et al. 2005). The biological effects of polyphenols rely largely on their bioavailability (Manach et al. 2004), but the bioavailability...
and metabolism of chlorogenic acid have been poorly studied. The \textit{in situ} perfusion of caffeic and chlorogenic acids into a segment of the upper intestinal tract of rats (jejunum plus ileum) allows us to compare their absorption in conditions close to normal physiology.

The esterification of caffeic acid, as with chlorogenic acid, more than halved its absorption: 19.5\% and 8\% of the perfused flux for caffeic and chlorogenic acids, respectively, were absorbed. Using the same model of perfusion as used here, the intestinal absorption of ferulic acid has been determined and was higher than that of caffeic acid (56.1\% v. 19.5\% of the perfused flux respectively; Adam \textit{et al.} 2002), showing that the O-methylation of caffeic acid increases its transfer through the intestinal barrier. Similar conclusions were obtained by perfusing an isolated rat intestinal segment (Spencer \textit{et al.} 1999). The values for net transfer of caffeic and chlorogenic acids through the small intestine also appear much lower than those observed for flavonoids, using the same \textit{in situ} perfusion model, which varied from 35.2\% for catechin to 78\% for kaempferol (Crespy \textit{et al.} 2003).

The absorption of different phenolic acids has also been studied with Caco-2 cell monolayers (Konishi \textit{et al.} 2003\textit{a,b}; Konishi & Kobayashi, 2004). The transepithelial flux of chlorogenic acid was found to be much lower than that of caffeic acid, which was itself lower than that of ferulic acid, in agreement with the present results. Free phenolic acids would largely be absorbed through an active mechanism involving monocarboxylic acid transporters, whereas the absorption of chlorogenic acid would follow the paracellular route. In experiments with volunteers who had undergone colonic ablation, chlorogenic acid absorption was also three times lower than that of caffeic acid, a proportion comparable to the one observed in the present work (Olthof \textit{et al.} 2001).

Chlorogenic acid has been identified by several authors in urine after the ingestion of coffee or pure chlorogenic acid, suggesting a direct absorption in the gut (Cremin \textit{et al.} 1999).

### Table 2. Biliary flux of phenolic acids after \textit{in situ} perfusion of caffeic acid or chlorogenic acid*

<table>
<thead>
<tr>
<th>Phenolic Acid</th>
<th>Caffeic Acid (pmol/min)</th>
<th>Chlorogenic Acid (pmol/min)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Mean</td>
<td>SE</td>
</tr>
<tr>
<td>Caffeic acid</td>
<td>109</td>
<td>23</td>
</tr>
<tr>
<td>Ferulic acid</td>
<td>35.0</td>
<td>5.0</td>
</tr>
<tr>
<td>Isoferulic acid</td>
<td>28.3</td>
<td>1.5</td>
</tr>
</tbody>
</table>

ND, not detected.

* Biliary flux = concentration of (free + conjugated) phenolic acid \times biliary flow. Biliary flow was 15.1 (S.E. 2.3) \mu l/min for caffeic acid and 15.6 (S.E. 2.6) \mu l/min for chlorogenic acid.

### Table 3. Concentrations of caffeic acid metabolites and chlorogenic acid metabolites in the mesenteric plasma of rats after perfusion with caffeic acid or chlorogenic acid

<table>
<thead>
<tr>
<th>Phenolic Acid</th>
<th>Caffeic Acid (\mu M)</th>
<th>Chlorogenic Acid (\mu M)</th>
</tr>
</thead>
<tbody>
<tr>
<td></td>
<td>Mean</td>
<td>SE</td>
</tr>
<tr>
<td>Caffeic acid</td>
<td>Trace</td>
<td></td>
</tr>
<tr>
<td>Ferulic acid</td>
<td>0.71</td>
<td>0.04</td>
</tr>
<tr>
<td>Isoferulic acid</td>
<td>0.14</td>
<td>0.01</td>
</tr>
<tr>
<td>Chlorogenic acid</td>
<td>ND</td>
<td></td>
</tr>
</tbody>
</table>

