Association of Genomic Features with Integration: Unselected vs. Puromycin-Selected HIV

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1 Introduction

In this document, I examine the association of integration siting with various genomic features.

The numbers are shown below:

Origin.of.data.set HIV-Burgess HIVPuro 290 525

The distribution of relative frequency of insertions across the chromosomes is given in this barplot:



Are there chromosomes that are particularly favored for integration by one group over the other? This was tested for statistical significance. The test performed used the likelihood ratio statistic for the logistic regression model (reviewed in [1]) as implemented by the glm function of R using the binomial family. The null hypothesis tested is the ratio of true integration events in the two groups is constant across all chromosomes. This test attains a p-value of 0.35886.

2 Preference for Genes

2.1 Acembly Genes

Here we examine the relative preference that integration events in the two groups have for genes. In the following plot we show the relative frequency of integrations in genes according to the 'Acembly' annotation. The bars grouped over the label "In Gene" give the relative frequency of integration events (compared to control sites) between bases located within Acembly gene annotations, while the label "Not in Gene" give the relative frequency of integration events (compared to control sites) between bases not located within Acembly gene annotations.



Is there is a difference in the tendency for insertions to occur in genes? A formal test of significance yields a p-value of 0.60225. In the following plot we show the relative frequency of insertions in exons according to the 'Acembly' annotation The bars grouped over the label "In Exon" give the relative frequency of integration events (compared to control sites) between bases located in exons according to the Acembly annotation, while the label "Not in Exon" give the relative frequency of integration events (compared to control sites) between bases located in exons according to the Acembly annotation, while the label "Not in Exon" give the relative frequency of integration events (compared to control sites) between bases not located in exons according to the Acembly gene annotation.



Here is the table of coefficients of the log ratio of intensities along with their standard errors, z statistics, and p-values:

	coef	se	Z	р
(Intercept)	0.488	0.215	2.270	0.0231
in.gene	0.179	0.231	0.772	0.4400
in.exon	-0.363	0.209	-1.740	0.0819

The model on which these coefficients are based include terms for whether the site is in a gene or not. Thus, the effect shown as 'in.exon' is net of that due to being in a gene. Note that in the barplot above the 'Not in Exon' bars include the both introns and intergenic regions, so the impression given by the table may differ from that for the barplot.

2.2 refGenes

Here we examine the relative preference that insertions of the two types have for genes. In the following plot we show the relative frequency of insertions in genes according to the 'refGene' annotation.



Is there is a tendency for insertions to occur in genes? A formal test of significance yields a p-value of 0.28323.

In the following plot we show the relative frequency of insertions in exons according to the 'refGene' annotation.



Here is the table of coefficients of the log ratio of intensities for along with their standard errors, z statistics, and p-values:

	coef	se	Z	р
(Intercept)	0.451	0.151	2.99	0.0028
in.gene	0.242	0.174	1.39	0.1640
in.exon	-0.847	0.333	-2.55	0.0109

The model on which these coefficients are based include terms for whether the site is in a gene or not. Thus, the effect shown as 'in.exon' is net of that due to being in a gene. Note that in the barplot above the 'Not in Exon' bars include the both introns and intergenic regions, so the impression given by the table may differ from that for the barplot.

2.3 ensGenes

Here we examine the relative preference that insertions of the two types have for genes. In the following plot we show the relative frequency of insertions in genes according to the 'ensGene' annotation.



Is there is a tendency for insertions to occur in genes? A formal test of significance yields a p-value of 0.22203.

In the following plot we show the relative frequency of insertions in exons according to the 'ensGene' annotation.



Here is the table of coefficients of the log ratio of intensities for along with their standard errors, z statistics, and p-values:

	coef	se	Z	р
(Intercept)	0.411	0.165	2.49	0.01280
in.gene	0.290	0.186	1.56	0.11900
in.exon	-0.883	0.315	-2.81	0.00501

The model on which these coefficients are based include terms for whether the site is in a gene or not. Thus, the effect shown as 'in.exon' is net of that due to being in a gene. Note that in the barplot above the 'Not in Exon' bars include the both introns and intergenic regions, so the impression given by the table may differ from that for the barplot.

2.4 genScan Genes

Here we examine the p erence that insertions have for genes. In the following plot we show the relative frequency of insertions in genes according to the 'genScan' annotation.



Is there is a tendency for insertions to occur in genes? A formal test of significance yields a p-value of 0.94896.

In the following plot we show the relative frequency of insertions in exons according to the 'genScan' annotation.



Here is the table of coefficients of the log ratio of intensities along with their standard errors, z statistics, and p-values:

	coef	se	Z	р
(Intercept)	0.5840	0.171	3.400	0.000664
in.gene	0.0269	0.190	0.142	0.887000
in.exon	-0.4280	0.436	-0.982	0.326000

The model on which these coefficients are based include terms for whether the site is in a gene or not. Thus, the effect shown as 'in.exon' is net of that due to being in a gene. Note that in the barplot above the 'Not in Exon' bars include the both introns and intergenic regions, so the impression given by the table may differ from that for the barplot.

2.5 uniGenes

Here we examine the preference that insertions have for genes. In the following plot we show the relative frequency of insertions in genes according to the 'uniGene' annotation.



Is there is a tendency for insertions to occur in genes? A formal test of significance yields a p-value of 0.33214.

In the following plot we show the relative frequency of insertions in exons according to the 'uniGene' annotation.



Here is the table of coefficients of the log ratio of intensities along with their standard errors, z statistics, and p-values:

	coef	se	Z	р
(Intercept)	0.476	0.141	3.39	0.00071
in.gene	0.226	0.168	1.34	0.18000
in.exon	-0.490	0.249	-1.97	0.04860

The model on which these coefficients are based include terms for whether the site is in a gene or not. Thus, the effect shown as 'in.exon' is net of that due to being in a gene. Note that in the barplot above the 'Not in Exon' bars include the both introns and intergenic regions, so the impression given by the table may differ from that for the barplot.

