

Supplementary Table 1. General demographic information of human postmortem brain samples. There were no significant differences between heroin addicts and control groups in pH and age. Student's *t*-tests, $p = n.s.$

| GROUP | Cause of Death | SEX | pH | AGE | Blood - Morphine | Blood - ETOH |
|---------|-----------------|-----|------|-----|------------------|--------------|
| HEROIN | heroin | m | 6.07 | 31 | 0.03 | 0 |
| HEROIN | addict:heroin | f | 6.48 | 32 | 0.30 | 0 |
| HEROIN | heroin overdose | m | 6.8 | 28 | 0.25 | 0 |
| HEROIN | heroin overdose | m | 6.66 | 27 | 0.24 | 0 |
| HEROIN | heroin overdose | f | 6.36 | 24 | 0.35 | 0 |
| HEROIN | heroin overdose | m | 6.52 | 20 | 0.74 | 0 |
| HEROIN | heroin overdose | m | 6.68 | 23 | 0.33 | 0 |
| HEROIN | heroin overdose | m | 6.47 | 27 | 0.96 | 0 |
| HEROIN | heroin overdose | m | 6.23 | 21 | 0.4 | 0 |
| CONTROL | cardiac failure | m | 6.33 | 30 | 0.00 | 0 |
| CONTROL | cardiac failure | f | 6.82 | 36 | 0.00 | 0 |
| CONTROL | cardiac failure | f | 6.98 | 31 | 0.00 | 0 |
| CONTROL | sudden death | m | 6.87 | 26 | 0.00 | 0 |
| CONTROL | sudden natural | m | 6.70 | 28 | 0.00 | 0 |

Supplementary Table 2. PCR Primers**Human qPCR primers**

| Target gene | Primer sequence (5' – 3') |
|-----------------------------|---------------------------|
| hs- <i>Rn18s</i> | CGCCGCTAGAGGTGAAATTC |
| hs- <i>Rn18s</i> _rev | CATTCTTGGCAAATGCTTTCG |
| hs- <i>Bdnf</i> exon IX | TTTCTCGTGACAGCATGAGCAG |
| hs- <i>Bdnf</i> exon IX_rev | GTCCTCATCCAACAGCTCTTCTATC |

Rat qPCR primers

| Target gene | Primer sequence (5' – 3') |
|---------------------------|----------------------------|
| <i>Gapdh</i> | GTGGACCTCATGGCCTACAT |
| <i>Gapdh</i> _rev | TGTGAGGGAGATGCTCAGTG |
| <i>Bdnf</i> exon I | AGTTGCTTTGTCTTCTGTAATCGC |
| <i>Bdnf</i> exon I_rev | ATTCACGCTCTCCAGAGTCCCATG |
| <i>Bdnf</i> exon II | TTGGCTTCCTAGCGGTGTAG |
| <i>Bdnf</i> exon II_rev | CGTGGACGTTTGCTTCTTTC |
| <i>Bdnf</i> exon III | CCGGGCTGGATGCTTC |
| <i>Bdnf</i> exon III_rev | TCCAGAGGAAGTGAAAGTTTTGACT |
| <i>Bdnf</i> exon IV | CTGCCTTGATGTTTACTTTGACAAG |
| <i>Bdnf</i> exon IV_rev | ACCATAGTAAGGAAAAGGATGGTCAT |
| <i>Bdnf</i> exon V | CATAACCCCGCACACTCTGT |
| <i>Bdnf</i> exon V_rev | GACCCTCTCTGTCCATCCAG |
| <i>Bdnf</i> exon VI | CAGGAGCGTGACAACAATGTGA |
| <i>Bdnf</i> exon VI_rev | ACCATAGTAAGGAAAAGGATGGTCAT |
| <i>Bdnf</i> exon VII | CCTGAAAGGGTCTGCGGAACTCCA |
| <i>Bdnf</i> exon VII_rev | GAAGTGTACAAGTCCGCGTCCTTA |
| <i>Bdnf</i> exon VIII | GTGTGTGTCTCTGCGCCTCAGTGGA |
| <i>Bdnf</i> exon VIII_rev | GAAGTGTACAAGTCCGCGTCCTTA |
| <i>Bdnf</i> exon IX | GGGTGAAACAAAGTGGCTGT |
| <i>Bdnf</i> exon IX_rev | ATGTTGTCAAACGGCACAAA |
| <i>Nurr1</i> | GAAGATTCTGGCTTTGCTG |
| <i>Nurr1</i> _rev | CTGGGTTGGACCTGTATGCT |

