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Supplementary appendix

This appendix formed part of the original submission and has been peer reviewed. We post it as supplied by the authors.

Supplement to: Piel FB, Patil AP, Howes RE, et al. Global epidemiology of sickle haemoglobin in neonates: a contemporary geostatistical model-based map and population estimates. *Lancet* 2012; published online Oct 25. [http://dx.doi.org/10.1016/S0140-6736\(12\)61229-X](http://dx.doi.org/10.1016/S0140-6736(12)61229-X).

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Web Appendix 1. HbS database and allele frequency map.

1. Database of HbS allele frequency survey data

Data sources

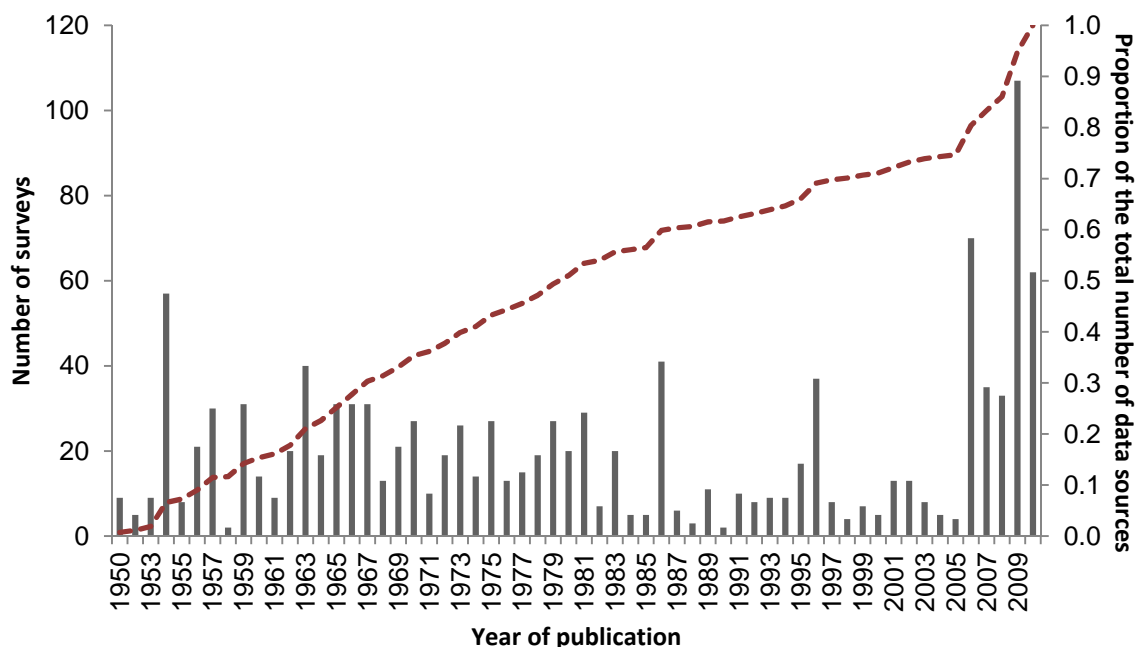
Systematic searches of the literature published since 1950 were undertaken following procedures developed by the Malaria Atlas Project (MAP, <http://www.map.ox.ac.uk>)¹⁻³ to create a contemporary database of available sickle haemoglobin (HbS) allele frequency surveys. Preliminary searches were conducted on 12 December 2007, using the following online bibliographic archives: PubMed (<http://www.pubmed.gov>), ISI Web of Knowledge (<http://isiwebofknowledge.com>) and Scopus (<http://www.scopus.com>). The keyword string used was: “sickle cell” or “haemoglobin S” or “hemoglobin S” or “Hb S”. Searches were last run on 20 October 2009. A total of 18,336 (in *Text terms*), 28,908 (in *Title/Keywords/Abstract*) and 22,732 (in *Article Title/Abstract/Keywords*) references were found in the three respective databases and exported using bibliographic management software (Endnote X4, Thomson-Reuters, Carlsbad, CA, USA). The 2,220 references from Livingstone's extensive, but out of date, database on frequencies of haemoglobin variants⁴ were then merged. Manual duplicate removal and abstract reviews of the amalgamated search results identified 3,531 references likely to contain data allowing calculation of HbS allele frequency. Despite using a conservative approach, the vast majority (91.5%) of the 41,445 unique references identified were irrelevant as they consisted of animal studies, case reports, technical descriptions of new protocols and methods, or reviews and general articles not including any primary data. Full text searches were conducted for each of the relevant references identified. When possible, authors were contacted for additional information concerning their studies in order to obtain missing information necessary to assess the quality of their results. In addition, direct contact with researchers working in Cambodia, Indonesia, Mali, Sri Lanka and Venezuela allowed inclusion of various sets of unpublished data. Finally, data collected by the Malaria Genomic Epidemiology Network Consortium (MalariaGEN, <http://www.malariagen.net>), representing a valuable source of standardised data from several malaria endemic countries,⁵ were also incorporated in the database.

Contributors are acknowledged on the MAP website (<http://www.map.ox.ac.uk/inherited-blood-disorders/acknowledgements/>). All sources from which data met the criteria for inclusion are listed in Web Dataset. Data are available on the MAP website (<http://www.map.ox.ac.uk>).

Inclusion criteria

Only population samples representative of the local communities were included. Studies where the sampling method was not explicitly stated to have been random or universal screening could be considered representative so long as other information in the article did not indicate that the sample was biased. Detailed data given for all ethnic groups found in an area (based on the information provided in the original source) were recorded as separate entries but spatial duplicates. Surveys targeting only specific ethnic groups (e.g. African Americans in North America) were excluded as in most cases the proportion of that particular ethnic group in the general population was unknown, as well as the allele frequency in the other ethnic groups present at the sampling site. Surveys focusing on hospital patients (e.g. minor ailments, fever or malaria) were also excluded, as they may represent biased estimates of the frequency in the general population. When several surveys were conducted in the same area, we kept only the most informative one (using a combination of year of survey, sample size and sampling and diagnostic methods criteria). The temporal distribution of surveys (not taken into account in our model due to the limited amount of data available) included in the model is shown in Web Figure 1.

Web Figure 1 Temporal distribution of the surveys included in the model (n=1,211). Vertical grey bars indicate the number of data sources for each year, shown on the left-hand axis. The red dashed line corresponds to a cumulative proportion of the total number of data sources, quantified on the right-hand y-axis of the plot.

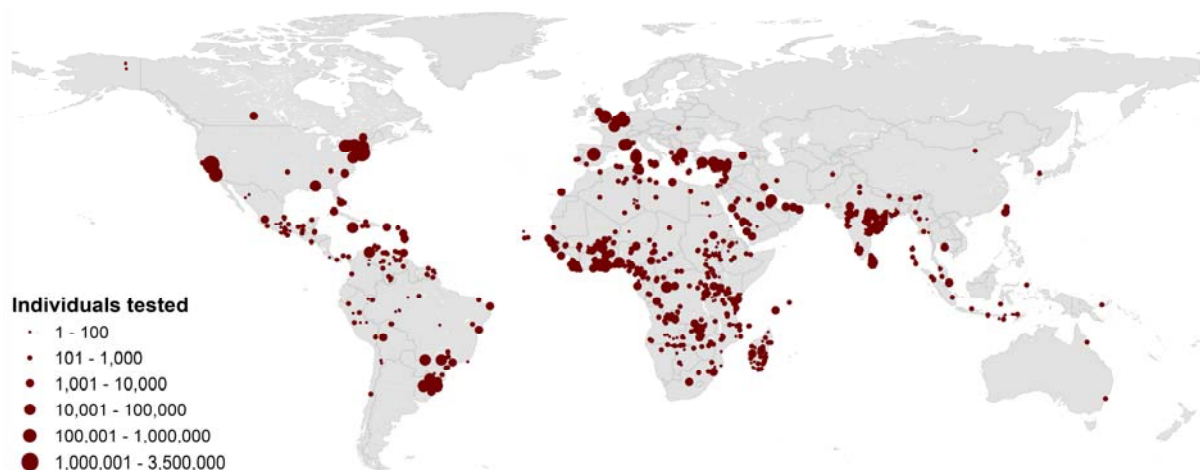


No constraints were placed on sample size (Web Figure 2), as there is no rationale for excluding small sample sizes in a model-based geostatistical (MBG) framework (see below). This is because, on one hand the model weighs the information content of each survey in accordance with a binomial sampling model, therefore

down-weighting the information from very small samples, and on the other hand the uncertainty in relation to the sample size is explicitly modelled by this technique.⁶ Nevertheless, case reports were excluded from this study as they did not match our criterion for representativeness of the local communities. Web Figure 3 shows both the temporal distribution of surveys and their sample size for each country within each WHO region (see Web Figure 8).

The number of normal (HbA or *neg*) and abnormal (HbS or *pos*) alleles observed was used as input for the model. For example, in a sample of N individuals tested in which n_{AA} , n_{AS} , n_{SS} were found to be AA, AS and SS individuals respectively, HbA allele frequency is $p = \frac{(2*n_{AA})+n_{AS}}{2N}$ and HbS allele frequency is $q = \frac{(2*n_{SS})+n_{AS}}{2N}$. Individuals identified as sicklers in earlier studies were all considered as carriers. We assumed all populations were at Hardy-Weinberg equilibrium (HWE).^{7,8}

Web Figure 2. Sample size of the HbS surveys. The size of the circles is proportional to the number of individuals tested.



