

Oligodendrocyte Lineage and the Motor Neuron Connection

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MRC L, M, C, L, C, B, L, D, K, B

KEY WORDS: a; r r ; ; c a

ABSTRACT: OL r r r c c r a b a b
a r c (OL) r a r r r c r b r c
r a b . T a r r b ca a a OL ar r a a r
c r a r , r a a r a c a r r c r r b
r r c . W a a ab ar OL a a
a a c (r r) c a r a r c a r r
(MN) a r a . T a ca r a ab a
a , c a , a a . I ar c r c r a ar

S c a r r a a b r r a c r -
 a ac r a r a b , c Pa 6.
 S r r , 6 RNA a , 6
 ab r r a - a c r r [S]
 , a r b a a r a - - r a (-
) c a c a r a r a r a r
 a c r . . , a r r c a r -
 r a S (E r c a . , 1997). Pa 6
 ar r a c ar ac r a r r
 a a r a r a b [S] a c r
 a a r r a c a c . I b
 r a a r r a a r a a a c a c a -
 r a - a a . F r a , r c a c a -
 r a Pa 6 ar c r r r r
 2.2, c r r c a Pa 6-
 a a a ac r a (E r c a
 a . , 1997). O r r a c r a c r a (. . , 3,
 2, 6.1) ar r a a a a
 a r a r a b r
 a ar a r c r Pa 6 (F . 1)
 (B r c a E r c a , r).
 T a a a b
 r a a r a c (F . 1). N r a
 r c r r a , 6 a , 2.2- r a
 a ac a r a r c r a r
 a r (MN) a a r a a c a a r a -
 a c ar a a a a a
 c a r I, r bra
 D . I a I . I bra ,
 MN c a a r r a , c a r a
 ar . I c r ca a c r a c r a c
 r a r , c a r a a r a .
 T r r r a r r
 r ar 6 r a a r a a c
 r r (MN). I bra r c
 a a ac a c (. . , a MN
 a a r a) a c r ca a
 c r a a a c a c
 r bra c . A b (bra c a a bar),
 r ar a a r a a r c
 b c .
 F r r r a , r r a
 V2 a V1 a r (IN) (F . 1). T a
 r a MN a IN r r ar r
 a c a a r a a r a
 r a a r r c ar r ac ar
 r a a r a a r a a c r
 a bra . F r a , MN r -
 a ac r I 6.14 I 9. (II 60366) . C 62 09 566 69 . . 2110 92 29366 29366 () - 9 29366 (1992 0) - 23366 X

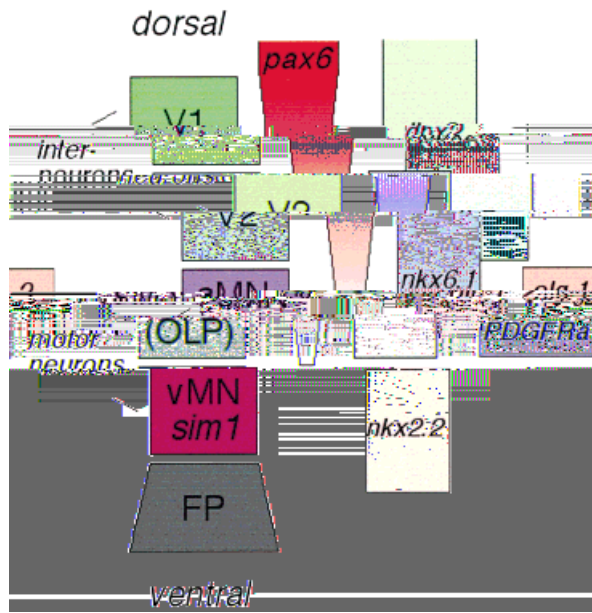


Fig. 1. Dorsal view of the brain showing gene expression patterns. The diagram is oriented with dorsal at the top and ventral at the bottom. Key regions and genes labeled include: V1, pax6, V2, V3, nkx2.2, VMN, (OLP), motor neurons, FP, nkx6.1, and PDGFRA. Other labels include inter-BURRO, nky2, 2, 3, 4, 5, 6, 7, 8, 9, 10, 11, 12, 13, 14, 15, 16, 17, 18, 19, 20, 21, 22, 23, 24, 25, 26, 27, 28, 29, 30, 31, 32, 33, 34, 35, 36, 37, 38, 39, 40, 41, 42, 43, 44, 45, 46, 47, 48, 49, 50, 51, 52, 53, 54, 55, 56, 57, 58, 59, 60, 61, 62, 63, 64, 65, 66, 67, 68, 69, 70, 71, 72, 73, 74, 75, 76, 77, 78, 79, 80, 81, 82, 83, 84, 85, 86, 87, 88, 89, 90, 91, 92, 93, 94, 95, 96, 97, 98, 99, 100.

