

# Chromatin accessibility and microRNA expression in nephron progenitor cells during kidney development

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**Manuscript Source:** <https://www.biorxiv.org/content/10.1101/2021.03.05.434138v1>

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- Always consult the original research paper as the true reference source for the text.

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### Research Paper Sections:

The sections of the research paper input text parsed in this audit.

[illegible]

Title      **Chromatin accessibility and microRNA expression in nephron progenitor cells during kidney development**

**S1 [001]      Abstract**

**S1 [002]**      Mammalian nephrons originate from a population of nephron progenitor cells (NPCs), and it is known that NPCs' transcriptomes change throughout nephrogenesis during healthy kidney development.

Mammalian nephrons originate ...  
... from a population ...  
... of nephron progenitor cells ...  
... (NPCs), ...  
... and it is known ...  
... that NPCs' transcriptomes change ...  
... throughout nephrogenesis ...  
... during healthy kidney development.

**S1 [003]**      To characterize chromatin accessibility and microRNA (miRNA) expression throughout this process, we collected NPCs from mouse kidneys at embryonic day 14.5 (E14.5) and postnatal day zero (P0) and assayed cells for transposase-accessible chromatin and small RNA expression.

To characterize chromatin accessibility ...  
... and microRNA ...  
... (miRNA) ...  
... expression ...  
... throughout this process, ...  
... we collected NPCs ...  
... from mouse kidneys ...  
... at embryonic day 14.5 ...  
... (E14.5) ...  
... and postnatal day zero ...  
... (P0) ...  
... and assayed cells ...  
... for transposase-accessible chromatin ...  
... and small RNA expression.

**S1 [004]**      We observe 46,374 genomic regions of accessible chromatin, with 2,103 showing significant changes in accessibility between E14.5 and P0.

We observe 46,374 genomic regions ...  
... of accessible chromatin, ...  
... with 2,103 showing significant changes ...  
... in accessibility ...  
... between E14.5 ...  
... and P0.

**S1 [005]** In addition, we detect 1,104 known microRNAs, with 114 showing significant changes in expression.

In addition, ...  
... we detect 1,104 known microRNAs, ...  
... with 114 showing significant changes ...  
... in expression.

**S1 [006]** Genome-wide, changes in DNA accessibility and microRNA expression highlight biological processes like cellular differentiation, cell migration, extracellular matrix interactions, and developmental signaling pathways such as Notch.

Genome-wide, ...  
... changes ...  
... in DNA accessibility ...  
... and microRNA expression highlight biological processes like cellular differentiation, ...  
... cell migration, ...  
... extracellular matrix interactions, ...  
... and developmental signaling pathways ...  
... such as Notch.

**S1 [007]** Furthermore, our data identify novel candidate cis-regulatory elements for Eya1 and Pax8, both genes with a role in NPC differentiation; we also associate expression-changing microRNAs, including let-7-5p, miR-125b-5p, miR-181a-2-3p, and miR-9-3p, with candidate cis-regulatory elements.

Furthermore, ...  
... our data identify novel candidate cis-regulatory elements ...  
... for Eya1 ...  
... and Pax8, ...  
... both genes ...  
... with a role ...  
... in NPC differentiation; ...  
... we also associate expression-changing microRNAs, ...  
... including let-7-5p, ...  
... miR-125b-5p, ...  
... miR-181a-2-3p, ...  
... and miR-9-3p, ...  
... with candidate cis-regulatory elements.

**S1 [008]** Overall, our data characterize NPCs during kidney development and point out new candidate regulatory elements for genes and microRNA with key roles in nephrogenesis.

Overall, ...  
... our data characterize NPCs ...  
... during kidney development ...  
... and point out new candidate regulatory elements ...  
... for genes ...  
... and microRNA ...  
... with key roles ...  
... in nephrogenesis.

