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## How electrostatic interactions drive nucleosome binding of RNF168 & PSIP : structural studies and their implications for rational drug design

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# How Electrostatic Interactions Drive Nucleosome Binding of RNF168 & PSIP1

Structural Studies and Their Implications for Rational Drug Design

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Velten Horn

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The research described in this thesis was performed in the Macromolecular Biochemistry Department of the Leiden Institute of Chemistry in Leiden, the Netherlands and the NMR Spectroscopy Research Group of the Bijvoet Center for Biomolecular Research in Utrecht, the Netherlands.



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# List of Abbreviations

<b>6-FAM</b>	5,6-carboxyfluoresceine
<b>AIR</b>	ambiguous interaction restraints
<b>ATP</b>	adenosine triphosphate
<b>Bp</b>	basepairs
<b>BS</b>	binding sequence
<b>CPMG</b>	Carr-Purcell-Meiboom-Gill sequence
<b>Cryo-EM</b>	cryogenic electron microscopy
<b>CSP</b>	chemical shift perturbation
<b>DCM</b>	dichloromethane
<b>DIC</b>	N,N'-Diisopropylcarbodiimide
<b>DSB</b>	double-strand break
<b>DTT</b>	dithiothreitol
<b>EDTA</b>	ethylenediaminetetraacetic acid
<b>EMSA</b>	electrophoretic mobility shift assay
<b>EPR</b>	electron paramagnetic resonance
<b>FMOC</b>	fluorenylmethyloxycarbonyl
<b>FRET</b>	Förster resonance energy transfer
<b>HEPES</b>	4-(2-hydroxyethyl)-1-piperazineethanesulfonic acid
<b>HOAt</b>	1-Hydroxy-7-azabenzotriazole
<b>HSQC</b>	heteronuclear single quantum coherence
<b>IEX</b>	ion exchange
<b>IM-MS</b>	ion-mobility spectrometry–mass spectrometry
<b>IPTG</b>	isopropyl β-D-1-thiogalactopyranoside
<b>KAc</b>	acetylated lysine
<b>Kme</b>	methylated lysine
<b>LC-MS</b>	liquid chromatography–mass spectrometry
<b>MD</b>	Molecular dynamics
<b>Mmt</b>	monomethoxytrityl

<b>MST</b>	microscale thermophoresis
<b>NCP</b>	nucleosome core particle
<b>NMR</b>	nuclear magnetic resonance
<b>NOE</b>	nuclear Overhauser effect
<b>PCS</b>	pseudocontact chemical shift
<b>PPI</b>	protein-protein interactions
<b>PRE</b>	paramagnetic relaxation enhancement
<b>PMSF</b>	phenylmethylsulfonyl fluoride
<b>PTM</b>	post translational modification
<b>SANS</b>	small-angle neutron scattering
<b>SASD</b>	surface accessible distance
<b>SAXS</b>	small-angle X-ray scattering
<b>SDS</b>	sodium dodecyl sulfate
<b>Sph</b>	phosphorylated serine
<b>ssNMR</b>	solid-state NMR
<b>TCEP</b>	tris(2-carboxyethyl)phosphine
<b>TFA</b>	trifluoroacetic acid
<b>TIPS</b>	triisopropylsilane
<b>TOCSY</b>	total correlation spectroscopy
<b>TROSY</b>	transverse relaxation optimized spectroscopy
<b>TSP</b>	trimethylsilylpropanoic acid
<b>Ub</b>	ubiquitin
<b>UPLC</b>	ultra performance liquid chromatography
<b>XL-MS</b>	cross-link-based mass spectrometry