3 CpG Island Neighborhoods

Here we study the effect of being in the neighborhood of CpG Islands. Following Wu et al [2], who found that the neighborhoods within ± 1 kb of CpG islands are enriched for MLV insertions, we study such neighborhoods.

3.1 1 kilobase neighborhoods

The following plot shows the effect of being in or within ± 1 kb of a CpG island:



A formal test of significance comparing the difference attains a p-value of 0.46878.

3.2 5 kilobase neighborhoods

The following plot shows the effect of being in or within ± 5 kb of a CpG island:



A formal test of significance comparing the difference attains a p-value of 0.21012.

3.3 10 kilobase neighborhoods

The following plot shows the effect of being in or within ± 10 kb of a CpG island:



A formal test of significance comparing the difference attains a p-value of 0.031535.

3.4 25 kilobase neighborhoods

The following plot shows the effect of being in or within ± 25 kb of a CpG island:



A formal test of significance comparing the difference attains a p-value of 0.35651.

3.5 50 kilobase neighborhoods

The following plot shows the effect of being in or within ± 50 kb of a CpG island:



A formal test of significance comparing the difference attains a p-value of 0.29107.

4 Gene Density, Expression 'Density', and CpG Island Density

In this section the association with gene density is examined. The 'genes' that are counted are the genes represented on the microarray. In addition, we the number of such genes expressed at various levels. The levels are

- **low.ex** Count genes whose expression is in the upper half and divide by number of bases
- **med.ex** Count genes whose expression is in the upper $1/8^{th}$ and divide by number of bases
- **high.ex** Count genes whose expression is in the upper $1/16^{th}$ and divide by number of bases

The bolded terms are used as abbreviations in what follows. The abbreviation **dens** is used to indicate gene density as number of genes per base.

4.1 25 kiloBase Window

In the barplot that follows we examine the association of insertion sites with gene density in a 25 kilobase window surrounding each locus. More such plots will follow and the method of their construction is always to try to divide the data according to the deciles of density. However, it often happens that there is a very skewed distribution of density and often even the 90^{th} percentile is zero. In that case, the barplots simply show the sites for which the density is zero and those for which it is non-zero. If there are fewer than ten groups of bars, then the groupings contain ten percent of the sites each except for the leftmost grouping which will contain all of the remaining sites.

Also note that the title of the plot contains clues as to its content; the prefix indicates the type of variable studied while the suffix indicates the window width in the number of bases. The p-value given is the result of fitting a cubic polynomial to the gene density values.

The following expression data and probe set were used for this report:

[1] "HeLa_exp_data-HU133a"

[1] "HG-U133"

Category limits

lower category upper 1 0e+00 group.1 0.00001 2 1e-05 group.2 0.00002 3 2e-05 group.3 0.00004 4 4e-05 group.4 0.00012



dens.25k - p-value = 0.41351

Here are the results for expression density. First, we count just genes that are in the upper half.

	lower	category	upper
1	0.00000e+00	group.1	1.333333e-05
2	1.333333e-05	group.2	3.880000e-05
3	3.880000e-05	group.3	8.00000e-05



low.ex.25k - p-value = 0.3645

Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0e+00	group.1	2.00000e-05
2	2e-05	group.2	6.666667e-05

med.ex.25k - p-value = 0.72577



And here we count genes in the upper $1/16^{th}$:

	lower	category	upper
0%	0.00000e+00	group.1	3.333333e-06
100%	3.333333e-06	group.2	6.00000e-05

high.ex.25k - p-value = 0.64274



Here the effect of density of CpG islands is studied:

Category limits

	lower	category	upper
1	0e+00	group.1	0.00002
2	2e-05	group.2	0.00004
3	4e-05	group.3	0.00024

cpg.dens.25k - p-value = 0.21420



4.2 50 kiloBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 50 kilobase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	0.00000e+00	group.1	4.00000e-06
2	4.00000e-06	group.2	1.000000e-05
3	1.00000e-05	group.3	1.333333e-05
4	1.333333e-05	group.4	2.00000e-05
5	2.00000e-05	group.5	3.00000e-05
6	3.00000e-05	group.6	9.666667e-05



dens.50k - p-value = 0.12079

Here are the results for expression density. First, we count just genes that are in the upper half.

	lower	category	upper
1	0.00000e+00	group.1	3.575758e-06
2	3.575758e-06	group.2	8.00000e-06
3	8.00000e-06	group.3	1.333333e-05
4	1.333333e-05	group.4	2.00000e-05
5	2.00000e-05	group.5	7.00000e-05





Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	6.666667e-06
2	6.666667e-06	group.2	1.666667e-05
3	1.666667e-05	group.3	7.00000e-05

med.ex.50k - p-value = 0.60381



And here we count genes in the upper $1/16^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	6.666667e-06
2	6.666667e-06	group.2	7.00000e-05

high.ex.50k - p-value = 0.62432



Here the effect of density of CpG islands is studied:

	lower	category	upper
1	0e+00	group.1	0.00001
2	1e-05	group.2	0.00002
3	2e-05	group.3	0.00003
4	3e-05	group.4	0.00005
5	5e-05	group.5	0.00026



cpg.dens.50k - p-value = 0.15212

4.3 100 kiloBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 100 kilobase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	0.00000e+00	group.1	4.00000e-06
2	4.00000e-06	group.2	6.666667e-06
3	6.666667e-06	group.3	1.000000e-05
4	1.00000e-05	group.4	1.100000e-05
5	1.100000e-05	group.5	1.666667e-05
6	1.666667e-05	group.6	2.500000e-05
7	2.50000e-05	group.7	8.833333e-05



dens.100k - p-value = 0.10897

Here are the results for expression density. First, we count just genes that are in the upper half.