Although compound individuals for haemoglobin C (HbC) and β -thalassaemia^{9, 10} are also of clinical and public health significance, we deliberately did not include these data in the present study. Limited data currently available on HbC¹¹, the complexity of the multiple variants for β -thalassaemias and the small number of individuals affected relative to HbAS, make the inclusion of these conditions difficult in the present modelling framework.

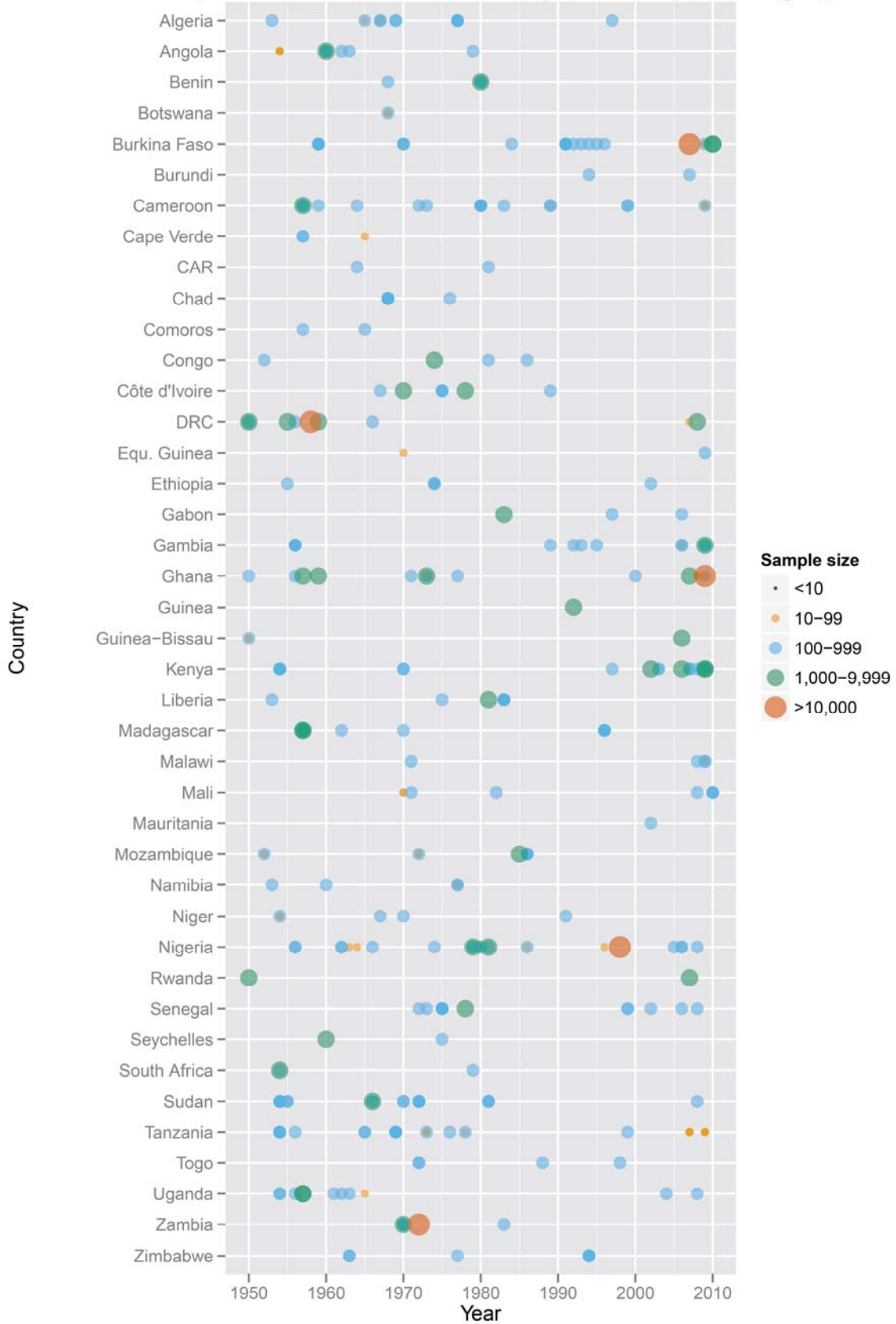
Georeferencing

The geographic location of each survey was determined as precisely as possible using a georeferencing protocol adapted from Guerra et al.³ Authors' descriptions of survey sites were used to locate the sampling

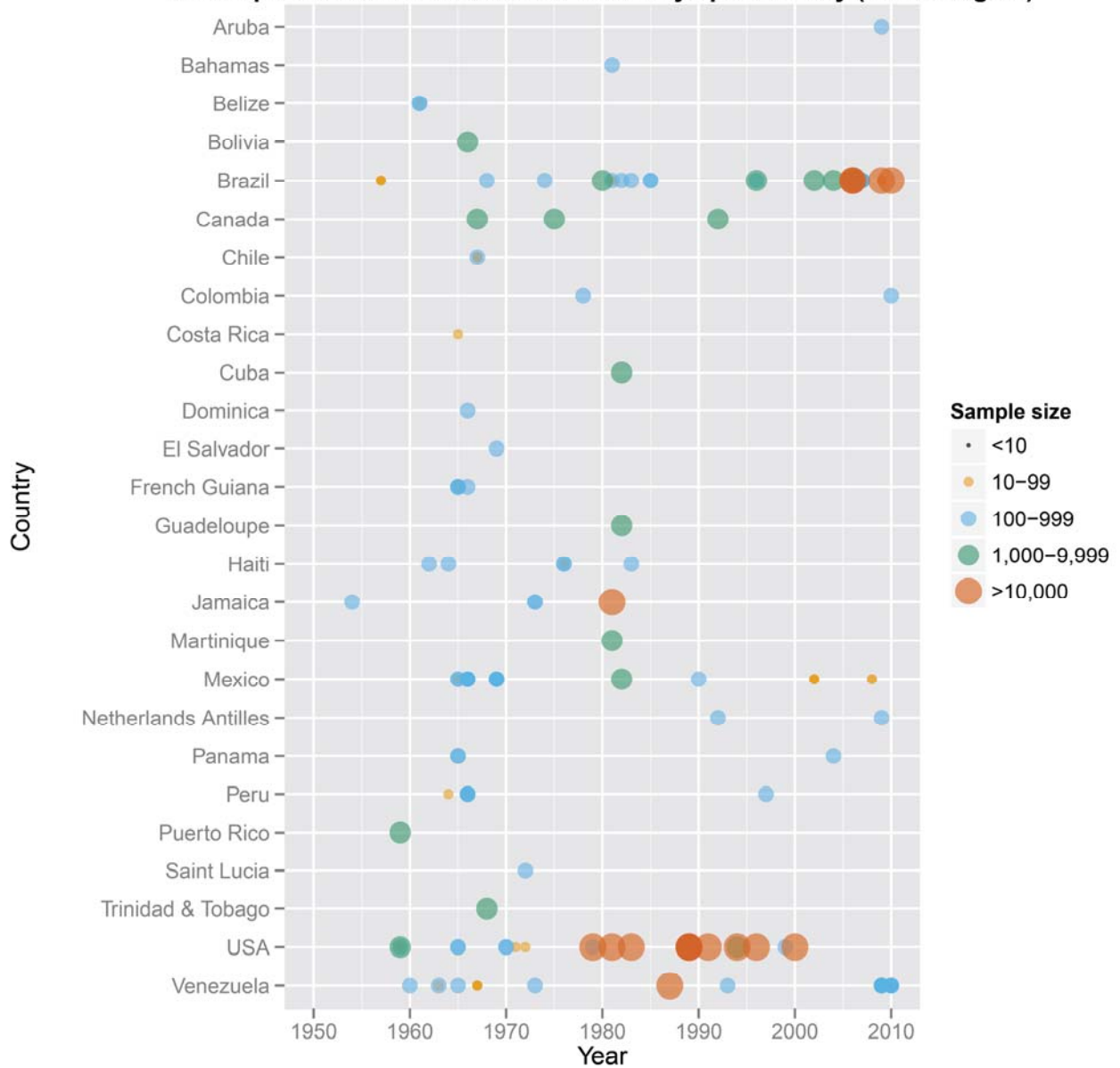
sites. Geographic coordinates (in decimal degrees) were identified in various global gazetteers including the Encarta Reference Library 2007 (Microsoft Corporation, Redmond, WA, USA), Geonames (National Geospatial-Intelligence Agency. <http://geonames.nga.mil/ggmagaz/>) and Global Gazetteer Version 2.2 (Falling Rain Genomics Inc. <http://www.fallingrain.com/world/index.html>). Surveys were categorised according to the extent of the area that they represented: points ($\leq 10\text{km}^2$), wide areas (>10 and $\leq 25\text{km}^2$), and small (>25 and $\leq 100\text{km}^2$) or large polygons ($>100\text{km}^2$). Polygons were digitised and centroids calculated in Geographic Information Systems (GIS) software (ArcView 3.2 and ArcMap 10.0, ESRI Inc., Redlands, CA, USA). A similar method was applied to surveys which could only be georeferenced to the district (admin2 unit) level. Surveys reported only to province (admin1 unit) or country (admin0 unit) level were considered to lack sufficient geographical specificity and were thus excluded. The geographic coordinates (latitude and longitude) were used as input in the model.

Web Figure 3. Temporal distribution and sample size of the HbS surveys per country within each WHO region. A. AFRO region, B. AMRO region, C. EMRO region, D. EURO region, E. SEARO region and F. WPRO region.

