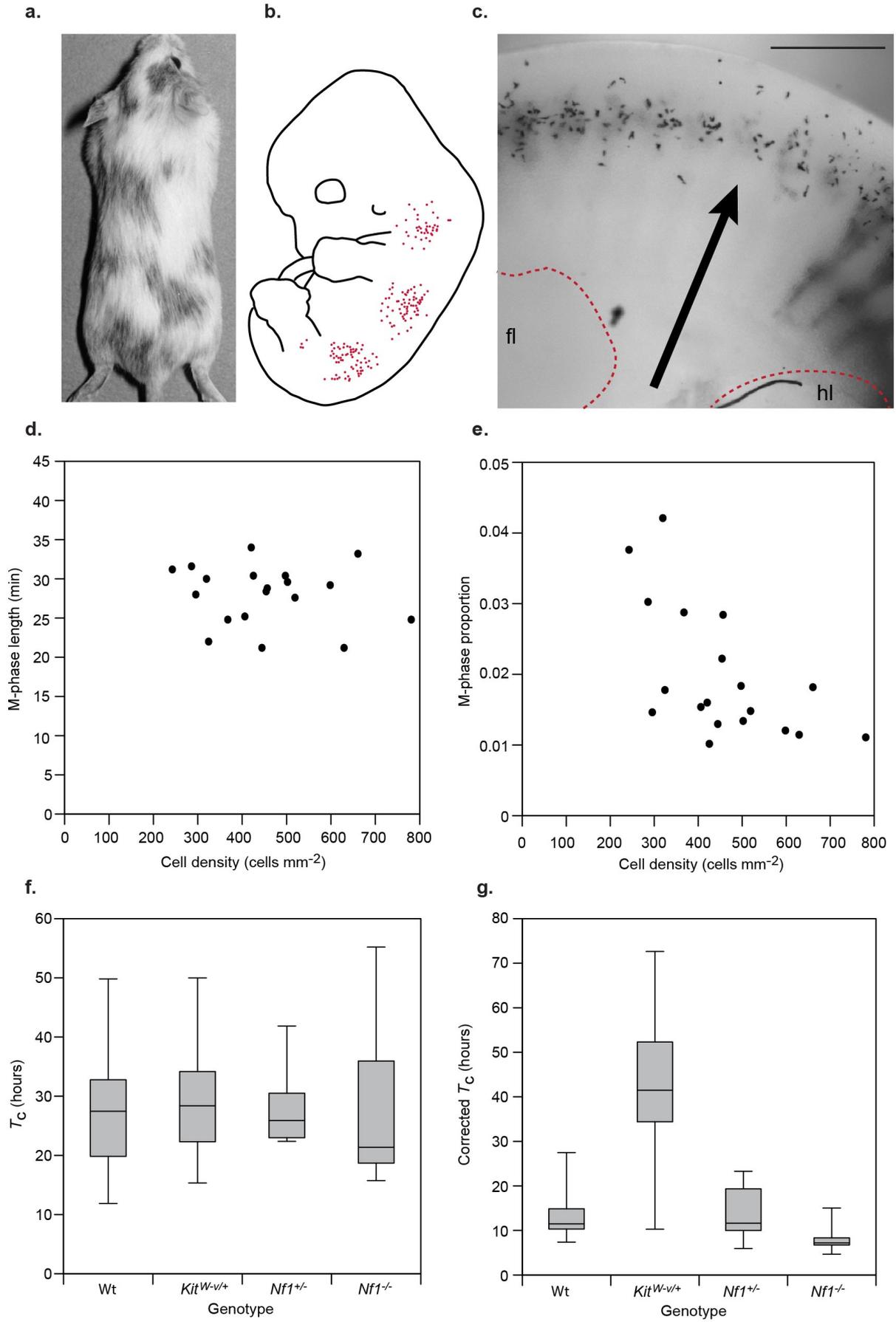