ND, not detected (limits of detection for caffeic, chlorogenic and isofurulic acids 0.06 \mu M and for ferulic acid 0.08 \mu M).
isoferulic acid, a reaction catalysed by catechol-
plasma.
phenolic acids absorbed from chlorogenic acid in the present
of phenolic acids in the plasma suggests absorption in the
intestinal lumen and in the bile. Isoferulic acid could
be detected in plasma. This can probably be explained by
the relatively low amounts of phenolic acid absorbed from
chlorogenic acid perfusion, only caffeic acid was found in the
mesenteric plasma.
A portion of the phenolic acids absorbed is quickly metab-
olized and in particular methylated into ferulic acid and
isoferulic acid, a reaction catalysed by catechol-O-methyl-
transferase previously identified in the intestine (Männistö
& Kaakkola, 1999). Some of this ferulic acid was secreted
back into the gut lumen during catechic acid perfusion. Ferulic
and isofeulic acids identified in the bile could originate from
both the secretion of ferulic acid formed in the gut mucosa and
from liver catechol-O-methyltransferase activity. With
chlorogenic acid perfusion, only caffeic acid was found in the
intestinal lumen and in the bile. Isoferulic acid could be
detected in plasma. This can probably be explained by
the relatively low amounts of phenolic acid absorbed from
chlorogenic acid.
The relative proportions of ferulic acid and isofeulic acid
in both bile and plasma show a preferential O-methylation in
the meta position, in agreement with previous studies with rats
(Gonthier et al. 2003; Wittmer & Veit, 2003). A similar prefer-
ence for methylation in the meta position is known
for other polyphenols with catechol groups such as
quercetin, catechin or cyanidin (Manach et al. 1999; Felgines
et al. 2002).
Other reactions involving phenolic acids also take place in
the gut mucosa. Phenolic acids were determined in the gut
effluent with and without enzymatic hydrolysis to look for
the presence of glucuronides and sulfate esters formed in
the mucosa and secreted back into the gut lumen. Enzymatic
decconjugation significantly increased the level of ferulic acid
recovered in the effluent after caffeic acid perfusion. However,
ferulic acid conjugates accounted for only 0.16 % of the
perfused flux of caffeic acid. Similarly, the level of caffeic
acid conjugates determined in the effluent after the perfusion
of chlorogenic acid represented 0.6 % of the perfused flux.
Similar experiments, carried out with different flavonoids,
have shown wide variations in the levels of conjugates
secreted back to the gut lumen, from 0 % of the perfused
flux for catechin to 52 % for quercetin (Crespy et al. 2003).
The present results show that the secretion of hydroxycym-
namic acid conjugates into the gut lumen is low compared
with that of some flavonoids, in agreement with a previous
study on ferulic acid (Adam et al. 2002).
The biliary excretion of hydroxycinnamic acids has pre-
viously been established by liver perfusion or intraperitoneal
injection of caffeic acid in the rat (Scheline, 1968; Gumbinger
et al. 1993). In the present work, biliary excretion did not
exceed 0.4 % of the perfused dose of caffeic acid, suggesting
a limited contribution of this route.
The splanchnic metabolism of caffeic and chlorogenic
acids is summarised in Fig. 3. A total of 19.5 % of the caffeic
acid and 8 % of the chlorogenic acid perfused through the
small intestine reached the mesenteric blood. A minute frac-
tion of the phenolic acids transferred into the gut mucosa was
secreted back to the intestinal lumen as ferulic acid for caf-
feeic acid perfusion and caffeic acid for chlorogenic acid per-
fusion. This result demonstrates for the first time the in vivo
hydrolysis of chlorogenic acid in the gut mucosa, but esterase
activity towards this phenolic acid appears very low. Pheno-
lic acids found in the mesenteric blood reached the liver,
where a low fraction was excreted into the bile in its
unchanged or O-methylated form, showing that biliary
excretion is not the major route of excretion for phenol-
inc acids. Therefore, the fraction of phenolic acids reaching
the peripheral tissues (19.1 % and 8 % for caffeic and chlorogenic
acid, respectively) depends largely on the absorption through
the gut barrier.
The present results for caffeic and chlorogenic acids can
be compared with those previously obtained with other poly-
phenols using the same in situ perfusion model (Crespy et al.
2003). For example, the fractions of eryodictiol and querce-
tin reaching the peripheral tissues were similar (20 % and
9 %, respectively) to that observed here for phenolic acids.
However, their splanchnic metabolism differs significantly
from that of caffeic and chlorogenic acids. Quercetin is
characterized by high intestinal uptake (67 %), high
secretion back into the gut lumen (52 %) and significant
secretion into the bile (6 %). Eryodictiol also showed a
high intestinal uptake (59 %), a moderate secretion back
into the gut lumen and a high biliary excretion (23 %). Caff-
eic and chlorogenic acids also differ from ferulic acid,
which showed a significant excretion through the bile
(6 %; Adam et al. 2002). Therefore, the factors determining
the bioavailability of polyphenols in inner body tissues vary
according to polyphenols. The bioavailability of caffeic and
chlorogenic acids appears to be largely determined by their
uptake into the gut mucosa, as previously reported for cate-
chin (Crespy et al. 2003).

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Intestinal metabolism of phenolic acids


References