	lower	category	upper
1	0.00000e+00	group.1	2.678571e-06
2	2.678571e-06	group.2	5.00000e-06
3	5.00000e-06	group.3	7.850000e-06
4	7.850000e-06	group.4	1.000000e-05
5	1.00000e-05	group.5	1.666667e-05
6	1.666667e-05	group.6	6.00000e-05



low.ex.100k - p-value = 0.83069

Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	3.333333e-06
2	3.333333e-06	group.2	6.666667e-06
3	6.666667e-06	group.3	1.000000e-05
4	1.00000e-05	group.4	5.00000e-05



med.ex.100k - p-value = 0.82232

And here we count genes in the upper $1/16^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	3.333333e-07
2	3.333333e-07	group.2	5.00000e-06
3	5.00000e-06	group.3	3.50000e-05

high.ex.100k - p-value = 0.43985



Here the effect of density of CpG islands is studied:

lower	category	upper
0.00e+00	group.1	5.00e-06
5.00e-06	group.2	1.00e-05
1.00e-05	group.3	1.50e-05
1.50e-05	group.4	2.05e-05
2.05e-05	group.5	3.20e-05
3.20e-05	group.6	5.50e-05
5.50e-05	group.7	1.95e-04
	lower 0.00e+00 5.00e-06 1.00e-05 1.50e-05 2.05e-05 3.20e-05 5.50e-05	lower category 0.00e+00 group.1 5.00e-06 group.2 1.00e-05 group.3 1.50e-05 group.4 2.05e-05 group.5 3.20e-05 group.6 5.50e-05 group.7



cpg.dens.100k - p-value = 0.516

4.4 250 kiloBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 250 kilobase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

upper	category	lower	
2.00000e-06	group.1	0.00000e+00	1
4.00000e-06	group.2	2.00000e-06	2
5.333333e-06	group.3	4.00000e-06	3
6.954545e-06	group.4	5.333333e-06	4
9.00000e-06	group.5	6.954545e-06	5
1.200000e-05	group.6	9.00000e-06	6
1.60000e-05	group.7	1.200000e-05	7
2.40000e-05	group.8	1.600000e-05	8
7.20000e-05	group.9	2.40000e-05	9



dens.250k - p-value = 0.89626
	lower	category	upper
1	0.00000e+00	group.1	1.333333e-06
2	1.333333e-06	group.2	2.135238e-06
3	2.135238e-06	group.3	4.00000e-06
4	4.00000e-06	group.4	4.647619e-06
5	4.647619e-06	group.5	6.671429e-06
6	6.671429e-06	group.6	9.066667e-06
7	9.066667e-06	group.7	1.400000e-05
8	1.40000e-05	group.8	4.280000e-05





Now we count genes in the upper $1/8^{th}$:

	lower	category	unner
	TOMOT	cancent	upper
1	0.000000e+00	group.1	1.333333e-06
2	1.333333e-06	group.2	2.106667e-06
3	2.106667e-06	group.3	4.00000e-06
4	4.00000e-06	group.4	5.333333e-06
5	5.333333e-06	group.5	8.00000e-06
6	8.00000e-06	group.6	2.666667e-05

med.ex.250k - p-value = 0.30575



And here we count genes in the upper $1/16^{th}\colon$

	lower	category	upper
1	0.00e+00	group.1	1.36e-06
2	1.36e-06	group.2	3.20e-06
3	3.20e-06	group.3	4.00e-06
4	4.00e-06	group.4	1.80e-05



high.ex.250k - p-value = 0.27463

Here the effect of density of CpG islands is studied:

	lower	category	upper
1	0.00e+00	group.1	2.00e-06
2	2.00e-06	group.2	4.00e-06
3	4.00e-06	group.3	8.00e-06
4	8.00e-06	group.4	1.00e-05
5	1.00e-05	group.5	1.20e-05
6	1.20e-05	group.6	1.80e-05
7	1.80e-05	group.7	2.40e-05
8	2.40e-05	group.8	3.40e-05
9	3.40e-05	group.9	5.94e-05
10	5.94e-05	group.10	1.98e-04

cpg.dens.250k - p-value = 0.8295



4.5 500 kiloBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 500 kilobase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	0.00000e+00	group.1	1.333333e-06
2	1.333333e-06	group.2	2.333333e-06
3	2.333333e-06	group.3	3.833333e-06
4	3.833333e-06	group.4	5.147619e-06
5	5.147619e-06	group.5	6.878571e-06
6	6.878571e-06	group.6	8.666667e-06
7	8.666667e-06	group.7	1.167000e-05
8	1.167000e-05	group.8	1.580000e-05
9	1.580000e-05	group.9	2.433333e-05
10	2.433333e-05	group.10	7.433333e-05



dens.500k - p-value = 0.74484

	lower	category	upper
1	0.00000e+00	group.1	1.000000e-06
2	1.000000e-06	group.2	2.00000e-06
3	2.00000e-06	group.3	2.50000e-06
4	2.50000e-06	group.4	3.50000e-06
5	3.50000e-06	group.5	4.666667e-06
6	4.666667e-06	group.6	6.273333e-06
7	6.273333e-06	group.7	8.802222e-06
8	8.802222e-06	group.8	1.40000e-05
9	1.40000e-05	group.9	3.633333e-05





Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	4.00000e-07
2	4.00000e-07	group.2	1.00000e-06
3	1.000000e-06	group.3	2.00000e-06
4	2.00000e-06	group.4	2.333333e-06
5	2.333333e-06	group.5	3.333333e-06
6	3.333333e-06	group.6	4.666667e-06
7	4.666667e-06	group.7	7.950000e-06
8	7.950000e-06	group.8	2.00000e-05





