

## **Metabolic reprogramming of *Klebsiella pneumoniae* exposed to serum and its potential implications in host immune system evasion and resistance**

Amanda Naiara Silva Moraes<sup>1</sup>, Juliana Miranda Tatara<sup>1</sup>, Rafael Lopes da Rosa<sup>1</sup>, Franciele Maboni Siqueira<sup>2</sup>, Guilherme Domingues<sup>3</sup>, Markus Berger<sup>4,5</sup>, Jorge Almeida Guimarães<sup>4</sup>, Afonso Luís Barth<sup>6</sup>, Patricia Orlandi Barth<sup>6</sup>, John R. Yates III<sup>7</sup>, Walter Orlando Beys-da-Silva<sup>1,8</sup>, Lucélia Santi<sup>1,8,\*</sup>

<sup>1</sup> Post-Graduation Program in Cellular and Molecular Biology, Federal University of Rio Grande do Sul. Porto Alegre, RS. Brazil. 91501-970

<sup>2</sup> Faculty of Veterinary, Federal University of Rio Grande do Sul. Porto Alegre, RS. Brazil. 91540-000

<sup>3</sup> Bruno Born Hospital. Lajeado, RS. Brazil. 95900-010

<sup>4</sup> Center of Experimental Research, Clinical Hospital of Porto Alegre. Porto Alegre, RS. Brazil. 90035-903

<sup>5</sup> Tick-Pathogen Transmission Unit, Laboratory of Bacteriology, National Institute of Allergy and Infectious Diseases, Hamilton, MT, USA. 20892

<sup>6</sup> Bacterial Resistance Research Laboratory, Clinical Hospital of Porto Alegre. Porto Alegre, RS. Brazil. 90035-903

<sup>7</sup> Department of Molecular Medicine, Scripps Research. La Jolla, CA. USA. 92037

<sup>8</sup> Faculty of Pharmacy, Federal University of Rio Grande do Sul. Porto Alegre, RS. Brazil. 90610-000

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**Supplementary Table 1. Exclusive proteins identified after 1 hour of native serum exposure.** Proteins were identified using the software Patternlab, module AAPV, using a probability 0.01.

| Accession number * | Spectral count | Protein identification  |
|--------------------|----------------|---|
| AEW63575.1         | 242            | 30S ribosomal subunit protein S12                                   |
| AEW60145.1         | 124            | enterobactin synthase subunit E                                     |
| AEW60659.1         | 62             | ribonuclease E  |
| AEW63547.1         | 51             | 50S ribosomal protein L15   |
| AEW62901.1         | 50             | GDP/GTP pyrophosphokinase   |
| AEW63488.1         | 48             | regulator of ftsI, penicillin binding protein 3, septation function |
| AEW63405.1         | 40             | ATP-dependent RNA helicase DeaD                                     |
| AEW59216.1         | 38             | leucyl aminopeptidase   |
| AEW59510.1         | 32             | UDP-N-acetylmuramate--L-alanine ligase                              |
| AEW63837.1         | 28             | 50S ribosomal protein L33   |
| AEW63565.1         | 27             | 50S ribosomal protein L4  |
| AEW60915.1         | 27             | DNA-binding transcriptional regulator TyrR                          |
| AEW58928.1         | 25             | isocitrate lyase  |
| AEW62887.1         | 24             | sulfate adenylyltransferase subunit 2                               |
| AEW60369.1         | 23             | glutamine ABC transporter permease protein                          |
| AEW62597.1         | 22             | polyphosphate kinase  |
| AEW61822.1         | 22             | cysteine desulfurase ATPase componente                              |
| AEW60144.1         | 20             | isochorismate synthase  |
| AEW59491.1         | 18             | isopropylmalate isomerase large subunit                             |
| AEW58701.1         | 17             | asparagine synthetase AsnA  |
| AEW60181.1         | 17             | putative aminotransferase   |
| AEW63797.1         | 17             | putative glutathione S-transferase                                  |
| AEW60138.1         | 16             | ATP-dependent serine activating enzyme                              |
| AEW62611.1         | 16             | GTP-binding protein EngA  |
| AEW59410.1         | 16             | bifunctional aspartokinase I/homoserine dehydrogenase I             |
| AEW59458.1         | 15             | dimethyladenosine transferase                                       |
| AEW58826.1         | 15             | ATP-dependent RNA helicase RhlB                                     |
| AEW62669.1         | 15             | L-aspartate oxidase   |
| AEW59810.1         | 14             | 1-deoxy-D-xylulose-5-phosphate synthase                             |
| AEW63776.1         | 14             | major cold shock protein  |
| AEW63769.1         | 14             | putative sugar phosphate isomerase/epimerase                        |
| AEW58785.1         | 13             | bifunctional aspartate kinase II/homoserine dehydrogenase II        |
| AEW58930.1         | 13             | transcriptional repressor IclR                                      |
| AEW62987.1         | 13             | N-acetylglutamate synthase  |
| AEW63426.1         | 12             | 23S rRNA methyltransferase J  |
| AEW92118.1         | 12             | fertility inhibition protein (plasmid)                              |
| AEW59469.1         | 12             | ATP-dependent helicase HepA   |
| AEW61883.1         | 12             | dsDNA-mimic protein   |
| AEW62915.1         | 12             | L-serine deaminase 2  |