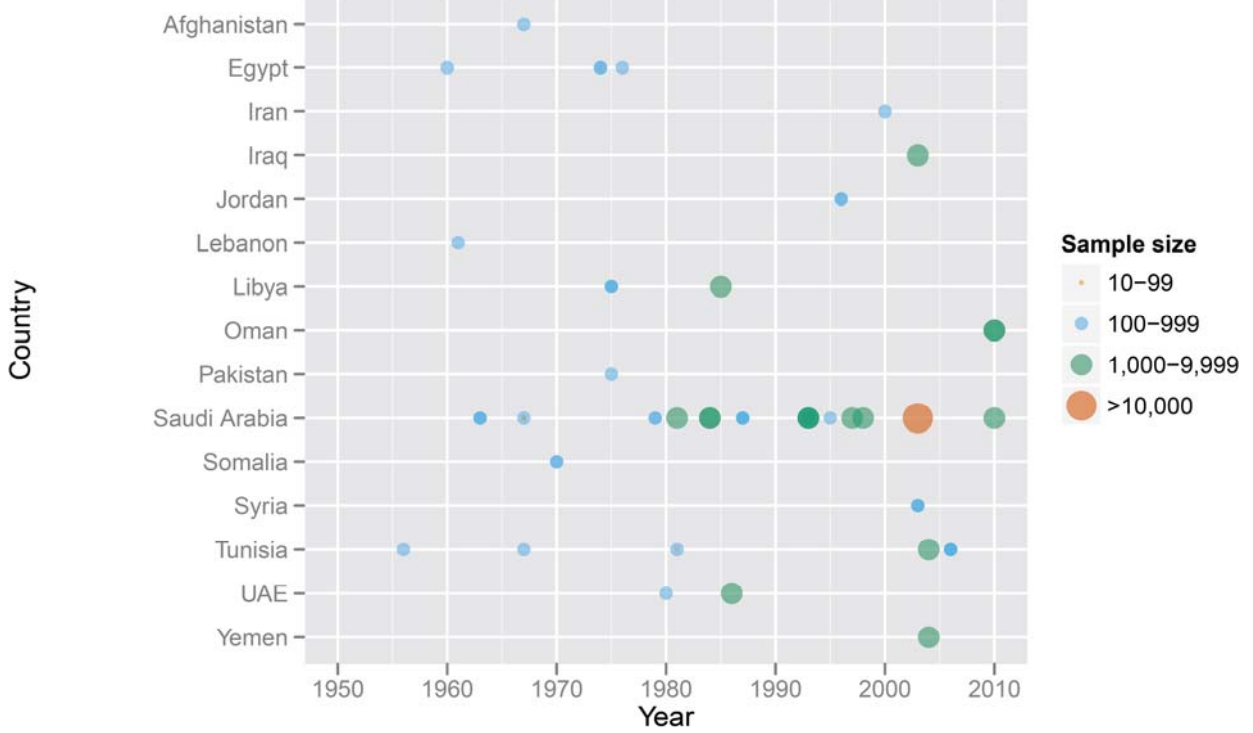
A. Temporal distribution and size of surveys per country (AFRO region)



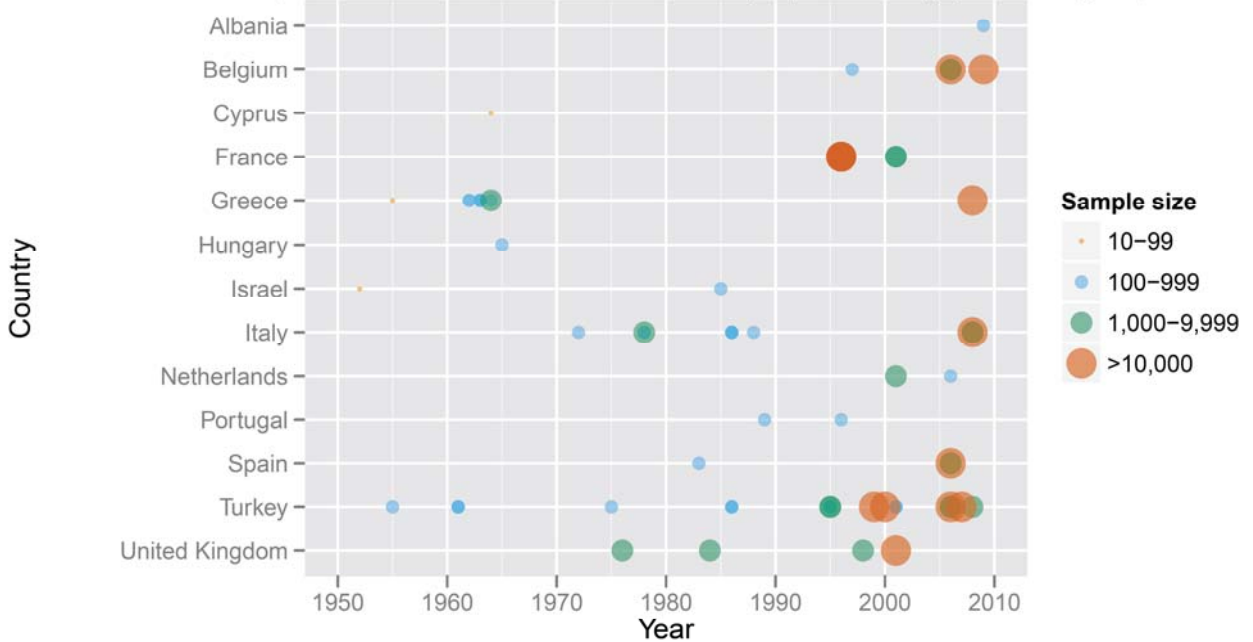
B. Temporal distribution and size of surveys per country (AMRO region)



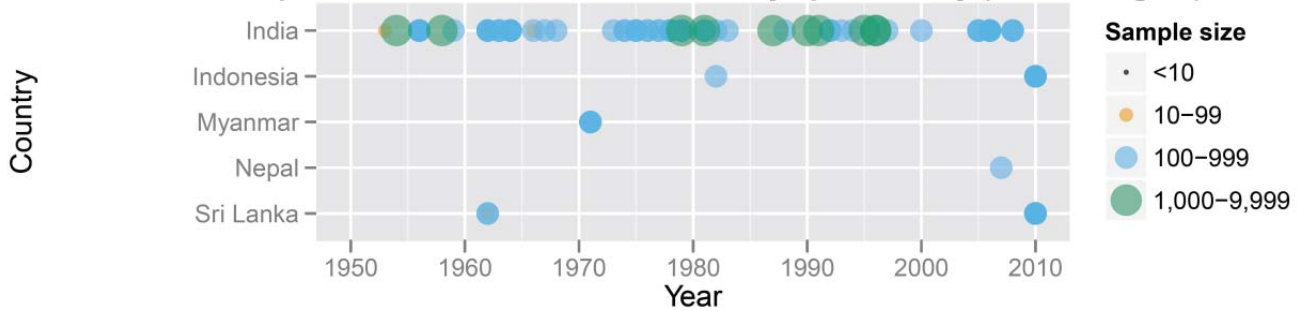
C. Temporal distribution and size of surveys per country (EMRO region)



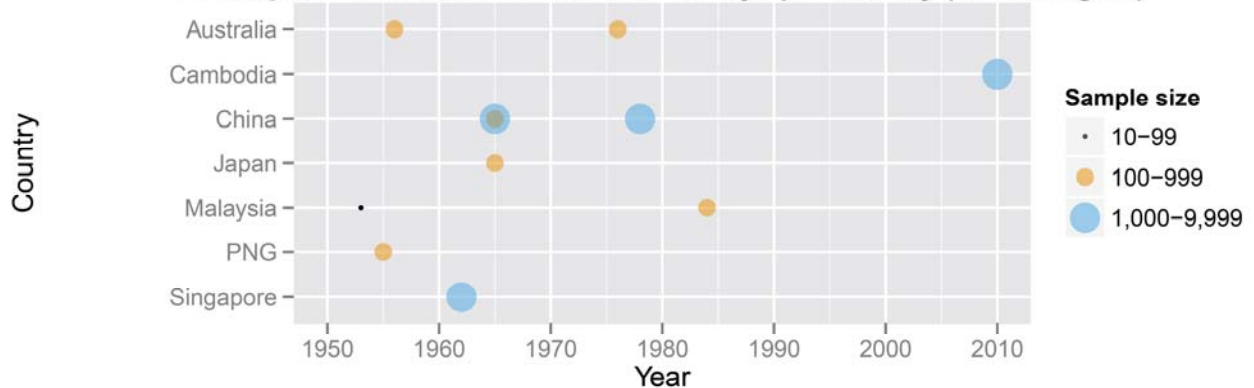
D. Temporal distribution and size of surveys per country (EURO region)



E. Temporal distribution and size of surveys per country (SEARO region)



F. Temporal distribution and size of surveys per country (WPRO region)



2. Creating a map of HbS allele frequency

Model

In this section, we describe our Bayesian spatial model for the HbS allele frequency surface $S(\cdot)$. S takes as its argument an arbitrary location on the Earth's surface. Because of the scale of this global study, the limited number of datapoints ($\sim 1,000$) and their relatively good spread, we assumed that the sampling bias towards areas of high-prevalence was reasonable. The posterior $[S]$ induces a posterior $[SS]$ for the SCA frequency surface $SS(\cdot)$ since, using the Hardy-Weinberg assumption^{7, 8} that an individual's two copies of each allele are chosen independently from a gene pool, $SS(x) = S(x)^2$. We computed summaries of $[SS(x)]$, such as $E(SS(x))$ and $Var(SS(x))$, at each location x , to produce the maps related to SCA frequency in neonates.