And here we count genes in the upper $1/16^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	5.00000e-07
2	5.00000e-07	group.2	1.000000e-06
3	1.000000e-06	group.3	2.00000e-06
4	2.00000e-06	group.4	2.333333e-06
5	2.333333e-06	group.5	4.00000e-06
6	4.00000e-06	group.6	1.40000e-05

high.ex.500k - p-value = 0.4721



Here the effect of density of CpG islands is studied:

	lower	category	upper
1	0.00e+00	group.1	2.00e-06
2	2.00e-06	group.2	4.00e-06
3	4.00e-06	group.3	7.00e-06
4	7.00e-06	group.4	9.00e-06
5	9.00e-06	group.5	1.20e-05
6	1.20e-05	group.6	1.70e-05
7	1.70e-05	group.7	2.30e-05
8	2.30e-05	group.8	3.40e-05
9	3.40e-05	group.9	5.57e-05
10	5.57e-05	group.10	1.83e-04

cpg.dens.500k - p-value = 0.91448



4.6 1 megaBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 1 megabase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	0.000000e+00	group.1	1.345833e-06
2	1.345833e-06	group.2	2.50000e-06
3	2.50000e-06	group.3	3.795000e-06
4	3.795000e-06	group.4	5.083333e-06
5	5.083333e-06	group.5	6.50000e-06
6	6.50000e-06	group.6	8.093333e-06
7	8.093333e-06	group.7	1.133385e-05
8	1.133385e-05	group.8	1.565429e-05
9	1.565429e-05	group.9	2.133333e-05
10	2.133333e-05	group.10	5.516667e-05



dens.1M - p-value = 0.3788

	lower	category	upper
1	0.000000e+00	group.1	3.333333e-07
2	3.333333e-07	group.2	1.000000e-06
3	1.000000e-06	group.3	1.730000e-06
4	1.730000e-06	group.4	2.500000e-06
5	2.500000e-06	group.5	3.308333e-06
6	3.308333e-06	group.6	4.333333e-06
7	4.333333e-06	group.7	5.750000e-06
8	5.750000e-06	group.8	8.00000e-06
9	8.00000e-06	group.9	1.248500e-05
10	1.248500e-05	group.10	2.691667e-05

low.ex.1M - p-value = 0.20182



Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	2.50000e-07
2	2.50000e-07	group.2	6.666667e-07
3	6.666667e-07	group.3	1.083333e-06
4	1.083333e-06	group.4	1.666667e-06
5	1.666667e-06	group.5	2.166072e-06
6	2.166072e-06	group.6	3.00000e-06
7	3.00000e-06	group.7	4.00000e-06
8	4.00000e-06	group.8	6.803333e-06
9	6.803333e-06	group.9	1.400000e-05





And here we count genes in the upper $1/16^{th}\colon$

	lower	category	upper
1	0.00000e+00	group.1	4.00000e-07
2	4.00000e-07	group.2	7.500000e-07
3	7.50000e-07	group.3	1.000000e-06
4	1.000000e-06	group.4	1.500000e-06
5	1.500000e-06	group.5	2.166667e-06
6	2.166667e-06	group.6	3.500000e-06
7	3.500000e-06	group.7	8.50000e-06



high.ex.1M - p-value = 0.82602

Here the effect of density of CpG islands is studied:

	lower	category	upper
1	0.000e+00	group.1	3.000e-06
2	3.000e-06	group.2	4.300e-06
3	4.300e-06	group.3	6.500e-06
4	6.500e-06	group.4	9.000e-06
5	9.000e-06	group.5	1.250e-05
6	1.250e-05	group.6	1.800e-05
7	1.800e-05	group.7	2.250e-05
8	2.250e-05	group.8	3.170e-05
9	3.170e-05	group.9	5.235e-05
10	5.235e-05	group.10	1.685e-04

cpg.dens.1M - p-value = 0.99526



4.7 2 megaBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 2 megabase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	0.000000e+00	group.1	1.566667e-06
2	1.566667e-06	group.2	2.50000e-06
3	2.50000e-06	group.3	3.541667e-06
4	3.541667e-06	group.4	4.626667e-06
5	4.626667e-06	group.5	6.021429e-06
6	6.021429e-06	group.6	8.185000e-06
7	8.185000e-06	group.7	1.120083e-05
8	1.120083e-05	group.8	1.518833e-05
9	1.518833e-05	group.9	1.941667e-05
10	1.941667e-05	group.10	3.912500e-05



dens.2M - p-value = 0.023753

	lower	category	upper
1	0.000000e+00	group.1	5.00000e-07
2	5.00000e-07	group.2	1.082857e-06
3	1.082857e-06	group.3	1.591905e-06
4	1.591905e-06	group.4	2.102619e-06
5	2.102619e-06	group.5	2.766667e-06
6	2.766667e-06	group.6	3.915000e-06
7	3.915000e-06	group.7	5.941667e-06
8	5.941667e-06	group.8	7.567857e-06
9	7.567857e-06	group.9	9.969167e-06
10	9.969167e-06	group.10	2.093333e-05

low.ex.2M - p-value = 0.031691



Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	1.461538e-07
2	1.461538e-07	group.2	4.707143e-07
3	4.707143e-07	group.3	7.50000e-07
4	7.500000e-07	group.4	1.041667e-06
5	1.041667e-06	group.5	1.416667e-06
6	1.416667e-06	group.6	1.958333e-06
7	1.958333e-06	group.7	2.810811e-06
8	2.810811e-06	group.8	3.995000e-06
9	3.995000e-06	group.9	5.570833e-06
10	5.570833e-06	group.10	1.126667e-05

med.ex.2M - p-value = 0.070378



And here we count genes in the upper $1/16^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	1.00000e-07
2	1.00000e-07	group.2	2.50000e-07
3	2.50000e-07	group.3	5.00000e-07
4	5.00000e-07	group.4	6.898810e-07
5	6.898810e-07	group.5	1.000000e-06
6	1.000000e-06	group.6	1.375000e-06
7	1.375000e-06	group.7	1.923333e-06
8	1.923333e-06	group.8	2.737500e-06
9	2.737500e-06	group.9	7.433333e-06