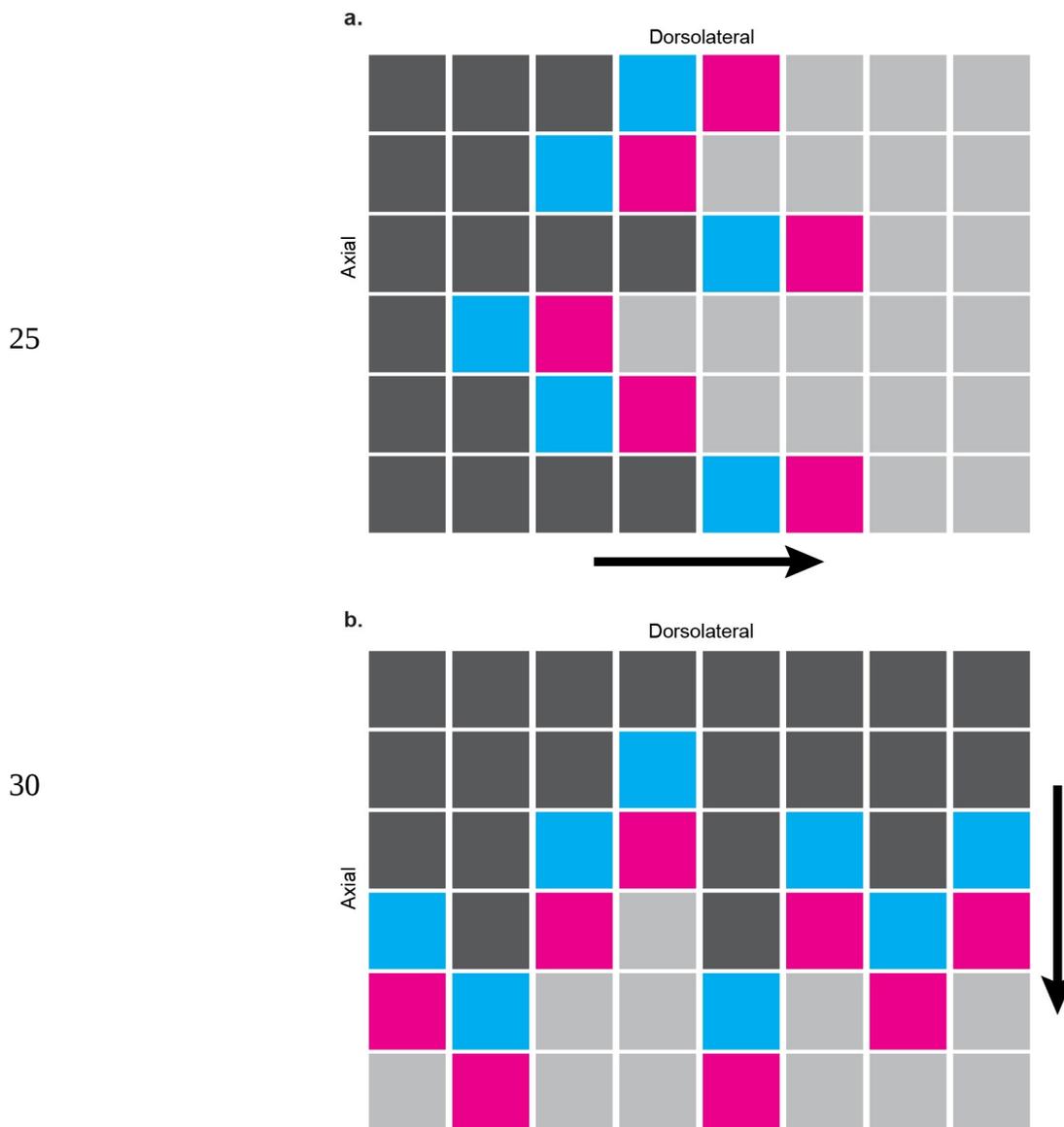