The model differs from the model employed by Piel et al.¹ because, while conducting the analysis for the current study, we diagnosed a lack of fit in our previous model that did not have a substantial effect on the main summaries of interest of $[S]$, but did cause serious errors in $[SS]$. This point is discussed below.

The national-level disease burden $D(A)$ in nation A can be computed from SS , the birth rate b_A in A and the global population density surface N using the areal integral

$$D(A) = \int_A b_A N(x) SS(x) da$$

This is a deterministic transformation of SS and therefore of S , so theoretically the posterior $[S]$ induces the posterior $[D(A)]$. However, for reasons discussed by Patil et al.,¹² it is prohibitively computationally expensive to sample from this posterior. We produce approximate samples using a method described below.

Prior for S

We model S as a nonlinear transformation of a Gaussian random field¹² $f(\cdot)$, plus a random field $\varepsilon(\cdot)$ that associates an independent normally distributed value with each location on the earth's surface. Specifically,

$$S(x) = g(f(x) + \varepsilon(x))$$

The link function g maps the random variable $f(x) + \varepsilon(x)$, which can be any real number, to the interval $(0,1)$, so $S(x)$ can be used as a probability or prevalence. We used a non standard link function, which is described below.

The prior for f is parameterized from the constant mean function $M(x) = m$, and the standard exponential covariance function $C(x, y) = \varphi^2 \exp\frac{|x-y|}{\theta}$ with amplitude parameter φ and range parameter θ , with suitable priors assigned to the scalar parameters m , φ and θ :

$$p(m) \propto 1$$

$$\varphi \sim \text{Exponential}(.1)$$

$$\theta \sim \text{Exponential}(.1)$$

$$f \sim \text{GP}(M, C)$$

GP indicates a Gaussian process. The units of x , y and θ are earth radii, and m and φ are unitless. The unstructured component $\varepsilon(x)$ is modelled as normally distributed with unknown variance V :

$$V \sim \text{Exponential}(.1)$$

$$\varepsilon(x) \stackrel{iid}{\sim} \text{Normal}(0, V, \alpha)$$

A global approach was preferred to a region-specific version of the model because of the limited number of datapoints available for this study.¹³ Furthermore, we privileged model stability over flexibility of the model parameters.

Although distances were calculated across oceans, the range parameter of the model was smaller than typical ocean width. Predictions on the eastern coast of Brazil were therefore not influenced by datapoints from West Africa, but strictly by Central and South American datapoints.

Likelihood

Adopting the assumption of Hardy-Weinberg equilibrium,^{7, 8} if n_i individuals are sampled at the i 'th observation location o_i (for a total of $2n_i$ chromosomes), the probability distribution for the number k_i of copies of the HbS allele that will be found is binomial, with probability $S(o_i)$:

$$k_i \sim \text{Binomial}(2n_i, S(o_i))$$

Flexible link function and empirical Bayesian analysis

The link function g for binomial data is usually taken to be the inverse logit function,

$$g(x) = \text{logit}^{-1}(x) = \frac{\exp x}{1 + \exp x}$$

Piel et al.¹ employed this model. Applying the change of variables formula,¹⁴ the induced prior for $S(x)$ is:

$$p(S(x)) = \frac{1}{S(x)(1 - S(x))} \text{Normal}(\text{logit}(S(x)); m, \varphi^2 + V)$$

Note that this is essentially a two-parameter family of probability distributions, since φ and V appear only in the sum.

When we initially attempted to fit Piel et al.'s model to the current dataset and predict SS , we found that, when the local distribution above is fitted in areas where datapoints are highly clustered, the best fitting values of m and $\varphi^2 + V$ result in implausibly long right-hand tails for the predictive distribution of prevalence in the next observation at x . Although the standard summary statistics, including the upper 95% credible interval, were consistent with the local dataset, strikingly high allele frequencies $S(x)$ (greater than 40%) were predicted with small but practically significant probability ($\sim 0.1\%$).

This particular type of lack of fit was not a major issue for predictions of $S(x)$ because the bulk of the predictive distribution was roughly consistent with the dataset, but the long right-hand tail translated to an even longer right-hand tail for $SS(x)$, which contained enough mass to skew all of the standard summary statistics. For example, the predicted mean of $SS(x)$ in some areas exceeded 5%, which is higher than all but the observed values in the dataset.

To remedy this problem, we attempted several strategies including employing Stukel's link function¹⁵ in place of the more standard inverse logit, and modelling $f(x) + \epsilon(x)$ as a skew-Gaussian process¹⁶ rather than the standard Gaussian. The skew-Gaussian approach indicated that it could solve the problem, but eliminated the crucial conjugate relationship between $f(x)$ and $f(x) + \epsilon(x)$, and we were unable to devise a successful Markov chain Monte Carlo (MCMC) scheme.

Ultimately, we employed an alternative flexible link function

$$h(x) = \sum_{i=0}^3 c_i x^i$$

$$g = \text{logit}^{-1} \circ h$$

We were unable to infer c_i jointly with the other model parameters in a fully Bayesian manner due to poor MCMC mixing, so we adopted an empirical fitting approach inspired by data pre-processing steps employed in classical geostatistics.

Empirical Bayesian approach to fitting the polynomial coefficients

For each observation (n_i, k_i) where n_i is the sample size and k_i is the number of HbS alleles, we first obtained the posterior expectation of the gene pool-wide prevalence of HbS with uniform prior density on $[0,1]$:

$$\hat{p}_i = \frac{k_i + 1}{2n_i + k_i + 2}$$

We discarded values where $2n_i$ was below 50. Then, we inferred the parameters \tilde{m} and \tilde{V} of the nonspatial Bayesian model:

$$p(\hat{p}) = \prod_i \frac{1}{\hat{p}_i(1 - \hat{p}_i)} \text{Normal}(\text{logit}(\hat{p}_i); \tilde{m}, \tilde{V})$$

We then plotted the posterior predictive cumulative distribution function (CDF) of $\text{logit}(\hat{p})$ against its empirical CDF, and fitted the coefficients c of the cubic polynomial function h to the points using least squares, subject to the constraint that h must be invertible (or, equivalently, monotone).

In the Bayesian analysis of the full spatial model, the fitted values of c were taken as known and fixed. Although this empirical procedure is admittedly informal, the resulting nonstandard link function did substantially improve the fit of the model to the data.

Prior predictive constraint

Relatively few datapoints are available in areas known to have very low HbS allele frequency, such as the high- and low-latitude regions. The observed allele frequency in these few observations is zero, but more zero observations would be required to rule out the possibility that an occasional area of high allele frequency occurs in these areas. As it happens, the striking local variation in allele frequency in more data-rich and higher-frequency areas in Africa leads the model to conclude that such foci probably *do* occur. Even with the more flexible model for $\varepsilon(x)$, the predictive distribution in areas of low data coverage exhibits long right-hand tails, and the predicted mean value of $S(x)$ in these regions is surprisingly high.

The ideal solution to this problem would be to locate the data behind the common knowledge that HbS allele frequency is low in the high- and low-latitude regions, such as in health service reports, and incorporate it in the model. Given the obvious logistical difficulties associated with this ideal approach, we found it more practical to supplement the dataset with expert opinion (based on authors' and technical advisory group members' knowledge of this gene).

Perhaps the best way to incorporate this expert opinion would be as 'soft data', as described by Christakos¹⁷ among others. However, producing defensible local pseudo-observations of HbS allele frequency all but requires the data collection process we sought to avoid. In addition, using soft datapoints would increase the number of spatial locations at which the Gaussian random field f has to be imputed, increasing the computational expense of fitting the model.

As a compromise, we elected to constrain φ , m , and V in such a way that the prior predictive distribution of $S(x)$, before the data are incorporated, puts probability mass of 1×10^{-3} or less on values in excess of 0.001. This constraint arguably induces a lack of fit by forcing $f(x)$ to depart from its prior mean by many standard deviations in areas where HbS allele frequency is known to be high; but it does remedy the implausibly high predictive values in the polar regions, and does not seem to adversely affect the fit in areas of nonzero allele frequency.

Fitting the model

The model was fitted using a MCMC algorithm¹⁸ implemented in the programming language Python using the Bayesian analysis package PyMC.¹⁹

The scalar parameters φ , m , θ and V were updated jointly using Haario, Saksman and Tamminen's adaptive Metropolis algorithm,²⁰ as implemented by PyMC's AdaptiveMetropolis step method. Each value $\varepsilon(o_i)$ at observation location o_i was updated separately using the standard one-at-a-time Metropolis algorithm. The

distribution of the Gaussian random field at the observation locations, $\{f(o_i)\}$, is conjugate to the distribution of $\{\epsilon(o_i) + f(o_i)\}$, so we updated $\{f(o_i)\}$ by sampling from its full conditional distribution. MCMC output parameter values based on 1,000,000 iterations are summarised in Web Table 5.

The polynomial coefficients for such a function are specific to the dataset. The set of coefficients used, corresponding to an invertible function and fitting the empirical cumulative distribution function was:

$$y = -1.48556762x^3 + 0.28125179x^2 + 0.02261485x + 0.02125477$$

The code implementing the analysis is available at <http://github.com/malaria-atlas-project/ibd-world> and <http://github.com/malaria-atlas-project/generic-mbg>. Traces for all variables are available upon request.