c r r S .I a r -
 a r r a a a c r r a c c
 a r r a a r a a r a c a r
 c r a r r c r c b a S ,
 a r a c a c r a c , MN ,
 OL ca b c c ca (R a ., 1994;
 Pr a ., 1996; P a ., 1996; Or a a .,
 1999). H r c c a r a S c r a
 c a , r c c a r a a r -
 a MN a OL . T S r MN
 r a a r OL (Pr a ., 1996; Or a
 a ., 1999), c c a a S
 a c a c MN-OL r c r c (. ., a ar
 a a). O r r r a a b . F r -
 a , a r r r MN b
 r r a c r ac a a ar a OL
 r c r r a r a r a r ca r a -
 . H r , a r r a MN
 b b a r r OL b ca OL
 a ar a r a a a a c r r I / -I
 a c , a c MN a (S a .,
 1998). I c b ar a MN, ar
 r a a I / -I c r , a r a r
 a a r MN a c OL r c . H r ,
 ac a PDGFR α OL P a ar a
 ar MN r c a c a a MN r r r
 N r , ca r MN-OL a c a .
 a c c . T r ar r b . F r -
 a , ar a MN a OL r c r r c
 -b - a r r a r r -
 , OL r c r r r a r r
 MN r c a a b c a ac a r a
 a r a c . T c a -
 a r b a ar MN-OL a , c
 a MN a OL br a c r ar r
 a a - c (F . 2). I
 c ar a r r r - a , a -
 a a c r c r . M r c r r a . (1998)
 -a c r c c r a a
 r a a c r , ar r r .

**MIXED NEURON-GLIAL LINEAGES
IN OTHER SYSTEMS**

C a a r - a r c r r c c r a r -
 bra a a r ar r bra a r
 . L D . , c r a a CNS r b a r -
 a r r r , a . F r a , r r -
 b a NB1 1 r r a a r c (c)
 a r a a a c a r
 c (GM1, GM2, c.) (F . 2). L ab a a ,
 GM1 a r a a r r (aCC) a a -
 r (CC) b a GM a r a b r -
 r a (a a) a a a r a (B
 a ., 1996). T , a r b a ca b a
 c r a a r a , a

MN/OL r c r r a a a
 c r . H r , a D . a ar r
 c - . L a a a r r
 a ca b a a r r r r .
 T acc r a r r a ar ac r c
 r a r c a b a r a -
 r c (F . 2).
 L r bra r a , a r a a r a , a
 a M r a , ar r r a r c r r
 a a a r a a r a ,
 r a a r ca a a r a
 r r r r a ar r r a
 c - a a r D . NB1 1 r b a .
 D r c ar r a / a r b a .
 r a a r ar , r r c r a ,
 r a r b r a
 c a a c - a .
 L c r bra c r , ca r r c a
 a r a c a a c (r a a r a
 r a r c , a) a b a a (Pa a a
 a ., 1991; Gr a ., 1993), b a
 r c r r a a r a b r a (W -
 a a ., 1991; Da a T , 1994). Pr -
 ab , a a r ca r c r -
 (ar a c ar r r a a ar r ,
 r a) . A a , a r r r
 ar r a a r a a a r r r
 r r a c - a a c
 a cc a . H r , r c a -
 a a c r ca c a c r
 a r a c r b a r c a
 (Q a ., 1998) a c a r - a a
 (S. T , r a c a a).

**A SINGLE OLIGODENDROCYTE LINEAGE
IN THE SPINAL CORD AND BRAINSTEM**

T r a b a c r r r c a r r
 r , r r a , OL a (S a
 a ., 1999).
 T a a c r r a a OL a
 c r c ar a OL a
 ar r : PDGFR α a r r
 PLP/DM-20 . L a a c r , PLP/DM-20
 r a a r c a r a VZ
 a r E14.5 (T a ., 1995). T r c
 r PDGFR α c r a ar 2 a ar r a
 E12.5. T r , r ac a E14.5