**S2 [010]**    The functional unit of the kidney is the nephron, which serves to filter waste and maintain normal homeostasis of water, acid-base, and electrolytes in the body.

The functional unit ...  
... of the kidney is the nephron, ...  
... which serves ...  
... to filter waste ...  
... and maintain normal homeostasis ...  
... of water, ...  
... acid-base, ...  
... and electrolytes ...  
... in the body.

**S2 [011]**    Nephron number varies widely in humans (typically between two hundred thousand and greater than one million nephrons (Hughson et al., 2003)) and is established prior to birth (birth in humans, approximately post-natal day 2-3 in mice (Hartman et al., 2007; Hinchliffe et al., 1991)).

Nephron number varies widely ...  
... in humans ...  
... (typically ...  
... between two hundred thousand ...  
... and greater ...  
... than one million nephrons ...  
... (Hughson et al., 2003)) ...  
... and is established ...  
... prior to birth ...  
... (birth ...  
... in humans, ...  
... approximately post-natal day 2-3 ...  
... in mice ...  
... (Hartman et al., 2007; ...  
... Hinchliffe et al., 1991)).

**S2 [012]**    Nephrons cannot regenerate after birth, and decreased nephron endowment is associated with an increased risk of chronic kidney disease and hypertension (Bertram et al., 2011).

Nephrons cannot regenerate ...  
... after birth, ...  
... and decreased nephron endowment is associated ...  
... with an increased risk ...  
... of chronic kidney disease ...  
... and hypertension ...  
... (Bertram et al., 2011).

**S2 [013]**    Nephron number, in turn, is in large part determined by a population of cells called nephron progenitor cells (Cebrian et al., 2014a).

Nephron number, ...  
... in turn, ...  
... is ...

... in large part determined ...  
... by a population ...  
... of cells called nephron progenitor cells ...  
... (Cebrian et al., 2014a).

**S2 [014]** During kidney development, one subset of nephron progenitor cells undergo mesenchymal-to-epithelial transitions (MET) to differentiate into cells of the early developing nephron (renal vesicle), while another continues to self-renew.

During kidney development, ...  
... one subset ...  
... of nephron progenitor cells undergo mesenchymal-to-epithelial transitions ...  
... (MET) ...  
... to differentiate ...  
... into cells ...  
... of the early developing nephron ...  
... (renal vesicle), ...  
... while another continues ...  
... to self-renew.

**S2 [015]** In the latter stages of embryonic development, nephron progenitors increase their propensity to differentiate, which gradually depletes their population and marks the eventual cessation of nephrogenesis (Hartman et al., 2007; Rumballe et al., 2011; Volovelsky et al., 2018).

In the latter stages ...  
... of embryonic development, ...  
... nephron progenitors increase their propensity ...  
... to differentiate, ...  
... which gradually depletes their population ...  
... and marks the eventual cessation ...  
... of nephrogenesis ...  
... (Hartman et al., 2007; ...  
... Rumballe et al., 2011; ...  
... Volovelsky et al., 2018).

**S2 [016]** Nephron progenitor differentiation is regulated by a series of coordinated events.

Nephron progenitor differentiation is regulated ...  
... by a series ...  
... of coordinated events.

**S2 [017]** Briefly, Bmp7-pSmad1/5/8 signaling induces the initial exit of self-renewing Cited1+/Six2+ nephron progenitors into a primed Cited1-/Six2+ state, followed by Wnt9b/β-catenin induction of differentiation (McMahon et al., 2016; Rumballe et al., 2011).

Briefly, ...  
... Bmp7-pSmad1/5/8 signaling induces the initial exit ...  
... of self-renewing Cited1+/Six2+ nephron progenitors ...  
... into a primed Cited1-/Six2+ state, ...  
... followed by Wnt9b/β-catenin induction ...  
... of differentiation ...  
... (McMahon et al., 2016; ...  
... Rumballe et al., 2011).

## **End of Sample Audit**

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