## Selective degradation of Aux/IAA proteins modulates plant development

Thomas Vain<sup>1</sup>, Noel Ferro<sup>2</sup>, Deepak Kumar Barange<sup>1</sup>, Qian Ma<sup>1</sup>, Mattias Thelander<sup>3</sup>, Barbora Pařízková<sup>4</sup>, Ondřej Novák<sup>4</sup>, Siamsa M. Doyle<sup>1</sup>, Alexandre Ismail<sup>5</sup>, Per Anders Enquist<sup>6</sup>, Adeline Rigal<sup>1</sup>, Yi Zhang<sup>7</sup>, Malgorzata Langowska<sup>1</sup>, Karin Ljung<sup>1</sup>, Judy Callis<sup>8</sup>, Fredrik Almqvist<sup>9</sup>, Mark Estelle<sup>7</sup>, Laurens Pauwels<sup>10</sup> and Stéphanie Robert<sup>1</sup>

<sup>1</sup>Umeå Plant Science Centre, Department of Forest Genetics and Plant Physiology, Swedish University of Agricultural Sciences, <sup>2</sup>Institute of Physical and Theoretical Chemistry, University of Bonn, <sup>3</sup>Department of Plant Biology, Uppsala BioCenter, Swedish University of Agricultural Sciences and Linnean Centre for Plant Biology in Uppsala, <sup>4</sup>Laboratory of Growth Regulators, Centre of the Region Haná for Biotechnological and Agricultural Research, Faculty of Science of Palacký University & Institute of Experimental Botany CAS, <sup>5</sup>Sup'Biotech, IONIS Education Group, <sup>6</sup>Laboratories for Chemical Biology Umeå, Chemical Biology Consortium Sweden, Department of Chemistry, Umeå University, <sup>7</sup>University of California San Diego and Howard Hughes Medical Institute, <sup>8</sup>University of California, Davis Department of Molecular and Cellular Biology, <sup>9</sup>Umeå University, Department of Chemistry, Umeå University, SE-90187 Umeå, Sweden, <sup>10</sup>Department of Plant Systems Biology, VIB

Auxin phytohormones control most aspects of plant development through a complex and interconnected signaling network. In the presence of auxin, AUXIN/INDOLE-3-ACETIC ACID (Aux/IAA) transcriptional repressors are targeted for degradation by the SKP1- CULLIN1-F-BOX (SCF) ubiquitin-protein ligases containing TRANSPORT INHIBITOR RESISTANT 1/AUXIN SIGNALING F-BOX (TIR1/AFB). Here, we report four small molecules named DEVELOPMENTAL REGULATORS (DRs) requiring AXR1 and SCF<sup>TIR1/AFB</sup> to modulate plant development. Three DR molecules trigger selective auxin responses at transcriptional, biochemical and morphological levels which are explained by their ability to promote the interaction between TIR1 and a specific subset of Aux/IAA proteins. These results demonstrate the potential of selective auxin agonists to reprogram plant development through a selective degradation of the Aux/IAA transcriptional repressors.