Mapping procedure

Interpolating spatially sparse survey data to predict an allele frequency across a wide region results in predictions for which the level of certainty (or uncertainty) varies spatially as a function of the density, quality, and sample size of survey data available. The underlying spatial heterogeneity of the frequency of the disorder studied, known to be high in some areas for HbS,¹⁻⁴ also influences this uncertainty. A Bayesian MBG framework²¹ generates a posterior predictive distribution (PPD) rather than a unique value, therefore allowing estimation of the uncertainty of the prediction for each pixel. In addition to the PPDs of HbS allele frequency, HbAS and HbSS genotype frequencies were also generated directly by the model. Because these are nonlinear functions of the allele frequency, it would be incorrect to produce summary maps of these quantities from those of allele frequency using GIS software.¹² Here, we used the mean as a per-pixel visual summary of the PPD. The summary map is shown at a 5 × 5 km resolution (Figure 2B).

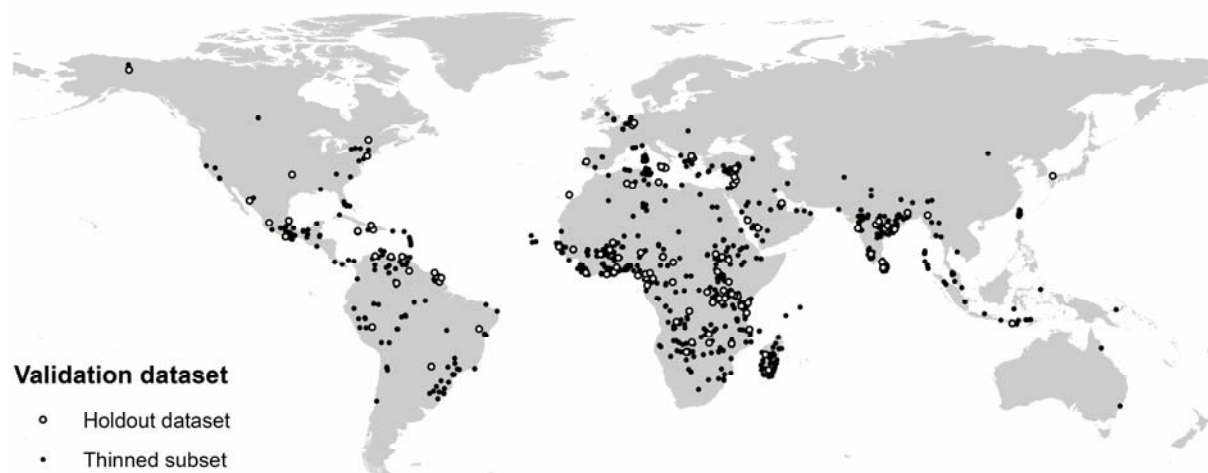
The uncertainty is a crucial measure of the accuracy of the prediction. It is based on the abundance of data, the sample sizes and the homogeneity or heterogeneity of the data around a pixel. From the range of possible uncertainty intervals available from the model's output, we chose here to use the interquartile range (IQR) of the posterior distribution,²² corresponding to a 50% probability (Figure 2C). This corresponds to the *mbg-map* command of the generic MBG package.

Validation and model performance metrics

In addition to the model-based representations of prediction uncertainty provided by the MBG framework, the model's predictive ability was quantified by assessing the disparity between the prediction and the observed allele frequency using a validation subset of the data. Ten percent of the data (n=121), randomly selected, were held out from the dataset (Web Figure 4). The model was run in full using the thinned data

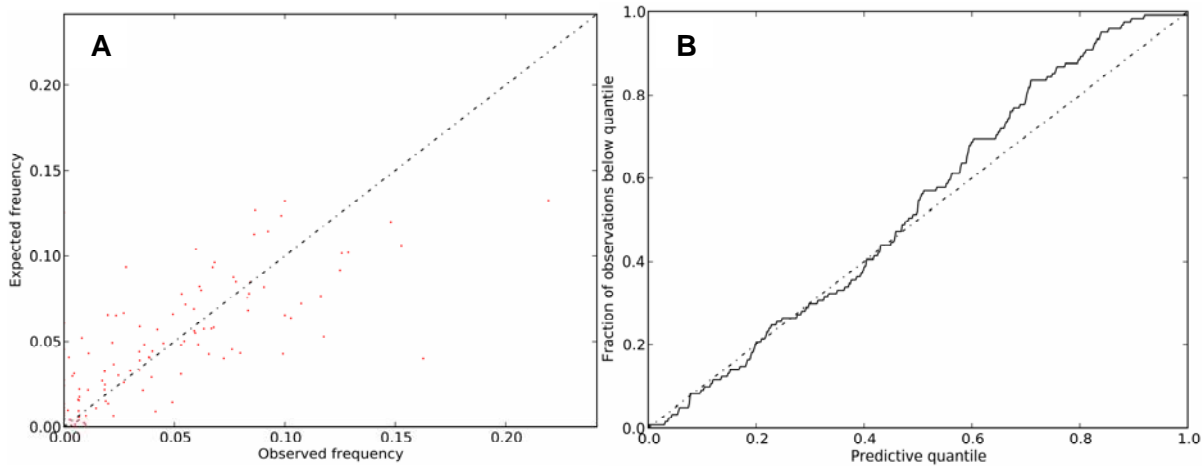
set (n=1,090) to generate HbS PPD for comparing the mean of the prediction with known values at the locations of the held-out data (Web Figure 5A).

Web Figure 4 Distribution of the validation subsets. Black dots represent datapoints used in the thinned dataset. Circles represent datapoints held out for the model validation.



The prediction's mean error (ME) and mean absolute error (MAE) were used to assess the model's overall bias and overall accuracy respectively. The ME is the average distance between the actual datapoints and the predicted values. The MAE is a measure of the average magnitude of the errors in the predicted values. A procedure was also implemented to test the extent to which PPDs at each location provided a suitable measure of uncertainty.^{23, 24} Working through 100 progressively narrower credible intervals (CIs), from the 99% CI to the 1% CI, each was tested by computing the actual proportion of held-out prevalence observations that fell within the predicted CI. In a perfect model, 95% of true values should fall within the 95% CI predicted at each location, 50% within the 50% CI, and so on. Plotting these actual proportions against each predicted CI level allows the overall fidelity of the posterior probability distributions predicted at the held-out data locations to be assessed (Web Figure 5B). This corresponds to the *mbg-validate* command of the generic MBG package.

Web Figure 5 Validation plots comparing the prediction with the observed allele frequency for the datapoints from the hold-out subset of the data (n=121). A. Scatter plot of the observed vs. predicted allele frequency; B. Plot of the observed vs. predicted quantiles.



3. Historical vs. contemporary maps

A map of the historical distribution of sickle haemoglobin has been published.¹ Although the methodology used to assemble the dataset, to identify surveys fulfilling inclusion criteria, to georeference the location of the survey and to create a continuous map within a Bayesian geostatistical framework are broadly similar to the methodology described here, fundamental differences exist between these two maps.

The historical map, published in 2010,¹ is based only on population samples which were representative of an indigenous population. We considered populations surveyed to be indigenous, if no information was available from the author(s) to indicate that the population did not evolve locally in relation to the historical prevalence of malaria. Non-native populations surveyed in the Americas or Western Europe, for example, were therefore excluded. A total of 773 surveys were included. This map aims at describing the distribution of sickle haemoglobin as a result of the selection pressure of malaria on the genetics of human populations.

The contemporary map, presented in this paper, is based on all representative population samples, whether indigenous or not. A much larger total of 1,211 surveys were included. This map aims at describing the current distribution of sickle haemoglobin as a result of both malaria selection and human diasporas, in order to estimate the number of individuals affected in 2010 at national, regional and global scales. This map is a continuous global surface of the frequency predictions made within our Bayesian geostatistical framework; frequency predictions combined with demographic data then allowed calculating population estimates and their associated uncertainty.

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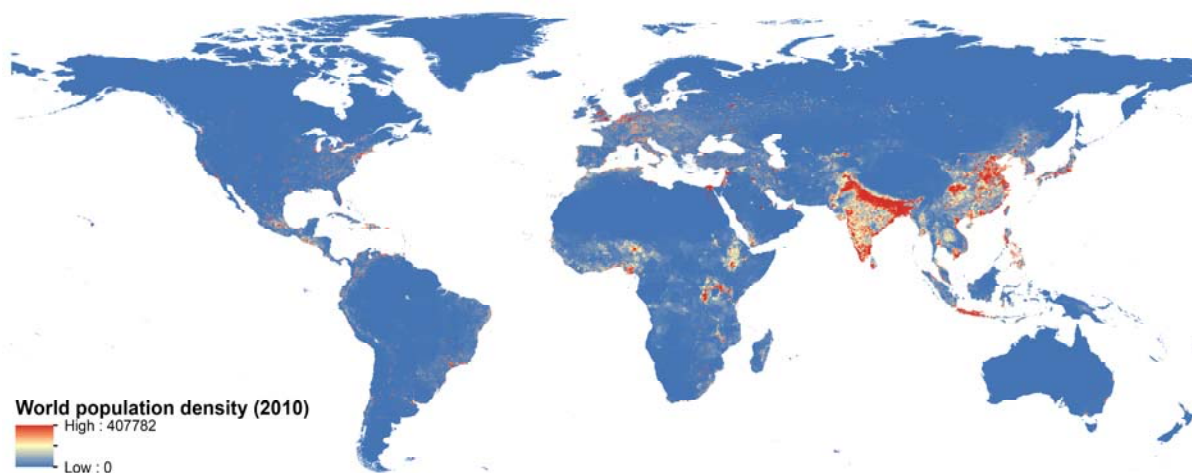
Web Appendix 2. Newborn estimates.