Here the effect of density of CpG islands is studied:

	lower	category	upper
1	0.00000e+00	group.1	2.825000e-06
2	2.825000e-06	group.2	4.50000e-06
3	4.50000e-06	group.3	5.750000e-06
4	5.750000e-06	group.4	8.253564e-06
5	8.253564e-06	group.5	1.225000e-05
6	1.225000e-05	group.6	1.770000e-05
7	1.770000e-05	group.7	2.218569e-05
8	2.218569e-05	group.8	2.835000e-05
9	2.835000e-05	group.9	4.785000e-05
10	4.785000e-05	group.10	1.476696e-04

cpg.dens.2M - p-value = 0.99862



4.8 4 megaBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 4 megabase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	3.571429e-08	group.1	1.822798e-06
2	1.822798e-06	group.2	2.557500e-06
3	2.557500e-06	group.3	3.279167e-06
4	3.279167e-06	group.4	4.209524e-06
5	4.209524e-06	group.5	5.728125e-06
6	5.728125e-06	group.6	7.947841e-06
7	7.947841e-06	group.7	1.042583e-05
8	1.042583e-05	group.8	1.331500e-05
9	1.331500e-05	group.9	1.725521e-05
10	1.725521e-05	group.10	3.462917e-05

dens.4M - p-value = 0.0084586



	lower	category	upper
1	0.000000e+00	group.1	7.720833e-07
2	7.720833e-07	group.2	1.097262e-06
3	1.097262e-06	group.3	1.474226e-06
4	1.474226e-06	group.4	1.910256e-06
5	1.910256e-06	group.5	2.598810e-06
6	2.598810e-06	group.6	3.825833e-06
7	3.825833e-06	group.7	5.262500e-06
8	5.262500e-06	group.8	6.631667e-06
9	6.631667e-06	group.9	8.235681e-06
10	8.235681e-06	group.10	1.792917e-05

low.ex.4M - p-value = 0.0065512



Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	2.562500e-07
2	2.562500e-07	group.2	5.416667e-07
3	5.416667e-07	group.3	7.291667e-07
4	7.291667e-07	group.4	9.782143e-07
5	9.782143e-07	group.5	1.330684e-06
6	1.330684e-06	group.6	1.851667e-06
7	1.851667e-06	group.7	2.725417e-06
8	2.725417e-06	group.8	3.380682e-06
9	3.380682e-06	group.9	4.661907e-06
10	4.661907e-06	group.10	9.337500e-06

med.ex.4M - p-value = 0.004282



And here we count genes in the upper $1/16^{th}\colon$

	lower	category	upper
1	0.00000e+00	group.1	8.333333e-08
2	8.333333e-08	group.2	2.50000e-07
3	2.50000e-07	group.3	3.333333e-07
4	3.333333e-07	group.4	5.208333e-07
5	5.208333e-07	group.5	6.666667e-07
6	6.666667e-07	group.6	9.350000e-07
7	9.350000e-07	group.7	1.209722e-06
8	1.209722e-06	group.8	1.666667e-06
9	1.666667e-06	group.9	2.276667e-06
10	2.276667e-06	group.10	5.766667e-06

high.ex.4M - p-value = 0.037411



Here the effect of density of CpG islands is studied:

	lower	category	upper
1	5.00000e-07	group.1	3.250000e-06
2	3.250000e-06	group.2	4.250000e-06
3	4.250000e-06	group.3	5.862500e-06
4	5.862500e-06	group.4	7.650000e-06
5	7.650000e-06	group.5	1.156250e-05
6	1.156250e-05	group.6	1.600000e-05
7	1.600000e-05	group.7	2.100611e-05
8	2.100611e-05	group.8	2.517500e-05
9	2.517500e-05	group.9	4.175000e-05
10	4.175000e-05	group.10	1.235785e-04

cpg.dens.4M - p-value = 0.92457



4.9 8 megaBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 8 megabase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	2.083333e-07	group.1	1.906598e-06
2	1.906598e-06	group.2	2.470119e-06
3	2.470119e-06	group.3	3.193542e-06
4	3.193542e-06	group.4	4.052008e-06
5	4.052008e-06	group.5	5.521181e-06
6	5.521181e-06	group.6	7.063116e-06
7	7.063116e-06	group.7	8.620625e-06
8	8.620625e-06	group.8	1.139873e-05
9	1.139873e-05	group.9	1.546811e-05
10	1.546811e-05	group.10	2.956875e-05

dens.8M - p-value = 0.012632



	lower	category	upper
1	0.000000e+00	group.1	7.374420e-07
2	7.374420e-07	group.2	1.060417e-06
3	1.060417e-06	group.3	1.385578e-06
4	1.385578e-06	group.4	1.766667e-06
5	1.766667e-06	group.5	2.516667e-06
6	2.516667e-06	group.6	3.314523e-06
7	3.314523e-06	group.7	4.458681e-06
8	4.458681e-06	group.8	5.663284e-06
9	5.663284e-06	group.9	7.561637e-06
10	7.561637e-06	group.10	1.221042e-05

low.ex.8M - p-value = 0.024777



Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	0.00000e+00	group.1	3.378125e-07
2	3.378125e-07	group.2	5.256250e-07
3	5.256250e-07	group.3	6.916667e-07
4	6.916667e-07	group.4	8.945734e-07
5	8.945734e-07	group.5	1.267708e-06
6	1.267708e-06	group.6	1.709524e-06
7	1.709524e-06	group.7	2.210625e-06
8	2.210625e-06	group.8	3.099444e-06
9	3.099444e-06	group.9	3.880208e-06
10	3.880208e-06	group.10	6.927083e-06

med.ex.8M - p-value = 0.010577



And here we count genes in the upper $1/16^{th}\colon$

	lower	category	upper
1	0.00000e+00	group.1	1.562500e-07
2	1.562500e-07	group.2	2.50000e-07
3	2.50000e-07	group.3	3.330476e-07
4	3.330476e-07	group.4	4.583333e-07
5	4.583333e-07	group.5	6.568452e-07
6	6.568452e-07	group.6	8.729167e-07
7	8.729167e-07	group.7	1.062500e-06
8	1.062500e-06	group.8	1.428467e-06
9	1.428467e-06	group.9	1.935417e-06
10	1.935417e-06	group.10	4.458333e-06