|            |    |  |
|------------|----|--|
| AEW62677.1 | 12 | putative NAD(P)-binding and ATP-binding acyl-CoA synthetase                                    |
| AEW58980.1 | 12 | excinuclease ABC subunit A   |
| AEW62861.1 | 11 | S-adenosylmethionine-dependent methyltransferase   |
| AEW62256.1 | 11 | dTDP-D-glucose 4,6-dehydratase   |
| AEW61020.1 | 11 | Oxidoreductase   |
| AEW59115.1 | 11 | putative GTPase HflX   |
| AEW58916.1 | 11 | hypothetical protein KPHS_02180  |
| AEW59112.1 | 11 | DNA mismatch repair protein  |
| AEW58821.1 | 11 | ATP-dependent DNA helicase Rep   |
| AEW63862.1 | 11 | bifunctional (p)ppGpp synthetase II/ guanosine-3',5'-bis pyrophosphate 3'-pyrophosphohydrolase |
| AEW61325.1 | 10 | 6-phosphogluconate dehydrogenase   |
| AEW62851.1 | 9  | iron ABC superfamily ATP binding cassette transporter, binding protein                         |
| AEW60206.1 | 9  | hypothetical protein KPHS_15080  |
| AEW62614.1 | 9  | histidyl-tRNA synthetase   |
| AEW59571.1 | 9  | sugar fermentation stimulation protein A   |
| AEW60760.1 | 9  | putative glycosyl transferase  |
| AEW62399.1 | 9  | OmpK36 porin   |
| AEW58782.1 | 8  | 50S ribosomal protein L31  |
| AEW60847.1 | 8  | putative hydroxylase   |
| AEW60146.1 | 8  | 2,3-dihydro-2,3-dihydroxybenzoate synthetase, isochroismatase                                  |
| AEW62446.1 | 8  | aminotransferase AlaT  |
| AEW63545.1 | 8  | 30S ribosomal protein S13  |
| AEW62598.1 | 8  | Exopolyphosphatase   |
| AEW58927.1 | 8  | malate synthase  |
| AEW61548.1 | 8  | cryptic nitrate reductase 2 alpha subunit  |
| AEW63718.1 | 8  | hypothetical protein KPHS_50200  |
| AEW63359.1 | 7  | hypothetical protein KPHS_46610  |
| AEW60517.1 | 7  | integration host factor subunit beta   |
| AEW62671.1 | 7  | ATP-dependent RNA helicase SrmB  |
| AEW59924.1 | 7  | phosphoribosylaminoimidazole carboxylase ATPase subunit  |
| AEW62886.1 | 7  | sulfate adenylyltransferase subunit 1  |
| AEW61342.1 | 7  | deoxyribose-phosphate aldolase   |
| AEW62634.1 | 7  | inositol monophosphatase   |
| AEW58816.1 | 7  | threonine dehydratase  |
| AEW63765.1 | 7  | biotin sulfoxide reductase   |
| AEW58971.1 | 7  | tRNA-dihydrouridine synthase A   |
| AEW59114.1 | 6  | RNA-binding protein Hfq  |
| AEW58942.1 | 6  | hypothetical protein KPHS_02440  |
| AEW62674.1 | 6  | putative methyltransferase   |
| AEW61628.1 | 6  | alcohol dehydrogenase class III  |
| AEW63467.1 | 6  | 30S ribosomal protein S9   |
| AEW60556.1 | 6  | ABC transport system ATP-binding component   |
| AEW60201.1 | 6  | D-alanyl-D-alanine carboxypeptidase DacA   |