PDGFRα c a b r, c r ra
 Br U, a r r (Ca r a.,
 1998), c OL r r
 c b a . I , ar OL r -
 r b ca ca b - r a a
 a b a a r ac ar a *PDGFRα*
 a ac c r , a a r a OL (Ha
 a ., 1996). C *PDGFRα* c
 c r ra a c r c ra a ca r c
 a b r OL a r (Ha a .,
 1996). M r r, ar a 10% r a -
 b r OL (a r) a
 c r c a ar r *PDGF-A*
 (Fr r a ., 1999). T r
 a r a r *PDGFRα* c ar a r,
 r bab a c OL a a c r .
 W a *PLP/DM-20* c ? S a ar
 b - c, r r r ar r
 a r OL (. ., CNP, MBP) (P r a .,
 1997), a
 ar c , r c -b ar c , ar
 r a , a a OL a r
 r a b r a a a r -
 a b ar b r . I b a
 r a a br " a " r a a -
 a ; , c r a b cr b a -
 c OL r r . A r a , a
 ca a c a a ab . I
 ca , ar r r c r a ,
 ca a a a c a c (a b-
 b r a r WDR' ab).
 A ab ar a a a
 r r *PLP/DM-20* c a a ar a
 VZ bra a r E12.5 (Fr r a .,
 1999). W c a ar r *PLP/DM-20*
 c a a c r a bra ar r r
 c a a , b ar r b ar
 r , r a b a a OL . I
 r a OL a a c r a bra
 a r a ca *PDGFRα*-
 r r MN-OL a cr b ab .
 T a b c acc (S a a ., 1999).
 H r, r a a . A ar r
 r *PLP/DM-20*- c c ab ,
 r a r c r a
 a a c r r *PLP/DM-20* (a)
 a ar a E12 ra (Y a ., 1994) b r
 a ar c *PDGFRα* E14. A ar
 b r . T *PLP/DM-20* c r -
 ab ar r c r r a r a r a ,
 a, r b . I b a ar a r a
 (r c r r) a r- r *PDGFRα*
 r r r VZ. I , *PLP/DM-20* c a
PDGFRα c c b b OL r c r r , b a
 cc a a a a (F .3). T a
 r r a r r a
 (. ., MN) a r r a *PLP/DM-20*,
 r a OL r a . A ar a
 b a b r b P r V a a . (1999).

OLIGODENDROGENESIS IN THE FOREBRAIN: ONE OR MORE LINEAGES?

T r ar a *PDGFRα* - c a r a
 ar a c a a (S a a ., 1998). T
 r a ar a VZ b a a a a c
 a a c (r - c ar a) ar a E12 ra (E10.5
) (Pr a R c ar a ., 1993; S a
 a ., 1998). T a a c r a a b r r a
 r a a a a a ,
 a ar a a a c r a r E16
 (ra) (Pr a R c ar a ., 1993). S ar *PDGFRα*
 c ar r a c c , a r ab c r r -
 a a b cr b a c c b O a . (1997). T
 b a r *PDGFRα* c r bra
 r a r a c *PDGFRα* OLP a a
 c r a b r r a a r a
 ar a a r . I ac , a c
 c r r a r bra a a
 ar OLP a r ar CNS. T a
 r a a ar a S r a a ra
 a c a a , a b c S ac a a -
 b ra OL a c r c c r c a b -
). T r r , r a ar b r a a
 b a OL a a r bra a a
 a a c r .
 I acc a MN a OL ar a r a
 a a c r OL r bra b
 a r a a a c r ar MN
 r bra . H r, b a OL
 r bra ar
 a ar

r br b a b OL OL a
PDGFRα 62629.7(62)-4T ((62)-4 ((62)-4c r r br362)-43(O.2

c (Fr r a., 1999), a a a
 PDGFR α PDGFR α - r r r . T
 . W a a PDGFR α r r
 r a ar a PDGF-A c , -
 ra a PDGFR α c
 ab PDGF-A (Fr r a., 1999). T
 a b a b PDGF-BB ar a
 b r (a a ab
 a), r b a - c r r r PDGF¹
 r b r /r c r c b a a r -
 a PDGF. I a , ac a r ar
 PDGFR α c r a PDGF-A c a
 b a PDGFR α c ar
 r b r a ar a PLP/DM-20 c
 a c . I ac , a ar
 CNS a a r a b r PDGFR α c a
 a c (. . , a) r a r a
 r a b r PLP/DM-20 c a
 a , c a a a a
 PLP/DM-20 c a ar r PDGFR α
 r r (Fr r a., 1999).
 N a PLP/DM-20 r a c ar a
 a a r a a bra a a r ar
 a ar a E9 ba a a
 a c a , r a (T a., 1992). T
 ar r a a a ab r
 a a , ca r PLP/DM-20 c a a ar
 a r VZ. T c c a a c a a
 a a a . A a r a a c r ,
 PLP/DM-20 a r a c b -
 ra r c r r r . J a b r ar
 a ar a c , ar r bab a ; a b
 ar r r r , ar
 a a ar r r c a r r c-
 bra r . I b a r
 OL a r c , a c a r a . I a a
 PLP/DM-20 a r a c a
 bra c b r ar a OL r c r r , b ar
 a ra OL . W a
 PLP/DM-20 a r a c
 a bra r PDGFR α - OLP , a
 ar r r a a c r (F .3). P r a
 r a r a a c r a r bra
 r ar a a .