high.ex.8M - p-value = 0.035593



Here the effect of density of CpG islands is studied:

	lower	category	upper
1	7.500000e-07	group.1	3.562500e-06
2	3.562500e-06	group.2	4.487279e-06
3	4.487279e-06	group.3	5.750000e-06
4	5.750000e-06	group.4	7.839015e-06
5	7.839015e-06	group.5	1.037500e-05
6	1.037500e-05	group.6	1.344793e-05
7	1.344793e-05	group.7	1.787500e-05
8	1.787500e-05	group.8	2.311342e-05
9	2.311342e-05	group.9	3.458812e-05
10	3.458812e-05	group.10	9.242333e-05

cpg.dens.8M - p-value = 0.79856



4.10 16 megaBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 16 megabase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	6.166667e-07	group.1	1.858780e-06
2	1.858780e-06	group.2	2.474033e-06
3	2.474033e-06	group.3	3.130491e-06
4	3.130491e-06	group.4	3.900208e-06
5	3.900208e-06	group.5	5.193123e-06
6	5.193123e-06	group.6	6.079664e-06
7	6.079664e-06	group.7	7.680484e-06
8	7.680484e-06	group.8	9.670616e-06
9	9.670616e-06	group.9	1.312768e-05
10	1.312768e-05	group.10	1.928512e-05



dens.16M - p-value = 0.0020961

	lower	category	upper
1	1.919643e-07	group.1	7.821875e-07
2	7.821875e-07	group.2	1.094980e-06
3	1.094980e-06	group.3	1.303125e-06
4	1.303125e-06	group.4	1.748333e-06
5	1.748333e-06	group.5	2.397900e-06
6	2.397900e-06	group.6	2.838750e-06
7	2.838750e-06	group.7	3.606250e-06
8	3.606250e-06	group.8	5.100166e-06
9	5.100166e-06	group.9	6.354021e-06
10	6.354021e-06	group.10	9.098263e-06

low.ex.16M - p-value = 0.00523



Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	6.250000e-08	group.1	3.794792e-07
2	3.794792e-07	group.2	5.270833e-07
3	5.270833e-07	group.3	6.779167e-07
4	6.779167e-07	group.4	9.320536e-07
5	9.320536e-07	group.5	1.184896e-06
6	1.184896e-06	group.6	1.496875e-06
7	1.496875e-06	group.7	1.897917e-06
8	1.897917e-06	group.8	2.681056e-06
9	2.681056e-06	group.9	3.113679e-06
10	3.113679e-06	group.10	4.852258e-06

med.ex.16M - p-value = 0.0013861



And here we count genes in the upper $1/16^{th}\colon$

	lower	category	upper
1	0.00000e+00	group.1	1.875000e-07
2	1.875000e-07	group.2	2.604167e-07
3	2.604167e-07	group.3	3.505878e-07
4	3.505878e-07	group.4	4.843750e-07
5	4.843750e-07	group.5	6.093750e-07
6	6.093750e-07	group.6	7.706250e-07
7	7.706250e-07	group.7	9.528362e-07
8	9.528362e-07	group.8	1.108333e-06
9	1.108333e-06	group.9	1.630218e-06
10	1.630218e-06	group.10	2.718750e-06

high.ex.16M - p-value = 0.0014961



Here the effect of density of CpG islands is studied:

	lower	category	upper
1	1.343750e-06	group.1	3.884375e-06
2	3.884375e-06	group.2	4.656250e-06
3	4.656250e-06	group.3	6.031250e-06
4	6.031250e-06	group.4	7.805925e-06
5	7.805925e-06	group.5	9.625000e-06
6	9.625000e-06	group.6	1.121250e-05
7	1.121250e-05	group.7	1.606562e-05
8	1.606562e-05	group.8	1.916041e-05
9	1.916041e-05	group.9	2.473723e-05
10	2.473723e-05	group.10	6.694779e-05

cpg.dens.16M - p-value = 0.7996



4.11 32 megaBase Window

In the barplot that follows we examine the association of insertion sites with expression density in a 32 megabase window surrounding each locus. First, we count just the number of genes on the represented on the chip.

	lower	category	upper
1	5.753393e-07	group.1	2.036203e-06
2	2.036203e-06	group.2	2.534345e-06
3	2.534345e-06	group.3	3.160329e-06
4	3.160329e-06	group.4	3.977480e-06
5	3.977480e-06	group.5	4.703423e-06
6	4.703423e-06	group.6	5.361487e-06
7	5.361487e-06	group.7	6.739752e-06
8	6.739752e-06	group.8	8.247105e-06
9	8.247105e-06	group.9	9.252270e-06
10	9.252270e-06	group.10	1.839120e-05

dens.32M - p-value = 0.008804



	lower	category	upper
1	2.840978e-07	group.1	8.776843e-07
2	8.776843e-07	group.2	1.091334e-06
3	1.091334e-06	group.3	1.396384e-06
4	1.396384e-06	group.4	1.752302e-06
5	1.752302e-06	group.5	2.099380e-06
6	2.099380e-06	group.6	2.551565e-06
7	2.551565e-06	group.7	3.279120e-06
8	3.279120e-06	group.8	4.097512e-06
9	4.097512e-06	group.9	4.536546e-06
10	4.536546e-06	group.10	8.754914e-06

low.ex.32M - p-value = 0.0055423


Now we count genes in the upper $1/8^{th}$:

	lower	category	upper
1	9.050701e-08	group.1	4.401184e-07
2	4.401184e-07	group.2	5.687500e-07
3	5.687500e-07	group.3	7.312846e-07
4	7.312846e-07	group.4	9.068750e-07
5	9.068750e-07	group.5	1.088542e-06
6	1.088542e-06	group.6	1.276027e-06
7	1.276027e-06	group.7	1.764750e-06
8	1.764750e-06	group.8	2.014504e-06
9	2.014504e-06	group.9	2.438892e-06
10	2.438892e-06	group.10	4.505173e-06