The best way to know with precision the number of neonates affected by a genetic disorder (such as sickle haemoglobin) at the national level is to conduct a universal national screening programme at birth. Although such programmes have proven their benefits,¹⁻³ they are in place only in a handful of countries, none of which are in Africa where the most affected populations are living.⁴ In addition to adding data from recent surveys and refining the modelling methodology (see Web Appendix 1), using the highest resolution available for demographic data can help refining estimates of neonates affected.

1. Demographic data

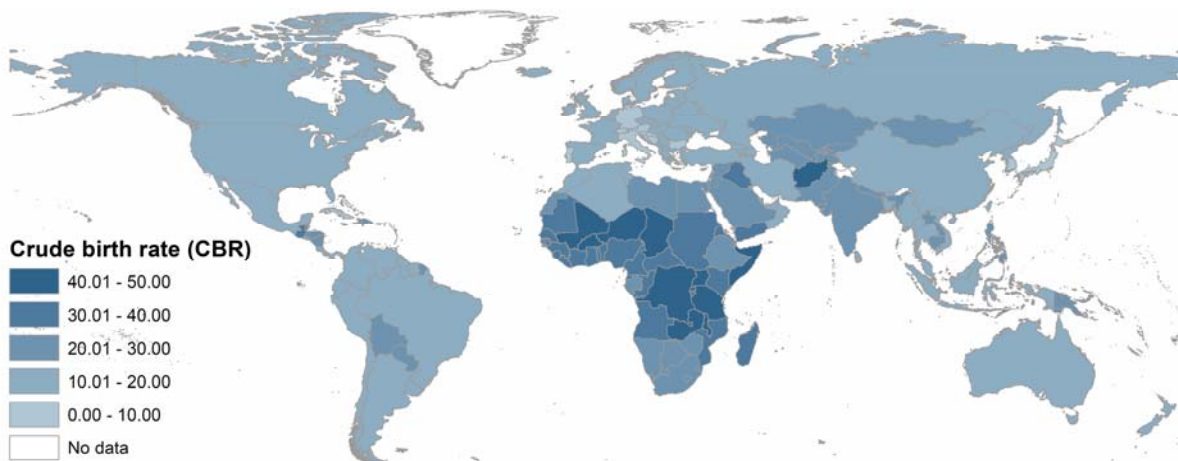
Both HbS allele frequency and population density are highly variable between pixels within countries. The HbS allele frequency map aims to include spatial heterogeneities revealed by population surveys. National estimates therefore depend on whether areas of high or low frequencies are highly populated or not. Rather than using crudely averaged data for each country, we combined our contemporary allele frequency map with high resolution population data to deal with this issue. The Global Rural Urban Mapping Project (GRUMP) *beta* version provides gridded population counts and population density estimates at 1×1 km spatial resolution for the years 1990, 1995 and 2000, both adjusted and unadjusted to the United Nations national population estimates.⁵ We used adjusted population counts for the year 2000 projected to 2010 described previously.⁶ The urban growth rates were applied to populations residing within the GRUMP-defined urban extents, and the rural rates were applied elsewhere. National 2010 totals were then adjusted to match those estimated by the United Nations.⁷ Population density is shown in Web Figure 6.

Web Figure 6. Global human population in 2010. Number of individuals per pixel (1x1 km) from the Global Rural and Urban Mapping Project, beta version.



Mortality data are currently insufficient for calculating prevalence estimates across whole populations, so we focus here on neonates using Hardy-Weinberg assumptions.^{8, 9} Assuming random mating and large population sizes, it is possible to estimate the HbS allele frequency and the proportions of each genotype (AA, AS and SS) from the number of heterozygote individuals observed in the population sample.¹⁰ Conceptually, the number of AS and SS babies born per year can be obtained by multiplying a function of HbS allele frequency ($2p(1 - p)$ and p^2 respectively) by the population living within the area of interest and the crude birth rate (CBR). CBRs are not consistently available globally at a finer resolution than the country level, hence the use of data from the United Nations Population Prospects for the 2010-2015 period¹¹ (Web Figure 7).

Web Figure 7. Crude birth rate per country. Data from the United Nations World Population Prospects: The 2010 Revision Population Database (<http://esa.un.org/unpd/wpp/>). The rates are for 2010 and are expressed in neonates per thousand (000s).



2. Areal predictions

As described previously,¹² one needs to be careful when predicting integrals over spatial areas. Using traditional geographic information system (GIS) methods, a researcher having access to a map of HbS allele frequency ($S(x)$) and desiring a national proportion of individuals with the SS genotype ($SS(x)$), would take the square of $S(x)$ and then average the values over the various pixels falling within the country of interest, weighted by population (Web Figure 6). This approach has limitations when the map of allele frequency is uncertain. Squaring the mean map for $S(x)$ does not yield the mean map for $SS(x)$ and it is impossible to produce any assessment of the uncertainty of the areal average from summary maps alone.¹²

To develop a correct procedure for producing predictive distributions for national proportions, we began by considering what we would do if we had the true map of HbS allele frequency in hand. As stated above, the national-level disease burden $D(A)$ in nation A can be computed from S , the birth rate b_A in A and the global population density surface N using the areal integral:

$$D(A) = \int_A b_A N(x) S(x)^2 da \quad (1)$$

In reality, the allele frequency map S is unknown. We do not know its exact value, but we have a posterior distribution $[S]$ for it, from which samples can be drawn. Because applying equation 1 to a sample from $[S]$ generates a sample from $[D(A)]$, many samples from $[S]$ can be used to build up a histogram approximating $[D(A)]$.¹² Summaries such as the mean, median, variance and credible intervals can be approximated using these samples.

Although it is mathematically correct, this procedure is impractical to implement. We have an approximation of $[S]$ in the form of the MCMC trace, but generating samples from it at an appropriately high resolution is extremely computationally expensive.¹² We use an approximate procedure based on the fact that, if z_1 is a single-element binomial process on A with intensity d ,

$$\frac{\int_A S(x)^2 d(x) da}{\int_A d(x) da} = E(S(z_1)^2) \quad (2)$$

Furthermore, if z_i is an l -element binomial process on a with intensity d ,

$$E(S(z_1)^2) = \lim_{l \rightarrow \infty} \frac{1}{l} \sum_{i=1}^l g(S(z_i)^2) \quad (3)$$

The expectation of the term inside the limit is equal to the left-hand term, but its variance is smaller than that of $S(z_1)$.

The pseudocode for our procedure, based on this approximation, was as follows:

1. Generate an l -element binomial process on A , z_i , with intensity N .
2. For each value in the thinned MCMC trace for the scalar parameters and the Gaussian random field f evaluated at the observation locations $\{o_i\}$, $\{f(o_i)\}$,
 - o Draw a value for the l -element random vector $\{f(z_i)\}$ from its full conditional distribution.
 - o Convert these values to $\{S(z_i)\}$ by applying the inverse-logit link function.

- Square these values to obtain a sample for the value of the desired genotype frequencies $\{S(z_i)^2\}$.
- Compute the arithmetic mean of this sample and store.

To estimate the Monte Carlo standard error (SE) attributable to the use of a set of l spatial locations rather than a high-resolution raster grid, we repeated all computations ten times and recorded the sample standard deviations of all summaries (mean, median, etc.). The point estimates that we report were obtained by aggregating the samples from all repetitions.

This procedure was conducted using $l = 30,000, 10,000, 1,000$ and 500 spatial points for the global, regional, national and sub-national areal estimates respectively; with 10% of the parameter samples in the dynamic trace, selected at random; and 1,000 iterations. As this resulted in a full PPD for each areal unit of interest, various parameters could be used to summarize the predicted estimates and their uncertainty. Here, we used the median and the interquartile range (IQR).

The modelled surfaces and population estimates presented here are made available in the public domain with the publication of this article via the Malaria Atlas Project (MAP) website (<http://www.map.ox.ac.uk>), with maps available at the global scale and population estimates at global, regional and national scales. The creation of an interactive online resource for open-access distribution of HbS modelled surfaces and estimates of newborns affected, is part of the ongoing development of the ROAD-MAP online database.