Mouse qPCR primers

| Target gene | Primer sequence (5' – 3') |
|----------------------------|---------------------------|
| m- <i>Gapdh</i> | AACTTTGGCATTGTGGAAGG |
| m- <i>Gapdh</i> _rev | ACACATTGGGGGTAGGAACA |
| m- <i>Bdnf</i> exon IX | CTCAGGCAGAATGAGCAATG |
| m- <i>Bdnf</i> exon IX_rev | AGCCGTCTGTGCTCTTCACT |
| m- <i>Ezh2</i> | ATCTGAGAAGGGACCGGTTT |
| m- <i>Ezh2</i> _rev | TGTGCACAGGCTGTATCCTC |
| m- <i>Creb1</i> | GTGTTACGTGGGGGAGAGAA |
| m- <i>Creb1</i> _rev | GGGCTAATGTGGCAATCTGT |
| m- <i>Nurr1</i> | GACCGGCTCTATGGAGATCA |
| m- <i>Nurr1</i> _rev | CTGGGTTGGACCTGTATGCT |

Supplementary Table 3. Rat qChIP PCR Primers

| Target gene | Primer sequence (5' – 3') |
|----------------------|---------------------------|
| <i>Bdnf</i> -p1 | TTCGATTCACGCAGTTGTTC |
| <i>Bdnf</i> -p1_rev | GCAGCCTCTCTGAGCCAGT |
| <i>Bdnf</i> -p2 | CGTGGAACAAACTTGGGACT |
| <i>Bdnf</i> -p2_rev | CCCTTGCGGATGTCAATTAT |
| <i>Bdnf</i> -p4 | GCGCGGAATTCTGATTCTGGTAAT |
| <i>Bdnf</i> -p4_rev | GAGAGGGCTCCACGCTGCCTTGACG |
| <i>Bdnf</i> -p6 | TGCAGGGGAATTAGGGATAC |
| <i>Bdnf</i> -p6_rev | TCTTCGGTTGAGCTTCGATT |
| <i>Bdnf</i> -eI | CAGTGACAGGCGTTGAGAAA |
| <i>Bdnf</i> -eI_rev | AACGCCCTCATTCTGAGAGA |
| <i>Bdnf</i> -eII | AAAGCGTCTTTTCCGAGGTT |
| <i>Bdnf</i> -eII_rev | CAGCCTACACCGCTAGGAAG |
| <i>Bdnf</i> -eIV | ACTGAAGGCGTGCGAGTATT |
| <i>Bdnf</i> -eIV_rev | AGTCTTTGGTGGCCGATATG |
| <i>Bdnf</i> -eVI | CTGGCAGGCTTTGATGAGAC |
| <i>Bdnf</i> -eVI_rev | CACATTGTTGTCACGCTCCT |
| <i>Th</i> -p | CTAATGGGACGGAGGCCTTT |
| <i>Th</i> -p_rev | GTCCCGAGTTCTGTCTCCAC |
| <i>Grial</i> -p | CTGGGAGAGAAGCAGAGGAA |
| <i>Grial</i> -p_rev | CTGCCAGCTCTCTATGCTGA |
| <i>Nurr1</i> -p | ATCGTCGAGCAAGGGTAGTG |
| <i>Nurr1</i> -p_rev | CCGCCCTTGAAAATATGTCT |