Supplementary Figures

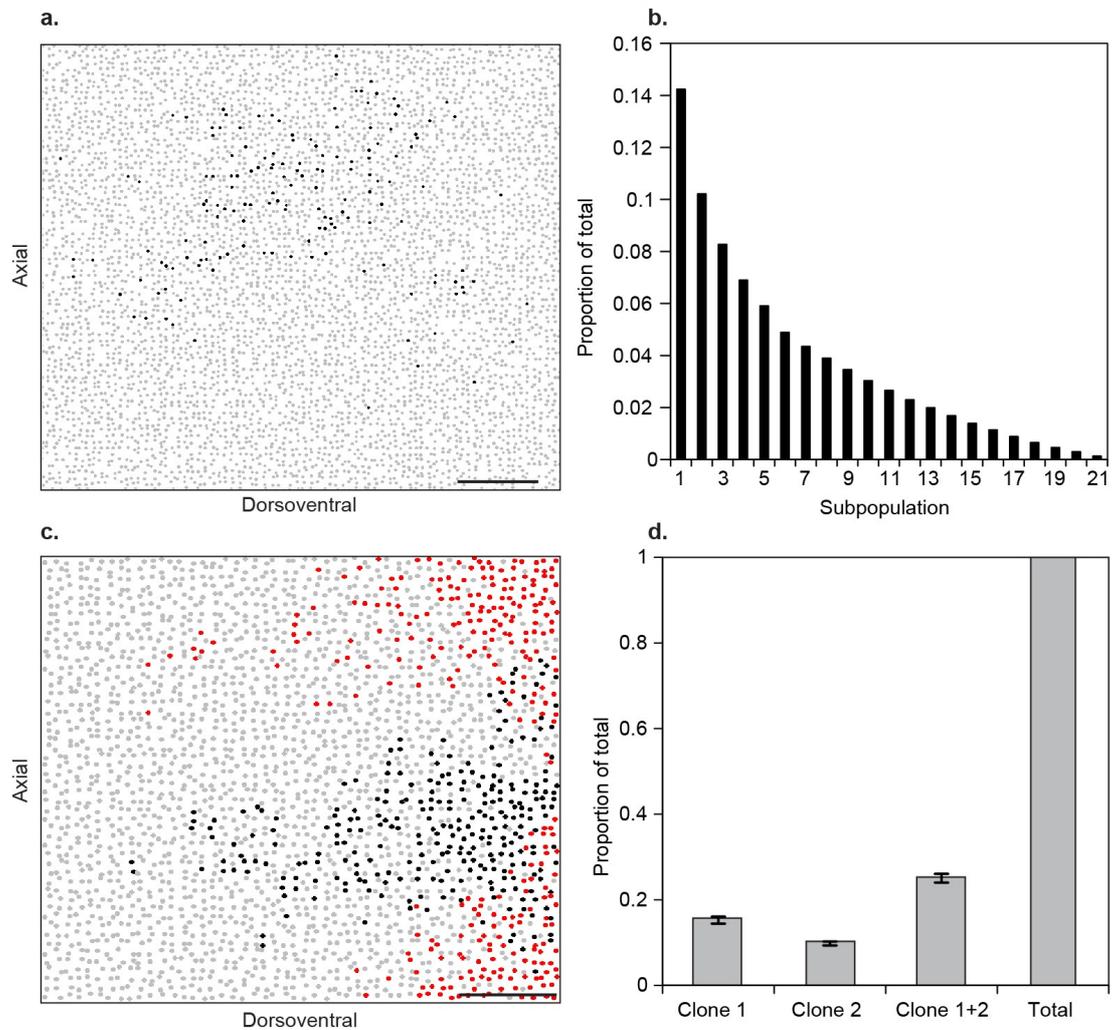


Supplementary Figure 1. Melanoblast patterning and cell cycle. a: A chimaera made by aggregation of cleavage stage embryos of two genotypes: pigment producing and non pigment producing. The patches of pigmentation extend dorsoventrally and generally do not cross the dorsal
5 midline. This can be taken as evidence of the dorsoventral migration in the developing embryo. **b:** Camera lucida generated image of rare (*Dct::lacZ*) melanoblast clones in the developing embryo at E13.5 exhibiting high levels of axial mixing (Redrawn from Wilkie *et al.*¹. **c:** X-Gal stained E11.5 wildtype *Dct::lacZ* embryo showing an under-representation of melanoblasts in the middle of the trunk (black arrow). **d:** Plot of the mean M-phase length (T_m) against density ($n = 19$ wildtype
10 Videos). Pearson product-moment correlation indicates no association between M-phase length and density ($r = -0.15$, d.f = 17, $P = 0.55$). **e:** Analysis of the proportion of cells in M-phase in E14.5 time-lapse sequences ($n = 19$ wildtype samples). Pearson product-moment correlation indicates a significant negative association between mitosis time and density ($r = -0.59$, d.f = 17, $P = 0.007$). **f:** Melanoblast cell cycle times in *Kit*^{+/+}; *Nf1*^{+/+} ($n = 19$), *Kit*^{W-v/+}; *Nf1*^{+/+} ($n = 12$), *Kit*^{+/+}; *Nf1*^{+/-} ($n = 7$)
15 and *Kit*^{+/+}; *Nf1*^{-/-} ($n = 14$) embryonic skin cultures. No group differs significantly from wildtype (*Kit*^{+/+}, *Nf1*^{+/+}) (One-way ANOVA $P = 0.963$). **g:** Melanoblast cell cycle times corrected for cell density in *Kit*^{+/+}; *Nf1*^{+/+} ($n = 20$), *Kit*^{W-v/+}; *Nf1*^{+/+} ($n = 12$), *Kit*^{+/+}; *Nf1*^{+/-} ($n = 7$) and *Kit*^{+/+}; *Nf1*^{-/-} ($n = 14$)
ex vivo E14.5 skin cultures. *Kit*^{W-v/+}, *Nf1*^{+/+} mice have an increased cell cycle time (One-way ANOVA $P < 0.0001$, TukeyHSD $P < 0.0001$). fl = forelimb, hl = hind limb, scale bar in c = 200
20 μm .

