Number of cells (N) = 5
Number of times variant is seen $(n)=4$

$$
f \approx 0.5-\sqrt{0.25-n / N^{2}}=0.2
$$



## Germline variant example

## Calculation of mosaic score:

Number of cells carrying mutation $\left(N^{\prime}\right)=f * N=0.2 * 5=1$
So, we take the 1 row with maximum hits and count it
$n_{m}=\sum_{i=3} n r_{i}=0$
$n_{m}=\sum_{i=1} n r_{i}=\sum_{i=2,} n r_{i}=\sum_{i=3} n r_{i}=\sum_{i=4} n r_{i}=\sum_{i=5} n r_{i}=1$
We take the maximum possible $n_{m}$
Mosaic score $=n_{m} / n=\frac{1}{4}=0.25$

## Calculation of germline score:

Number of cells not carrying germline variant $\left(N^{\prime}\right)=f * N=0.09 * 11=1$
So, we take the 1 column with maximum hits and count it

$$
\begin{aligned}
& n_{g}=\sum_{i=1,} n c_{i}=\sum_{i=2} n c_{i}=\sum_{i=4} n c_{i}=\sum_{i=5} n c_{i}=0 \\
& n_{g}=\sum_{i=3} n c_{i}=4 \quad \begin{array}{l}
\text { We take the maximum possible } n_{g} \\
\text { Germline score }=n_{g} / n=\frac{4}{4}=1
\end{array}
\end{aligned}
$$

Fig S3. Example of calculating mosaic and germline scores for a germline variant.