**POSTNATAL OLIGODENDROGENESIS
 IN THE FOREBRAIN**

W a a c r OL a a
 br c CNS. G a a c a a
 a r r r c r r c a ca a ra
 a r , OL , a a r c , a b r c ar
 a a a r r bra (L a a

¹T r c a a c r "PDGF-C" a c . T PDGFA
 PDGFR α PDGFR α r r r . T PDGFA
 c c r PDGF-A/PDGF-B b - c (C. B , r a a
 c a ca) : r a , r ar r ar ac a a
 c a r c r a a a ar b a c
 T a ca r c r a c a r a c r PDGFR α .

G a , 1993, 1997). I a c ar , r ,
 r c r r ar r a a a bra r ca-
 a a a a a bra c a r -
 a c . T a b r a (r
 ca) a ra c a r a
 CNS r a . H br c r c r-
 r , a a r c r r , a a c ar
 c a c a a r a r r .

**EVOLUTIONARY IMPLICATIONS
 OF THE MN-OL CONNECTION**

I c r a OL a c r a a
 MN , a OL a r r c r a a -
 a ra r a a a . W a
 b r a a a c a c a b a MN
 OL r c a a c r a a (R c -
 ar a ., 1997); r a OL a a
 ac r a r MN a
 b r . T c a c r r a c a a a
 a a b a c a c r r a ca
 r r a r , r a . T c a a
 OL a a c r b r a c a -
 ca a c (a), c ra (a c /
 ara a c), MN . I a a
 a a r a a ca a a r r r
 ra ca r (R , 1993; Da a ., 1999).
 M r , a (M), c a a (r
 a , r a), a a c a , a ra a
 a b acc ra a ca r (WDR, a b-
 b r a a).

CONCLUSION

T r r a a c a OL a a
 c r a bra a a r PDGFR α r -
 r a a ar a a ra r
 a c r a a c r r a . F r r
 a r r , a a c ar, b r ar
 b a a b a r a a
 r bra a a a c r . I r a a ab
 a c ca b r r r r a a
 a r a a b , a ab c
 c a c a c ar , r r r a
 c a a a br a c OL a a
 (r a a ar a a) a a r r - r r -
 a ra .

ACKNOWLEDGMENTS

D a r a a ba ca a c ar a
 a r a a (a r a), a a a a
 c c a c r c . F r a r
 a r a a ra a a ,

ca (c r r c a) B r Z a c , J a -
 L a T a , K a I a a a , B b M r , a r
 c a , a a r r b c a . T a a r -
 a c a r c a a r c a
 T r r a a r a . S a c c a r a a
 a r a r a a a a
 c a c a . W r W D R ' a b r a r r b
 UK M ca R a r c C c a W c
 Tr .