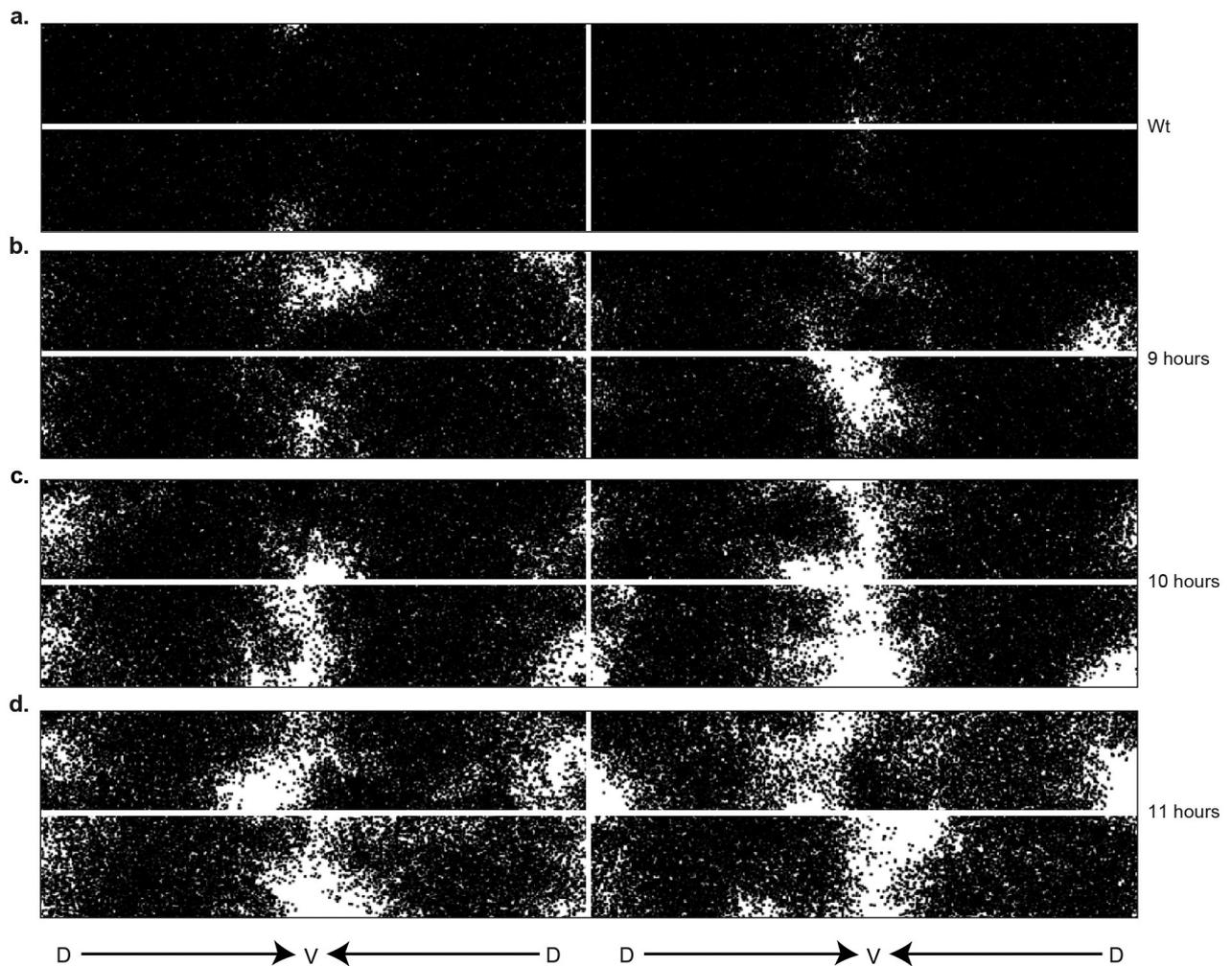


35 **Supplementary Figure 2: Pushing growth mechanism of the discrete model. a:** Growth in the horizontal direction (dorsoventral) for a two-dimensional lattice. The arrow indicates the direction of growth. For each row, a column has been chosen uniformly at random. Together the row and column index specify a site (magenta) that undergoes a growth event: the red site moves one lattice spacing to right, carrying with it any contents, and a new empty site (blue) is inserted in its place.