3. Model validation and performance metrics

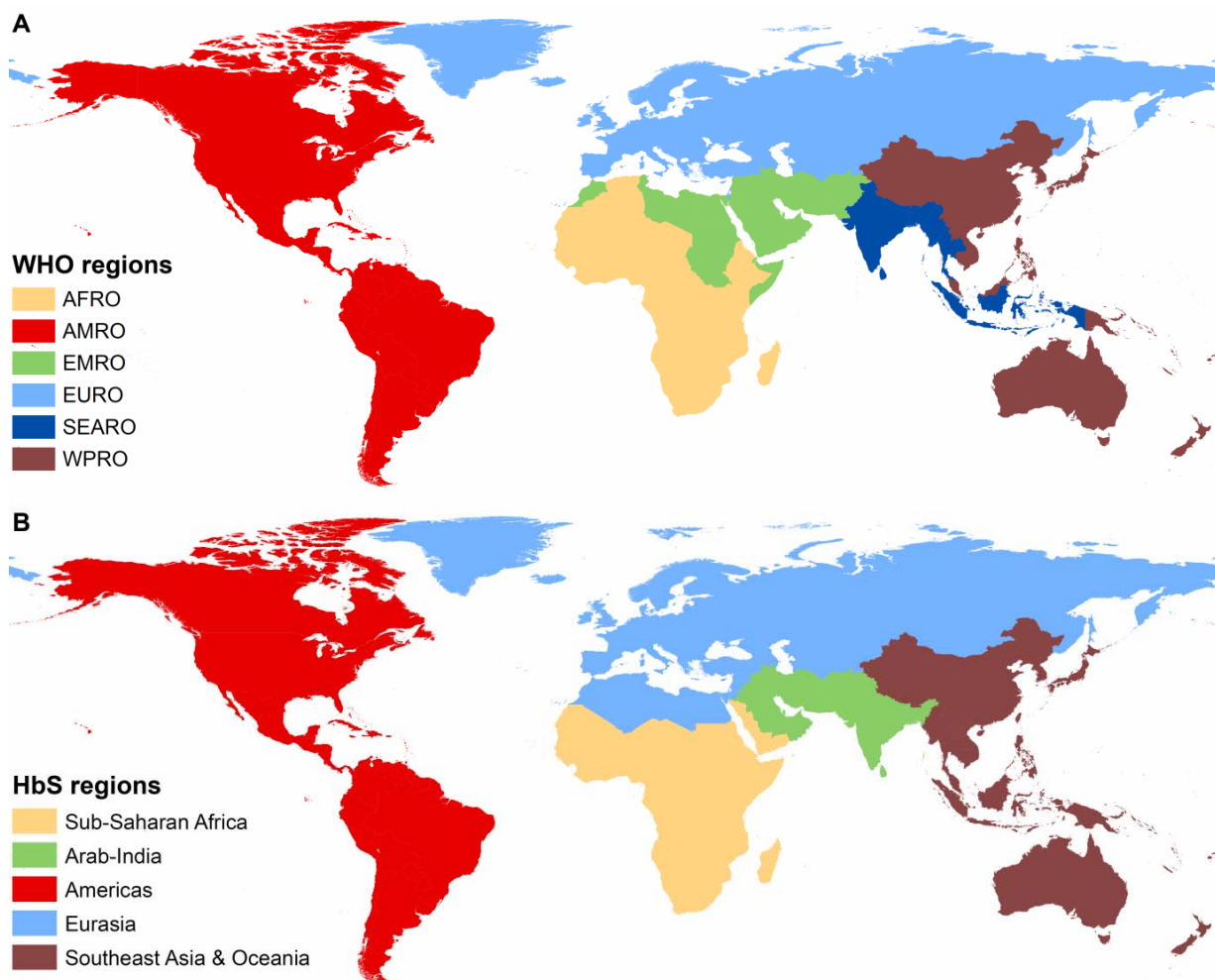
In order to quantify the error in our areal estimation process, we performed the procedure described above ten times. Each repetition used a different subset of the dynamic tracefile and was therefore independent from previous repetitions. This method allowed us to calculate the Monte Carlo SE^{13, 14} for any summary parameter chosen. The results are summarised in Table 1, Web Table 2 and Web Table 3 for the global, national and sub-national estimates respectively.

4. The use of modified WHO regions

The historical distribution of sickle haemoglobin has been explained by the selective advantage that it confers against malaria. Frequency maps of historical malaria and HbS frequencies therefore correlate on a global scale.¹⁵ Recent changes however, are unbalancing this relationship, both in malaria-endemic and non-endemic regions. Many historically malaria endemic regions are undergoing epidemiological shifts, and it is

projected that genetic disorders in these regions will have increasingly prominent public health burdens in future years.¹⁶⁻¹⁸ Increases in genetic disorder burdens are also expected in non-malaria endemic regions as human migrations continue to import the sickle gene into these regions. It is important, therefore, to consider estimates of sickle haemoglobin within these epidemiologically-relevant regions. Maps of malaria endemic countries have been recently published by the MAP.^{19,20} In addition, several haplotypes for the HbS mutation have been identified and designated by their geographic area of origin: Senegal, Benin, Cameroon and Arab-Indian.²¹⁻²³ Based on current knowledge, it seems that the Arab-Indian haplotype, which has been observed in India and the eastern part of Saudi Arabia and surrounding countries, has milder clinical severity than the African haplotypes.^{21,24}

Web Figure 8. WHO and HbS regions. A. WHO regions. AFRO: regional office for Africa; AMRO: regional office for the Americas; EMRO: regional office for Eastern Mediterranean; EURO: regional office for Europe; SEARO: regional office for South-East Asia; WPRO: regional office for the Western Pacific. See <http://www.who.int/about/regions/en/index.html> for further detail. B. HbS regions. WHO modified regions based on sickle cell haplotypes²¹⁻²³ and malaria endemicity.²⁰



Because of the influence of these two factors on the dynamics, and complications related to HbS respectively, which both have implications for public health policies, we have presented our estimates within modified WHO regions (Web Figure 8) alongside estimates within WHO regions. These regions are: i) the Americas, corresponding exactly to the AMRO region; ii) Eurasia which includes Russia, Mongolia, as well as European and other Mediterranean countries; iii) sub-Saharan Africa which also includes the western part of Saudi Arabia and Yemen; iv) Arab-India which extends from eastern Saudi Arabia and Oman to India; v) Southeast Asia, including China, and Oceania. To allow comparisons with previous estimates,²⁵ all results are presented within both WHO and modified WHO regions.

5. Comparison of national estimates with Modell and Darlison's (M&D)

We compared our national estimates of SS neonates with unpublished estimates calculated by Modell and Darlison (M&D) in 2008.²⁵ M&D aimed at presenting “the most conservative [estimates] permitted by the data (i.e. give minimum figures)”, based on demographic data from the 2003 UN Demographic Yearbook. In the present study, we aimed at providing the best summary of our current knowledge and of the precision associated with these estimates. It is therefore logical that our national estimates for 2010 tend to be higher than those calculated by M&D as illustrated in Figure 3. This can be explained by various methodological differences including: i) the use of an updated dataset based on strict inclusion criteria; ii) the use of a state-of-the-art modelling geostatistical framework, unconstrained by national boundaries and accounting for spatial heterogeneities; iii) the use of up to date demographic data; and iv) weighting each pixel estimate by population density data, so that, for example, highly populated areas with high frequencies are better represented.

We first compared absolute numbers of neonates affected. Within the AFRO region (Figure 3A), the biggest difference was in the DRC with our median estimate being approximately twice that of M&D. Their estimate for Nigeria fell within our IQR. Important differences appeared for Sudan and Kenya for which relatively large numbers of surveys (32 and 51 respectively) were used in our model. For the AMRO countries (Figure 3B), our median estimates for Brazil and the USA are approximately three times higher than M&D's. Although 47 and 25 surveys respectively were included in our model, they tended to be clustered and targeted areas of high prevalence. This contributes to the large IQR associated with our predictions. Large differences were also observed in Haiti, Mexico, Venezuela and most Central American countries. Within the EMRO region (Figure 3C), M&D's estimate for Saudi Arabia was more than five times higher than ours which has a small IQR. For Egypt, the Syrian Arab Republic and Iran, our median estimates are higher, and the IQRs were relatively large. In the EURO countries (Figure 3D), our median estimates are higher for most Western

European countries, as well as for Turkey and Israel, but IQRs are large. Within the SEARO region (Figure 3E), the only difference appeared in India for which our median estimate is 1.6 times higher than M&D's. For the WPRO countries (Figure 3F), numbers are low but differences appeared for Australia and New Zealand.

We also looked at the rate of affected neonates per thousand overall conceptions as a relative comparison measure. This rate, calculated as the ratio between the number of affected neonates (AS or SS) and the product of the population and crude birth rate within the area considered, given in per thousand (Web Table 4), can facilitate comparisons with previous and future estimates, as well as with estimates for other disorders. M&D global and regional rates were calculated using the sum of regional and national estimates respectively. MAP global and regional estimates are based on our areal predictions. These rates have also been included in Web Table 4.

6. National newborn screening programme data

Worldwide, only a few high-income, non-malarious countries, with relatively low haemoglobinopathy prevalence, have implemented a national universal newborn screening for sickle haemoglobin. These countries include the United States of America (USA), England and the French Overseas Departments and Territories. A targeted screening programme focussing on populations at risk is in place in France. Some African countries (e.g. Ghana)²⁶ have recently started to implement pilot screening programmes for haemoglobinopathies but no screening-based national estimates were available for this study.

In the USA, universal newborn screening for sickle cell homozygotes (SS), sickle cell carriers (AS) and compound heterozygotes with HbS and HbC (SC) is now either required by law, by an official rule or fully implemented in all states.²⁷ Since 2005, part of this data is centralized into the National Newborn Screening Information System (NNSIS, <http://nnsis.uthscsa.edu/>) on a voluntary reporting basis. Although incomplete, due to the absence of any reports from some states and the lack of spatial resolution, this is the only national resource currently available. Haemoglobin reports from the NNSIS included the total number of cases detected per year. Such data were available for most states from 2001 to 2010. Based on reported data, the annual number of SS neonates in the USA was 1,213 in 2001 (50 states/territories), 1,166 in 2002 (47 states/territories), 1,141 in 2003 (48 states/territories), 1,056 in 2004 (58 states/territories), 1,047 in 2005 (50 states/territories), 1,057 in 2006 (49 states/territories), 988 in 2007 (44 states/territories), 1,029 in 2008 (42 states/territories), 883 in 2009 (36 states/territories), and 783 in 2010 (32 states/territories). Using all data available, the annual average number of SS neonates over that period was 1,123. Based on the 25 population representative surveys used in our model, our estimate for the USA was 2,843 SS (IQR: 2,038-

3,968) neonates. Our estimate is therefore at least twice as high as suggested by the NNSIS reported data. Potential explanations for this include i) the relatively small number of surveys available for such a large country; ii) the heterogeneous distribution of ethnic groups mostly affected (not included in our model); and iii) potential discrepancies between data collected locally and data reported in the NNSIS data.