med.ex.32M - p-value = 0.00047359



And here we count genes in the upper $1/16^{th}\colon$

	lower	category	upper
1	3.016900e-08	group.1	2.124740e-07
2	2.124740e-07	group.2	2.992708e-07
3	2.992708e-07	group.3	3.632552e-07
4	3.632552e-07	group.4	4.630208e-07
5	4.630208e-07	group.5	5.781250e-07
6	5.781250e-07	group.6	7.018929e-07
7	7.018929e-07	group.7	8.080357e-07
8	8.080357e-07	group.8	9.372024e-07
9	9.372024e-07	group.9	1.207031e-06
10	1.207031e-06	group.10	2.171921e-06

high.ex.32M - p-value = 0.00060393



Here the effect of density of CpG islands is studied:

	lower	category	upper
1	2.734375e-06	group.1	4.433832e-06
2	4.433832e-06	group.2	5.643750e-06
3	5.643750e-06	group.3	6.611517e-06
4	6.611517e-06	group.4	7.296875e-06
5	7.296875e-06	group.5	8.469715e-06
6	8.469715e-06	group.6	1.043897e-05
7	1.043897e-05	group.7	1.246406e-05
8	1.246406e-05	group.8	1.665107e-05
9	1.665107e-05	group.9	2.010270e-05
10	2.010270e-05	group.10	4.036208e-05

cpg.dens.32M - p-value = 0.23685



5 Juxtaposition with Gene Start and End Positions

5.1 Acembly Annotations

In this section we study the effect of juxtaposition in terms of gene start and end positions. The first barplot shows the effect of gene width for those insertions that are located within an Acembly gene.

Category	limits
oucepor,	TTmT0D

	lower	category	upper
1	319.0	group.1	14085.2
2	14085.2	group.2	23312.6
3	23312.6	group.3	34922.3
4	34922.3	group.4	46396.8
5	46396.8	group.5	61817.0
6	61817.0	group.6	77882.0
7	77882.0	group.7	105342.3
8	105342.3	group.8	142274.6
9	142274.6	group.9	204679.4
10	204679.4	group.10	777203.0

acembly gene.width - p-value = 0.27387



The next plot uses the width of a non-gene region for insertions that fall into such regions.

	lower	category	upper
1	1831.0	group.1	10067.8
2	10067.8	group.2	18439.8
3	18439.8	group.3	28426.9
4	28426.9	group.4	55108.4
5	55108.4	group.5	69209.5
6	69209.5	group.6	100116.4
7	100116.4	group.7	161656.4
8	161656.4	group.8	204695.0
9	204695.0	group.9	304640.0
10	304640.0	group.10	765281.0



acembly non-gene width - p-value = 0.11199

The next plot studies the distance to the nearest boundary between a gene and a non-gene region. The distance is expressed as a fraction of the length of the region. Thus, '0.25' refers to one quarter of the distance from the site to nearest boundary divided by the total width of the region.

	lower	category	upper
1	0.0001694987	group.1	0.05353284
2	0.0535328379	group.2	0.10995390
3	0.1099538989	group.3	0.15953682
4	0.1595368165	group.4	0.21857310
5	0.2185731007	group.5	0.25746174
6	0.2574617365	group.6	0.31155298
7	0.3115529771	group.7	0.35748191
8	0.3574819092	group.8	0.40318291
9	0.4031829149	group.9	0.44503803
10	0.4450380305	group.10	0.49994218



acembly boundary.dist - p-value = 0.37758

This plot studies the effect of nearness to the beginning of a transcript. For sites in genes, it is the distance to the start of the gene divided by the width of the gene. For other sites it is the distance from the site to the nearer gene if that gene boundary is also a transcription starting point. Locations near '0' are relatively near the beginning of transcription, while those near '1' are near the termination of the transcript.

	lower	category	upper
1	0.0009294105	group.1	0.09256817
2	0.0925681712	group.2	0.18756371
3	0.1875637105	group.3	0.26255787
4	0.2625578735	group.4	0.34918394
5	0.3491839386	group.5	0.43963176
6	0.4396317624	group.6	0.54339454
7	0.5433945379	group.7	0.63546206
8	0.6354620571	group.8	0.74275959
9	0.7427595947	group.9	0.86813987
10	0.8681398661	group.10	0.99983050



acembly start.dist - p-value = 0.45331

5.2 RefSeq Annotations

Category limits

	lower	category	upper
1	4339.0	group.1	23732.9
2	23732.9	group.2	35378.4
3	35378.4	group.3	46472.1
4	46472.1	group.4	63481.8
5	63481.8	group.5	80910.0
6	80910.0	group.6	102225.0
7	102225.0	group.7	131988.4
8	131988.4	group.8	166140.0
9	166140.0	group.9	247961.7
10	247961.7	group.10	992672.0

refSeq gene.width - p-value = 0.73186



	lower	category	upper
1	3478.0	group.1	24812.3
2	24812.3	group.2	48950.8
3	48950.8	group.3	76135.6
4	76135.6	group.4	119032.0

5	119032.0	group.5	185385.5
6	185385.5	group.6	337935.4
7	337935.4	group.7	540105.2
8	540105.2	group.8	923251.2
9	923251.2	group.9	1682141.6
10	1682141.6	group.10	21293005.0