40 All sites to the right of a red site's initial position move one space to the right to accommodate the new site. **b:** For growth in the axial domain the same mechanism is invoked except that cells move downwards rather than to the right after a growth event.



Supplementary Figure 3: Investigation of dominant sub-clones. **a:** To generate rare clones we chose a single agent from the entire population at a time point chosen uniformly at random and followed its progeny for the rest of the simulation, the example cropped to show the extent of a single rare clone (black cells). The clone was initialised at $t = 8\text{h } 20\text{min}$ and plotted at $t = 120\text{hrs}$ (the end of the simulation). **b:** Plot of the mean agent proportions for all lineages from 100 repeats of the discrete model simulation showing weak selection bias towards a smaller number of dominant lineages. **c:** A dominant lineage pattern from a single simulation of the non-growing discrete model. The two dominant lineages are shown in red and black, all other clones are in grey. **d:** Clone counts from 100 repeats of the discrete simulation without domain growth show that the top 2 dominant clones tend to contribute around 25% of the total number of cells. Error bars in d = s.e.m.



Supplementary Figure 4: Effect of increasing cell cycle time on belly spot formation. **A:** Four
 55 examples of left and right domains showing colonisation using the wildtype parameters (7-hour cell
 cycle time, the ventrum is at the centre of each panel). Some areas of slightly lower density are
 present at the ventrum consistent with experimental data. **b-d:** As the cell cycle time increases
 (from 7-11 hours) melanoblast density at the ventrum and the dorsal region decreases. In the model
 both ends of the domain are sensitive to proliferation the ventrum more so than the dorsum. D =
 60 dorsal, V = ventral.

Supplementary Tables

Supplementary Table 1: Analysis of the distribution of melanoblast angles

Age	Melanoblast marker	Number of Cells	Kolmogorov-Smirnov <i>P</i>
E11.5	<i>Dct::lacZ</i>	325	0.2955
E12.5	<i>Dct::lacZ</i>	232	0.2817
E13.5	<i>Dct::lacZ</i>	331	0.2717
E14.5	<i>Tyr::Cre/R26R-EYFP</i>	414 independent samples	> 0.25 in all cases
E15.5	<i>Dct::lacZ</i>	318	0.2925

Supplementary Table 2: Growth of the mouse trunk between E10.5 and E15.5

Age	N	Mean axial width ($\mu\text{m} \pm 95\% \text{ CI}$)	Mean dorsoventral length ($\mu\text{m} \pm 95\% \text{ CI}$)
E10.5	6	1641.4 \pm 160.3	1496.9 \pm 95.4
E11.5	6	1845.7 \pm 146.1	2461.8 \pm 248.0
E12.5	9	2196.1 \pm 128.1	4684.8 \pm 116.2
E13.5	7	2322.5 \pm 90.2	5409.1 \pm 168.0
E14.5	6	2701.4 \pm 172.7	6590.6 \pm 205.4
E15.5	6	3080.9 \pm 190.2	7901.8 \pm 449.0

65

Supplementary Table 3: Parameters used in the discrete model

Parameter	Value
P_{ga}	0.00526 min^{-1} (one site added to each column per growth event)
P_{gd}	0.0246 min^{-1} (one site added to each row per growth event)
P_m	0.0412 min^{-1}
P_p	0.00165 min^{-1}
Site length, Δ	38 μm
Initial axial length	\approx 1634 μm
Initial dorsoventral length	\approx 1178 μm
Initial agents	21

Agent movement events occur in the model with rate P_m per unit time. Agent proliferation events occur in the model with rate P_p per unit time. The insertion of new lattice sites into the domain occurs with rates P_{ga} and P_{gd} per unit time, for growth in the axial and dorsoventral direction,

70 respectively.

Supplementary References

1. Wilkie, A. L., Jordan, S. A. & Jackson, I. J. Neural crest progenitors of the melanocyte lineage: coat colour patterns revisited. *Development* **129**, 3349–3357 (2002).