REFERENCES

B T, U G, D CQ, T c a GM. 1996. T b r c c
 c a r a r a a D r a a a r . I.
 N r b a a r r a a a r c -
 r . D B 179:41 64.
 B r c J, E r c a J. 1999. T c a a r a a b
 r a c . S a r c C D B . 10:353
 362.
 B AM, H b MF, I b r a M, K r S, G r a a A, B r r M. 1997.
 PDGF-a a r c r a b a c r r a RNA a r c
 c - r b r c r a : a b b r a -
 C N r c 8:311 322.
 Ca r AR, Ha AC, Y W-P, Wa FS, H a JK, B C,
 R c a r WD. 1998. O r c a a a c a
 r PDGF . N r a 20:869 882.
 Da AA, T S. 1994. A r a a c a
 b r c r a c r b r a c r . N a r 372:263 266.
 Da AD, W a r b TM, H a r DK, L a PH. 1999. M -
 a a c a a r . N a r 398:571 571.
 E r c J, R a b a P, S c A, B r c r - M r a S, K a a a A, a
 H a a V, J TM, B r c J. 1997. P a 6 c r r a r
 c a a a r a a r a S a a .
 C 90:169 180.
 F c r M, K a r L, H a AC, A b c a A, C a r AR, B r
 H, W K, B r C-H, H a JK, B C, R c a r WD.
 1999. D c r c r c a a a r
 a a a PDGF-A c c c . D 126:457 467.
 G r EA, W a BP, L DQ, H a M, F r c A, P r c J.
 1993. M r c a a b r c c r a c r b r a
 c r . D 117:553 561.
 Ha A, G NA, R c a r WD. 1996. S a c r r c
 a a r c r a - r c r c a a r PDGF
 a a r c r . D 122:4085 4094.
 Ha r RJ, F r c VJ. 1996. O a r c r r a r r -
 a r b r c b r a a , b r a a
 r r c c . D 122:2059 2069.
 Ka a A, H b K, R a MS. 1997. N r a c r
 b r c a a c r : a a , c a r a c r a a , a c a
 a a . D B 186:202 223.
 L b r SM, B r SM, S a JR. 1990. L a a , a r r a a , a
 a c a a r a a r a c c a a c r . J
 N r c 10:2451 2462.
 L b r SM, S a JR. 1995. M r a r a a r a a
 b r c c c a c r . J N r c 15:1236 1248.
 L a SW, G JE. 1993. B r c a r c a a r c
 r r r a b a r c a r a a a r a
 r b r a . N r a 10:201 212.
 L a SW, G JE. 1997. M a a a a r c r c
 r c r c a a a a b r c a r a .
 J N r c R 48:83 94.

M r RH. 1996. O r c r a . T r a N r c 19:92 96.
 Q a K, Y a Y, R a r U, M r RH. 1997. F c a a r c a r
 r a a r a a r a r c r c r c r c r c
 c a r . N r a 19:283 292.
 O r a DM, H a JE, D r KL, M r RH. 1999. S a c
 r a a r r r a a a r c c a c r
 r c r c r c r . D 126:2419 2429.
 P a a a JG, B a J A, F r a E, L a MB. 1991. S a r a
 r r c r c r a a a a a a a r a a
 r a c c a a . C r b r a C r 1:463 468.
 P r V a EM, O r C, S a N, P a C, C c a r P, Z a c B,
 T a J-L, M a S. 1999. E a r c a a r -
 c a c c b r a c b r a . D B (a r).
 P r a F, T S, T a J-L, K a a a T, I a a K, Z a c B. 1997.
 l a r a PLP/DM-20, MBP, a CNP r a b r c
 a a a a a a a a a a a a a a a
 c r a a PLP. J N r c R 50:190 201.
 P a C, S a C, T r F, K a P, H r a E, P r O, D r a
 A-M, C c a r P. 1996. l a c r a b b r a : c c r c r
 a a r a r a , a a c . M c D 60:13 32.
 P r a NP, R c a r WD. 1993. A a r PDGF a a -
 r c r r a a r a a a r a b a
 a a a a a r c a a . D
 117:525 533.
 P r a NP, Y W-P, G r S, R a H, L a A, P r a AC,
 R c a r WD. 1996. D r a a a a r a c a a :
 c a a r c r c a a b a r a a c a
 S a c . D B 177:30 42.
 Q a X, G r SK, S a Q, S a JH, T S. 1998. l a c
 r a a a a a a a a a r b r a CNS
 r c a r c . D 125:3143 3152.
 R a MS, N b M, M a r -P r c M. 1998. A r a a
 r c r c c r a a a a a a c r . P r c Na
 Aca Sc USA 95:3996 4001.
 R c a r WD, P r a NP, Y W-P, H a AC. 1997. O r a a
 c r a a r c : b r a a a a a a
 r a a a r a . D N r c 19:54 64.
 R a H, A b r r A, H r J, K r V, N r a S, R
 A a b a A, T a a b Y, P a c M, E a T, J TM, D J. 1994.
 F r a a r r a a c c b - l, a r b r a
 r b c c r . C 76:761 775.
 R a BI. 1993. T -Z a c C, P a r a . A N r a Sc 1:187 213.
 S a N, G -Z a c C, P a r a E, O r C, M a r a S,
 I a a A, I a a K, M a c a W, C r r I, Z a c B, T a J-L.
 1998. M r c r c r a a r c . J N r c
 18:8331 8343.
 S a N, O r C, G -Z a c C, M a r a S, T a J-L, Z a c B.
 1999. S a r a a a a : a c r r r .
 G a 29:143 148.
 S a T, H a r AP, R c a r WD, S HK. 1998. P a 6 a a c
 a a a a c c r r a a a r a a a
 b . M C N r c 12:228 239.
 T a a b Y, J TM. 1996. D r a a a a
 a a c r . S c a c 274:1115 1123.