|            |  |
|------------|--|
| AEW63840.1 | 6 bifunctional phosphopantethenoylcysteine decarboxylase/phosphopantetheate synthase |
| AEW62643.1 | 6 putative NAGC-like transcriptional regulator                                       |
| AEW60135.1 | 6 ferrienterobactin receptor   |
| AEW61659.1 | 5 transcriptional regulator SlyA   |
| AEW63160.1 | 5 tRNA (guanine-N(7))-methyltransferase  |
| AEW61505.1 | 5 hypothetical protein KPHS_28070  |
| AEW62318.1 | 5 putative ATP-binding protein   |
| AEW60143.1 | 5 iron-enterobactin transporter periplasmic binding protein                          |
| AEW63998.1 | 5 phosphate transporter subunit  |
| AEW60575.1 | 5 Acylphosphatase  |
| AEW60136.1 | 5 enterochelin esterase  |
| AEW58910.1 | 5 thiamine-phosphate pyrophosphorylase   |
| AEW61934.1 | 5 4-diphosphocytidyl-2-C-methyl-D-erythritol kinase                                  |
| AEW62685.1 | 5 23S rRNA pseudouridine synthase D  |
| AEW63681.1 | 5 cell division protein FtsE   |
| AEW58777.1 | 5 ATP-dependent protease peptidase subunit   |
| AEW63976.1 | 5 chromosomal replication initiator protein DnaA                                     |
| AEW63832.1 | 5 putative glycosyltransferase   |
| AEW59142.1 | 5 putative cell envelope opacity-associated protein A                                |
| AEW63410.1 | 5 tRNA pseudouridine synthase B  |
| AEW58752.1 | 5 repressor of galETK operon   |
| AEW60530.1 | 5 condesin subunit B   |
| AEW59642.1 | 5 DL-methionine transporter substrate-binding subunit                                |
| AEW63262.1 | 5 DNA topoisomerase IV subunit A   |
| AEW62862.1 | 4 hemin ABC superfamily ATP binding cassette transporter, binding protein HmuT       |
| AEW63500.1 | 4 DNA-binding protein Fis  |
| AEW61184.1 | 4 dithiobiotin synthetase  |
| AEW61869.1 | 4 hypothetical protein KPHS_31710  |
| AEW59124.1 | 4 23S rRNA (guanosine-2'-O)-methyltransferase  |
| AEW62986.1 | 4 N-acetylmuramoyl-L-alanine amidase   |
| AEW59498.1 | 4 LacI family transcriptional regulator, fructose operon transcriptional repressor   |
| AEW59378.1 | 4 16S ribosomal RNA m2G1207 methyltransferase  |
| AEW59772.1 | 4 recombination associated protein   |
| AEW62777.1 | 4 ribonucleotide-diphosphate reductase alpha subunit                                 |
| AEW60170.1 | 4 hypothetical protein KPHS_14720  |
| AEW63283.1 | 4 bifunctional heptose 7-phosphate kinase/heptose 1-phosphate adenyltransferase      |
| AEW58911.1 | 4 thiamin biosynthesis protein for pyrimidine moiety                                 |
| AEW60383.1 | 4 putative transport system ATP-binding component                                    |
| AEW61853.1 | 4 phospho-2-dehydro-3-deoxyheptonate aldolase  |
| AEW58716.1 | 4 serine/threonine protein kinase RdoA   |
| AEW63864.1 | 4 ATP-dependent DNA helicase RecG  |