Supplementary Table 4. Rat qChIP PCR Primers for validation

| Target gene | Location (from TSS) | Primer sequence (5' – 3') | Related epigenetic modification |
|-------------------|---------------------|---------------------------|--|
| <i>Actb-1</i> | -250 | TCGCCAAACTCTTCATCCTC | Pol II ⁴⁹⁻⁵¹ , H3K36me ³⁵¹ |
| <i>Actb-1_rev</i> | -52 | GCCATAAAAGGCAACTTTTCG | |
| <i>Actb-2</i> | 1639 | CGATCGCCTTTCTGACTAGG | Pol II ⁴⁹⁻⁵¹ , H3K36me ³⁵¹ (Within intron3) |
| <i>Actb-2_rev</i> | 1808 | GCTGGGAGTATCAGGACAGG | |
| <i>Actb-3</i> | 2770 | GATCATTGCTCCTCCTGAGC | Pol II ⁴⁹⁻⁵¹ , H3K36me ³⁵¹ (Within exon6) |
| <i>Actb-3_rev</i> | 2922 | AGTCCGCCTAGAAGCATTTCG | |
| <i>Gapdh</i> | -156 | TGAGAGAGGCCCAGCTACTC | Pol II ⁵² , acH3 ⁵² , acH4 ⁵² , H3K4me ³⁵³ , H3K36me ³⁵³ |
| <i>Gapdh_rev</i> | -4 | AGGGCTGCAGTCCGTATTTC | |
| <i>Rpl30</i> | 498 | GGAAAGTACGTGCTGGGGTA | acH3 ⁵⁴ , H3K4me ³⁵⁵ (Within exon3) |
| <i>Rpl30_rev</i> | 650 | TTCTTCCCAATGCTGGAATC | |
| <i>Myod1</i> | -115 | CAAGCTCCGCCCTACTACAC | H3K27me ³⁵² , EZH2 ⁵⁶ , SUZ12 ⁵⁷ |
| <i>Myod1_rev</i> | 49 | TGTGGTGAAGAAAGCAGTCG | |
| <i>Afm</i> | 444 | TGTACACACTCACACACAAACACA | H3K9me ²⁵² , H3K27me ³⁵² , G9a ⁵⁸ (Within intron1) |
| <i>Afm_rev</i> | 632 | GCTTTGCTCTTGCGTAGTC | |
| <i>Myt1-1</i> | -184 | GATGGGAAAGTGGATCTGGA | H3K9me ³⁵⁹ , H3K27me ³⁵⁹ |
| <i>Myt1-1_rev</i> | -26 | GGGCCAGGAAGGATAGTCTC | |
| <i>Myt1-2</i> | -1903 | TCCCCTTTAGAGGAGGCACT | H3K9me ³⁵⁹ , H3K27me ³⁵⁹ |
| <i>Myt1-2_rev</i> | -1732 | CCGGGAAAGGGAATAACATT | |
| <i>Hoxa2</i> | -165 | CCCCATACGGCTGTAATCAG | H3K27me ³⁶⁰ , RING1A ⁶¹ , RING1B ⁶² , BMI1 ⁶³ |
| <i>Hoxa2_rev</i> | 10 | CGTAATTCATGGCCTTCTCC | |
| <i>Esr1</i> | -203 | GGGGGCTGGAGTTTCTTCTA | mSIN3a ⁶⁴ |
| <i>Esr1_rev</i> | 10 | TATGTGGAGTGGCAGGGACT | |
| <i>Nr4a3</i> | -674 | GGGCCAGAGAGACGTAAACA | CREB ⁶⁵ Containing CREB binding motif |
| <i>Nr4a3_rev</i> | -524 | CTTCCGAAAGGGAAGGCTAC | |
| <i>Bdnf</i> | 1219 | CGCACGTGACAAAACGTAAG | NURR1 ³² *Containing NBRE (NGFI-B response element)-like sequence ³⁰ -Within exon2 |
| <i>Bdnf_rev</i> | 1294 | CTACCACCTCGGACAAATCC | |
| <i>Pitx3</i> | -1293 | GGTGGGAAGTCCTGAAGGT | NURR1 ³⁰ *Containing NBRE (NGFI-B response element)-like sequence ³⁰ |
| <i>Pitx3_rev</i> | -1095 | ATTTGCAAGGGACAGGACAG | |
| <i>Th</i> | -1055 | AGGAAATGACTCCAGGGACA | NURR1 ⁶⁶ *Containing NBRE (NGFI-B response element)-like sequence ³⁰ |
| <i>Th_rev</i> | -865 | CATGGTACCCCGAGGTCTAA | |

* Putative Nurr1 binding motif, 5'-AAAGGTCA-3' or one base deviation