0.14 0.12 0.10 0.00 0.08 0.00 0.04 0.02 0.00 group.1 group.3 group.5 group.7 group.9

refSeq non-gene width - p-value = 0.43048

	lower	category	upper
1	0.001127961	group.1	0.06017484
2	0.060174840	group.2	0.11592495
3	0.115924954	group.3	0.16239860
4	0.162398602	group.4	0.21729528
5	0.217295283	group.5	0.26540012
6	0.265400123	group.6	0.31453438
7	0.314534378	group.7	0.35773016
8	0.357730160	group.8	0.40532204
9	0.405322039	group.9	0.45348723
10	0.453487226	group.10	0.49990109



refSeq boundary.dist - p-value = 0.015346

	lower	category	upper
1	0.002402913	group.1	0.09925882
2	0.099258817	group.2	0.18405402
3	0.184054021	group.3	0.27356617
4	0.273566167	group.4	0.35470220
5	0.354702204	group.5	0.45746632
6	0.457466318	group.6	0.54922751
7	0.549227514	group.7	0.64173437
8	0.641734368	group.8	0.73845920
9	0.738459204	group.9	0.86435298
10	0.864352976	group.10	0.99887204



refSeq start.dist - p-value = 0.080218

5.3 genScan Annotations

Category limits

	lower	category	upper
1	4684.0	group.1	24682.5
2	24682.5	group.2	39421.0
3	39421.0	group.3	53523.5
4	53523.5	group.4	72479.0
5	72479.0	group.5	88359.0
6	88359.0	group.6	113194.0
7	113194.0	group.7	139572.5
8	139572.5	group.8	184425.0
9	184425.0	group.9	230325.5
10	230325.5	group.10	644368.0

genScan gene.width - p-value = 0.75324



	lower	category	upper
1	690.0	group.1	7004.7
2	7004.7	group.2	10520.4
3	10520.4	group.3	17265.9
4	17265.9	group.4	22417.6

5	22417.6	group.5	29233.0
6	29233.0	group.6	34344.4
7	34344.4	group.7	43923.5
8	43923.5	group.8	56102.0
9	56102.0	group.9	79819.1
10	79819.1	group.10	20489465.0

genScan non-gene width - p-value = 0.27779



	lower	category	upper
1	0.0005360666	group.1	0.0739102
2	0.0739101980	group.2	0.1333521
3	0.1333520781	group.3	0.1750946
4	0.1750945709	group.4	0.2191114
5	0.2191114320	group.5	0.2702058
6	0.2702058292	group.6	0.3190237
7	0.3190236584	group.7	0.3703712
8	0.3703711563	group.8	0.4080653
9	0.4080653451	group.9	0.4529110
10	0.4529109601	group.10	0.4993297



genScan boundary.dist - p-value = 0.37131

	lower	category	upper
1	0.001081510	group.1	0.1227651
2	0.122765116	group.2	0.2001852
3	0.200185174	group.3	0.2909930
4	0.290992951	group.4	0.3781252
5	0.378125208	group.5	0.4806164
6	0.480616353	group.6	0.5672257
7	0.567225686	group.7	0.6356691
8	0.635669075	group.8	0.7596218
9	0.759621794	group.9	0.8533575
10	0.853357467	group.10	0.9994639



genScan start.dist - p-value = 0.15889

5.4 uniGene Annotations

Category limits

	lower	category	upper
1	4684.0	group.1	24682.5
2	24682.5	group.2	39421.0
3	39421.0	group.3	53523.5
4	53523.5	group.4	72479.0
5	72479.0	group.5	88359.0
6	88359.0	group.6	113194.0
7	113194.0	group.7	139572.5
8	139572.5	group.8	184425.0
9	184425.0	group.9	230325.5
10	230325.5	group.10	644368.0

uniGene gene.width - p-value = 0.75324



	lower	category	upper
1	690.0	group.1	7004.7
2	7004.7	group.2	10520.4
3	10520.4	group.3	17265.9
4	17265.9	group.4	22417.6

5	22417.6	group.5	29233.0
6	29233.0	group.6	34344.4
7	34344.4	group.7	43923.5
8	43923.5	group.8	56102.0
9	56102.0	group.9	79819.1
10	79819.1	group.10	20489465.0

uniGene non-gene width - p-value = 0.27779



	lower	category	upper
1	0.0005360666	group.1	0.0739102
2	0.0739101980	group.2	0.1333521
3	0.1333520781	group.3	0.1750946
4	0.1750945709	group.4	0.2191114
5	0.2191114320	group.5	0.2702058
6	0.2702058292	group.6	0.3190237
7	0.3190236584	group.7	0.3703712
8	0.3703711563	group.8	0.4080653
9	0.4080653451	group.9	0.4529110
10	0.4529109601	group.10	0.4993297



uniGene boundary.dist - p-value = 0.37131

	lower	category	upper
1	0.001081510	group.1	0.1227651
2	0.122765116	group.2	0.2001852
3	0.200185174	group.3	0.2909930
4	0.290992951	group.4	0.3781252
5	0.378125208	group.5	0.4806164
6	0.480616353	group.6	0.5672257
7	0.567225686	group.7	0.6356691
8	0.635669075	group.8	0.7596218
9	0.759621794	group.9	0.8533575
10	0.853357467	group.10	0.9994639



uniGene start.dist - p-value = 0.15889

6 GC content

Here we study the effect of GC content on insertion. The GC content is taken from the Human Genome Draft at GoldenPath from the table

 $\verb+http://genome.ucsc.edu/goldenPath/hg17/database/gc5Base.txt.gz.$

Following the plot is a table of fitted coefficients based on splitting the GC percent data at the median.

	lower	category	upper
1	28.49609	group.1	33.75000
2	33.75000	group.2	35.34766
3	35.34766	group.3	36.48828
4	36.48828	group.4	38.07812
5	38.07812	group.5	39.43359
6	39.43359	group.6	40.94531
7	40.94531	group.7	42.75781
8	42.75781	group.8	44.94531
9	44.94531	group.9	49.17188
10	49.17188	group.10	64.10156



gcpct - p-value = 0.70992

7 Cytobands

Here we study the association of cytoband with insertion intensity. The data are obtained from

http://genome.ucsc.edu/goldenPath/hg17/database/cytoBand.txt.gz.



A formal test of significance attains a p-value of 0.41123.

References

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