|            |  |
|------------|--|
| AEW59411.1 | 4 homoserine kinase  |
| AEW62664.1 | 4 GTP-binding protein LepA   |
| AEW62355.1 | 4 ferric iron-catecholate outer membrane transporter                                     |
| AEW60544.1 | 4 nicotinate phosphoribosyltransferase   |
| AEW60649.1 | 4 Dihydroorotase   |
| AEW60161.1 | 4 methylthioribose kinase  |
| AEW60328.1 | 3 dithiobiotin synthetase  |
| AEW60284.1 | 3 quinolinate synthetase   |
| AEW58915.1 | 3 endonuclease V   |
| AEW62061.1 | 3 ProP effector  |
| AEW59794.1 | 3 S-adenosylmethionine:tRNA ribosyltransferase-isomerase                                 |
| AEW61679.1 | 3 riboflavin synthase subunit alpha  |
| AEW60147.1 | 3 2,3-dihydro-2,3-dihydroxybenzoate dehydrogenase  |
| AEW59335.1 | 3 4-hydroxyphenylacetate catabolism  |
| AEW61865.1 | 3 integration host factor subunit alpha  |
| AEW59764.1 | 3 hypothetical protein KPHS_10660  |
| AEW60849.1 | 3 short-chain dehydrogenase/reductase SDR  |
| AEW61939.1 | 3 peptidyl-tRNA hydrolase  |
| AEW60154.1 | 3 putative alcohol dehydrogenase   |
| AEW62352.1 | 3 GTP cyclohydrolase I   |
| AEW63964.1 | 3 hypothetical protein KPHS_52660  |
| AEW60098.1 | 3 putative aminotransferase  |
| AEW60859.1 | 3 bifunctional indole-3-glycerol phosphate synthase/phosphoribosylanthranilate isomerase |
| AEW59902.1 | 3 inosine-guanosine kinase   |
| AEW61823.1 | 3 cysteine desulfurase activator complex subunit SufB                                    |
| AEW61142.1 | 3 hypothetical protein KPHS_24440  |
| AEW63253.1 | 3 cystathionine beta-lyase   |
| AEW61658.1 | 2 MarR family transcriptional regulator, transcriptional regulator for hemolysin         |
| AEW60200.1 | 2 hypothetical protein KPHS_15020  |
| AEW59806.1 | 2 transcription antitermination protein NusB   |
| AEW58882.1 | 2 FMN reductase  |
| AEW59807.1 | 2 thiamine monophosphate kinase  |
| AEW63331.1 | 2 hypothetical protein KPHS_46330  |
| AEW61876.1 | 2 transport protein TonB   |
| AEW60629.1 | 2 transcriptional regulator, LysR family   |
| AEW61859.1 | 2 putative outer membrane lipoprotein  |
| AEW60868.1 | 2 short chain dehydrogenase  |
| AEW62046.1 | 2 PTS enzyme IIC, mannose-specific   |
| AEW61761.1 | 2 gluconate 2-dehydrogenase subunit gamma  |
| AEW60470.1 | 2 arginine transporter permease subunit ArtQ   |
| AEW59374.1 | 2 ferric iron reductase involved in ferric hydroximate transport                         |
| AEW62267.1 | 2 polysaccharide export protein  |
| AEW60572.1 | 2 Methyltransferase  |

|            |  |
|------------|--|
| AEW63653.1 | 2 regulator of gluconate (gnt) operon  |
| AEW64014.1 | 2 tRNA uridine 5-carboxymethylaminomethyl modification enzyme GidA                                 |
| AEW60327.1 | 2 biotin biosynthesis protein BioC   |
| AEW59479.1 | 2 thiamin/thiamine pyrophosphate ABC superfamily ATP binding cassette transporter, binding protein |
| AEW59895.1 | 2 DNA polymerase III subunits gamma and tau  |
| AEW60654.1 | 2 hypothetical protein KPHS_19560  |
| AEW58974.1 | 2 replicative DNA helicase   |
| AEW60330.1 | 2 excinuclease ABC subunit B   |
| AEW60387.1 | 2 hypothetical protein KPHS_16890  |
| AEW62290.1 | 2 putative protease  |
| AEW60848.1 | 2 catecholate siderophore receptor Fiu   |
| AEW62503.1 | 2 Aminotransferase   |
| AEW60555.1 | 2 23S rRNA m(2)G2445 methyltransferase   |
| AEW59103.1 | 2 ribosome-associated GTPase   |
| AEW63586.1 | 2 putative ATP-binding component of a transport system   |
| AEW63683.1 | 2 RsmD family RNA methyltransferase  |
| AEW60308.1 | 2 putative regulatory protein MerR   |
| AEW60736.1 | 2 hypothetical protein KPHS_20380  |
| AEW63329.1 | 2 putative enzyme, ferredoxin reductase-like, FAD-linked, NADP-linked                              |
| AEW62997.1 | 2 thymidylate synthase   |
| AEW58722.1 | 2 coproporphyrinogen III oxidase   |
| AEW63637.1 | 2 glycogen phosphorylase   |

\* according to database GenBank accession: GCF\_000240185.1

**Supplementary Table 2. Exclusive proteins identified after 4 hours of native serum exposure.** Proteins were identified using the software Patternlab, module AAPV, using a probability 0.01.

