# Protein allocation and utilization in the versatile chemolithoautotroph Cupriavidus necator

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Section No.	Headings	Sentences
Section: 1	Summary	14
Section: 2	Introduction	13
N/A		0

# Protein allocation and utilization in the versatile chemolithoautotroph Cupriavidus necator

\$1 [001]	Summary	

**S1 [002]** Bacteria must balance the different needs for substrate assimilation, growth functions, and resilience in order to thrive in their environment.

Bacteria must balance the different needs ...
... for substrate assimilation, ...
... growth functions, ...
... and resilience ...

... in order ...

... to thrive ...

... in their environment.

**S1 [003]** Of all cellular macromolecules, the bacterial proteome is by far the most important resource and its size is limited.

Of all cellular macromolecules, ...

- ... the bacterial proteome is ...
- ... by far the most important resource ...
- ... and its size is limited.

**S1 [004]** Here, we investigated how the highly versatile 'knallgas' bacterium Cupriavidus necator reallocates protein resources when grown on different limiting substrates and with different growth rates.

Here, ...

- $\dots$  we investigated how the highly versatile 'knallgas' bacterium Cupriavidus necator reallocates protein resources  $\dots$
- ... when grown ...
- ... on different limiting substrates ...
- ... and with different growth rates.

**S1 [005]** We determined protein quantity by mass spectrometry and estimated enzyme utilization by resource balance analysis modeling.

We determined protein quantity ...

- ... by mass spectrometry ...
- $\dots$  and estimated enzyme utilization  $\dots$
- ... by resource balance analysis modeling.
- **S1 [006]** We found that C. necator invests a large fraction of its proteome in functions that are hardly utilized.

We found ...

- ... that C. necator invests a large fraction ...
- ... of its proteome ...

```
... that are hardly utilized.
S1 [007]
              Of the enzymes that are utilized, many are present in excess abundance.
                    Of the enzymes ...
                    ... that are utilized, ...
                    ... many are present ...
                   ... in excess abundance.
S1 [008]
              One prominent example is the strong expression of CBB cycle genes such as Rubisco
              during growth on fructose.
                   One prominent example is the strong expression ...
                   ... of CBB cycle genes ...
                   ... such as Rubisco ...
                   ... during growth ...
                   ... on fructose.
S1 [009]
              Modeling and mutant competition experiments suggest that CO2-reassimilation through
              Rubisco does not provide a fitness benefit for heterotrophic growth, but is rather an
              investment in readiness for autotrophy.
                   Modeling ...
                   ... and mutant competition experiments suggest ...
                   ... that CO2-reassimilation ...
                    ... through Rubisco does not provide a fitness benefit ...
                   ... for heterotrophic growth, ...
                    ... but is rather an investment ...
                   ... in readiness ...
                    ... for autotrophy.
S1 [010]
              Highlights
                   Highlights
S1 [011]
              A large fraction of the C. necator proteome is related to environmental readiness
                   A large fraction ...
                   ... of the C. necator proteome is related ...
                    ... to environmental readiness
S1 [012]
              Highly utilized enzymes are more abundant and less variable
                   Highly utilized enzymes are more abundant ...
                    ... and less variable
S1 [013]
              Autotrophy related enzymes are largely underutilized
                    Autotrophy related enzymes are largely underutilized
```

Knockout of Calvin cycle genes increases growth rate on sugar but decreases affinity

... in functions ...

S1 [014]

Knockout ...

```
... of Calvin cycle genes increases growth rate ...
... on sugar ...
... but decreases affinity
```

# S2 [015] Introduction

**S2 [016]** Cupriavidus necator (formerly Ralstonia eutropha) is a model aerobic lithoautotroph and formatotroph, and is notable for production of the storage polymer polyhydroxybutyrate (PHB) [Yishai et al., 2016, Brigham, 2019].

```
Cupriavidus necator ...
... (formerly Ralstonia eutropha) ...
... is a model aerobic lithoautotroph ...
... and formatotroph, ...
... and is notable ...
... for production ...
... of the storage polymer polyhydroxybutyrate ...
... (PHB) ...
... [Yishai et al., 2016, ...
... Brigham, 2019].
```

**S2 [017]** Cupriavidus necator H16 (hereafter abbreviated C. necator) is a soil-dwelling bacterium with a large genome (~6,600 genes) distributed on two chromosomes and one megaplasmid [Pohlmann et al, 2006].

```
Cupriavidus necator H16 ...
... (hereafter abbreviated C. necator) ...
... is a soil-dwelling bacterium ...
... with a large genome ...
... (~6,600 genes) ...
... distributed ...
... on two chromosomes ...
... and one megaplasmid ...
... [Pohlmann et al, 2006].
```

**S2 [018]** It features a wide arsenal of metabolic pathways for xenobiotics degradation, hydrogen and formate oxidation, carbon fixation via the Calvin-Bensson-Bassham (CBB) cycle, and utilization of nitrate/nitrite as alternative electron acceptors (de-nitrification) [Cramm, 2008].

```
It features a wide arsenal ...
... of metabolic pathways ...
... for xenobiotics degradation, ...
... hydrogen ...
... and formate oxidation, ...
... carbon fixation ...
... via the Calvin-Bensson-Bassham ...
... (CBB) ...
... cycle, ...
... and utilization ...
... of nitrate/nitrite ...
```

```
... as alternative electron acceptors ...
... (de-nitrification) ...
... [Cramm, 2008].
```

**S2 [019]** Several operons for substrate assimilation are present in multiple copies, often on different chromosomes (e.g. cbb operon, hydrogenases, formate dehydrogenases).

Several operons ...
... for substrate assimilation are present ...
... in multiple copies, ...
... often ...
... on different chromosomes ...
... (e.g. cbb operon, ...
... hydrogenases, ...
... formate dehydrogenases).

**S2 [020]** A detailed reconstruction of its metabolic network suggested that it can metabolize 229 compounds [Park et al., 2011].

```
A detailed reconstruction ...
... of its metabolic network suggested ...
... that it can metabolize 229 compounds ...
... [Park et al., 2011].
```

**S2** [021] Interestingly, C. necator prefers organic acids as growth substrate over sugars.

```
Interestingly, ...
... C. necator prefers organic acids ...
... as growth substrate ...
... over sugars.
```

**S2 [022]** The only sugar that supports growth is fructose, which is metabolized via the Entner-Doudoroff (ED) pathway [Alagesan et al., 2018].

```
The ...
... only sugar ...
... that supports growth is fructose, ...
... which is metabolized ...
... via the Entner-Doudoroff ...
... (ED) ...
... pathway ...
... [Alagesan et al., 2018].
```

**S2 [023]** Although the metabolic versatility of C. necator is interesting from a biotechnological point of view, we wondered if it does not come at a considerable cost for the cell.

```
Although the metabolic versatility ...
... of C. necator is interesting ...
... from a biotechnological point ...
... of view, ...
... we wondered ...
... if it does not come ...
... at a considerable cost ...
... for the cell.
```

# **End of Sample Audit**

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