| Accession number * | Spectral count | Protein identification   |
|--------------------|----------------|--|
| AEW63837.1         | 82             | 50S ribosomal protein L33  |
| AEW63998.1         | 61             | phosphate transporter subunit  |
| AEW62851.1         | 35             | iron ABC superfamily ATP binding cassette transporter, binding protein |
| AEW63565.1         | 17             | 50S ribosomal protein L4   |
| AEW60387.1         | 17             | hypothetical protein KPHS_16890  |
| AEW63781.1         | 11             | putative outer membrane protein  |
| AEW61158.1         | 10             | sensory histidine kinase DcuS  |
| AEW63547.1         | 8              | 50S ribosomal protein L15  |
| AEW62760.1         | 8              | aldehyde dehydrogenase B (lactaldehyde dehydrogenase)                  |

|            |  |
|------------|--|
| AEW60177.1 | 6 sugar ABC transporter, periplasmic sugar-binding protein   |
| AEW61842.1 | 6 5-methyltetrahydropteroylglutamate-homocysteine methyltransferase                                |
| AEW59411.1 | 6 homoserine kinase  |
| AEW58911.1 | 6 thiamin biosynthesis protein for pyrimidine moiety   |
| AEW59722.1 | 6 phosphonate ABC superfamily ATP binding cassette transporter, binding protein                    |
| AEW61235.1 | 6 5-methyltetrahydropteroylglutamate--homocysteine methyltransferase                               |
| AEW63575.1 | 6 30S ribosomal subunit protein S12  |
| AEW58782.1 | 5 50S ribosomal protein L31  |
| AEW60492.1 | 5 translation initiation factor IF-1   |
| AEW62691.1 | 5 phospho-2-dehydro-3-deoxyheptonate aldolase  |
| AEW61869.1 | 4 hypothetical protein KPHS_31710  |
| AEW59479.1 | 4 thiamin/thiamine pyrophosphate ABC superfamily ATP binding cassette transporter, binding protein |
| AEW59378.1 | 4 16S ribosomal RNA m2G1207 methyltransferase  |
| AEW61628.1 | 4 alcohol dehydrogenase class III  |
| AEW61679.1 | 4 riboflavin synthase subunit alpha  |
| AEW62061.1 | 4 ProP effector  |
| AEW62916.1 | 4 putative ABC-type transport protein  |
| AEW60614.1 | 4 hypothetical protein KPHS_19160  |
| AEW59410.1 | 4 bifunctional aspartokinase I/homoserine dehydrogenase I  |
| AEW63467.1 | 3 30S ribosomal protein S9   |
| AEW62466.1 | 3 lysine-arginine-ornithine-binding periplasmic protein  |
| AEW59490.1 | 3 isopropylmalate isomerase small subunit  |
| AEW60154.1 | 3 putative alcohol dehydrogenase   |
| AEW59650.1 | 3 UbiE/COQ5 family methyltransferase   |
| AEW59152.1 | 3 hypothetical protein KPHS_04540  |
| AEW59058.1 | 3 putative DSBA oxidoreductase   |
| AEW60544.1 | 3 nicotinate phosphoribosyltransferase   |
| AEW63399.1 | 3 hypothetical protein KPHS_47010  |
| AEW59027.1 | 3 ribose ABC superfamily ATP binding cassette transporter, binding protein                         |
| AEW60145.1 | 3 enterobactin synthase subunit E  |
| AEW63089.1 | 2 flavodoxin   |
| AEW62685.1 | 2 23S rRNA pseudouridine synthase D  |
| AEW63545.1 | 2 30S ribosomal protein S13  |
| AEW58910.1 | 2 thiamine-phosphate pyrophosphorylase   |
| AEW59642.1 | 2 DL-methionine transporter substrate-binding subunit  |
| AEW58926.1 | 2 homoserine O-succinyltransferase   |
| AEW62862.1 | 2 hemin ABC superfamily ATP binding cassette transporter, binding protein HmuT                     |
| AEW58995.1 | 2 periplasmic maltose-binding protein  |
| AEW59659.1 | 2 acyl-CoA dehydrogenase   |
| AEW63671.1 | 2 high-affinity branched-chain amino acid ABC transporter periplasmic substrate-binding protein    |
| AEW58927.1 | 2 malate synthase  |

|            |   |   |
|------------|---|---|
| AEW59806.1 | 2 | transcription antitermination protein NusB                    |
| AEW60649.1 | 2 | dihydroorotase  |
| AEW61250.1 | 2 | putative oxidoreductase                                       |
| AEW61325.1 | 2 | 6-phosphogluconate dehydrogenase                              |
| AEW60146.1 | 2 | 2,3-dihydro-2,3-dihydroxybenzoate synthetase, isochroismatase |
| AEW61220.1 | 2 | beta-lactamase SHV-11   |
| AEW58928.1 | 2 | isocitrate lyase  |

\* according to database GenBank accession: GCF\_000240185.1