In England, the National Health System (NHS) Sickle Cell and Thalassaemia Screening Programme was launched in 2001. It collects data from 152 antenatal laboratories and 13 newborn laboratories.²⁸ Based on data from April 2005 to March 2007, 10,000 AS neonates were identified amongst 1,069,173 babies screened (89% of all babies born in England).¹ The report estimates that this represents 90% of the total actual number of carriers. Considering 748,990 neonates for 2010,¹¹ applying the same ratio would result in 7,783 AS neonates. No number for SS neonates was available. Based on the four population surveys included in our global model, our estimate for England was 14,509 (IQR: 10,263-20,013) AS and 269 (IQR: 150-466) SS neonates. Our estimate of AS neonates appeared therefore slightly higher than what could be expected from national screening programme data. This probably results from the opportunistic sampling of published surveys used in our model, which focus mostly on populations at risk.

In France and the French Overseas Departments and Territories, there is currently no central database to give precise figures on the number of neonates affected by HbS. Cumulative data from 1996 to 2007, based on the screening of 2,622,870 neonates, identified 64,269 AS and 2,917 SS or S β ⁰ thalassaemia babies respectively.²⁹ Using the total number of neonates in 2010 (n=832,799),³⁰ the annual number of AS neonates based on the above ratio would be 20,416 which is consistent with our estimate of 22,040 (IQR: 15,621-30,597) AS neonates. For Guadeloupe, Martinique and Réunion, our estimates were 459 (IQR: 343-594) AS and 14 (IQR: 8-22) SS; 391 (IQR: 300-496) AS and 12 (IQR: 8-19) SS; 6 (IQR: 0-110) AS and 0 (IQR: 0-1) SS neonates respectively. No detailed data from the universal screening programme in place in these French Overseas Departments and Territories could be accessed for comparison.

The inclusion of the complete data from national screening programmes for each maternity or testing centre would obviously result in much better estimates, but collecting these data was beyond the scope of the present study. Moreover, it appeared that existing infrastructures do not allow optimal management and centralisation of data collected locally. Further work on improving the management of these data would therefore be useful, particularly for developing countries aiming at launching a national screening programme.

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Web Tables

Web Table 1: National demographic indicators and estimates of HbS allele frequency and neonates affected. The color code indicates high uncertainty (IQR>1.5*median) in red; medium-high uncertainty (IQR>median) in orange; medium low uncertainty (IQR>0.5*median) in blue; and low uncertainty (IQR<0.5*median) in green. An asterisk (*) indicates Modell and Darlison's 2008 (M&D) estimates falling outside our 50% credible interval (IQR). A double asterisk (**) indicates M&D estimat+C237es falling outside our 90% credible interval. The full legend is shown at the bottom of the Table (p.33).

Country ¹	Population ²	CBR ³	WHO/HbS region ⁴	Surveys	HbS AF (IQR ⁵)	AS neonates/year (IQR5)	SS neonates/year (IQR5)	M&D ⁶
Afghanistan	29,117	0.0423	EMRO/Arab-India	1	0.000 (0.000-0.001)	144 (13-1,078)	0 (0-8)	0
Albania	3,158	0.0128	EURO/Eurasia	1	0.014 (0.009-0.022)	1,391 (863-2,112)	30 (14-58)	11 *
Algeria	35,423	0.0192	AFRO/Eurasia	18	0.007 (0.005-0.010)	10,486 (7,351-15,100)	191 (115-332)	50 **
Angola	18,994	0.0399	AFRO/Sub-Saharan Africa	18	0.137 (0.097-0.192)	112,443 (90,764-135,659)	8,364 (5,880-11,735)	9,560
Argentina	40,668	0.0168	AMRO/Americas	0	0.000 (0.000-0.001)	444 (131-1,273)	4 (1-22)	0 **
Armenia	3,093	0.0150	EURO/Eurasia	0	0.000 (0.000-0.001)	0 (0-8)	0 (0-0)	0
Aruba	107	0.0111	AMRO/Americas	1	0.016 (0.010-0.024)	48 (30-72)	1 (0-2)	0 **
Australia	21,510	0.0135	WPRO/Southeast Asia	2	0.001 (0.001-0.002)	663 (281-1,415)	5 (1-18)	0 *
Austria	8,403	0.0086	EURO/Eurasia	0	0.002 (0.000-0.005)	143 (19-691)	1 (0-10)	0 *
Azerbaijan	8,932	0.0195	EURO/Eurasia	0	0.000 (0.000-0.001)	4 (0-92)	0 (0-0)	4 *
Bahamas	346	0.0152	AMRO/Americas	1	0.024 (0.011-0.054)	287 (130-559)	8 (3-24)	34 *
Bahrain	807	0.0179	EMRO/Arab-India	0	0.050 (0.031-0.076)	1,548 (1,104-2,016)	59 (32-98)	81
Bangladesh	164,424	0.0195	SEAR/Arab-India	0	0.001 (0.001-0.002)	3,459 (847-11,777)	16 (2-101)	0 **
Barbados	257	0.0109	AMRO/Americas	0	0.025 (0.010-0.063)	179 (66-350)	4 (1-15)	4
Belarus	9,579	0.0112	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-7)	0 (0-0)	0
Belgium	10,715	0.0114	EURO/Eurasia	18	0.008 (0.006-0.011)	2,302 (1,695-3,094)	29 (17-49)	5 **
Belize	313	0.0240	AMRO/Americas	3	0.075 (0.051-0.112)	937 (719-1,178)	47 (31-70)	5 **
Benin	9,219	0.0383	AFRO/Sub-Saharan Africa	9	0.159 (0.130-0.191)	64,207 (58,876-69,623)	4,543 (3,808-5,362)	3,561 *
Bhutan	723	0.0196	SEAR/Southeast Asia	0	0.001 (0.000-0.002)	4 (0-26)	0 (0-0)	0
Bolivia (Plurinational State of)	9,995	0.0258	AMRO/Americas	0	0.000 (0.000-0.000)	6 (0-45)	0 (0-0)	0
Bosnia and Herzegovina	3,746	0.0082	EURO/Eurasia	0	0.003 (0.001-0.007)	120 (18-509)	1 (0-8)	0 *
Botswana	1,977	0.0229	AFRO/Sub-Saharan Africa	4	0.003 (0.002-0.007)	300 (120-693)	4 (1-15)	0 **
Brazil	195,453	0.0150	AMRO/Americas	47	0.017 (0.013-0.022)	101,138 (81,605-124,941)	3,244 (2,312-4,570)	764 **
Brunei Darussalam	410	0.0182	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Bulgaria	7,491	0.0100	EURO/Eurasia	0	0.007 (0.003-0.015)	1,076 (377-2,510)	17 (4-64)	0 **
Burkina Faso	16,250	0.0424	AFRO/Sub-Saharan Africa	51	0.056 (0.049-0.064)	71,083 (65,562-76,864)	3,124 (2,681-3,609)	1,065 **
Burundi	8,519	0.0333	AFRO/Sub-Saharan Africa	2	0.040 (0.029-0.057)	24,365 (19,070-30,317)	889 (577-1,322)	2,449 **
Cambodia	15,056	0.0216	WPRO/Southeast Asia	1	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Cameroon	19,957	0.0349	AFRO/Sub-Saharan Africa	34	0.120 (0.105-0.138)	109,857 (102,502-117,187)	6,915 (6,069-7,853)	5,906 *
Canada	33,871	0.0113	AMRO/Americas	3	0.004 (0.002-0.006)	3,004 (1,678-5,163)	45 (17-104)	43
Cape Verde	513	0.0199	AFRO/Sub-Saharan Africa	3	0.026 (0.018-0.036)	625 (463-800)	18 (11-28)	6 *
Central African Republic	4,506	0.0345	AFRO/Sub-Saharan Africa	2	0.077 (0.052-0.113)	18,315 (14,060-22,742)	976 (638-1,437)	677
Chad	11,509	0.0434	AFRO/Sub-Saharan Africa	6	0.051 (0.038-0.069)	43,553 (35,604-52,360)	2,045 (1,459-2,788)	908 *
Chile	17,135	0.0140	AMRO/Americas	4	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
China	1,381,710	0.0119	WPRO/Southeast Asia	8	0.000 (0.000-0.000)	43 (3-384)	0 (0-1)	0
Colombia	46,305	0.0189	AMRO/Americas	2	0.008 (0.004-0.014)	14,464 (7,675-25,504)	321 (126-742)	537
Comoros	691	0.0357	AFRO/Sub-Saharan Africa	2	0.018 (0.011-0.031)	1,105 (660-1,736)	25 (10-56)	0 **
Congo	3,760	0.0346	AFRO/Sub-Saharan Africa	4	0.145 (0.112-0.190)	22,773 (19,572-25,999)	1,560 (1,218-2,008)	2,042 *

Country ¹	Population ²	CBR ³	WHO/Hbs region ⁴	Surveys	Hbs AF (IQR ⁵)	AS neonates/year (IQR5)	SS neonates/year (IQR5)	M&D ⁶
Congo, the Democratic Republic of the	67,829	0.0421	AFRO/Sub-Saharan Africa	11	0.165 (0.142-0.191)	489,745 (455,733-524,014)	38,217 (32,838-44,870)	20,113 **
Costa Rica	4,640	0.0153	AMRO/Americas	1	0.016 (0.008-0.033)	2,731 (1,211-5,281)	60 (18-173)	8 *
Côte d'Ivoire	21,571	0.0330	AFRO/Sub-Saharan Africa	7	0.062 (0.047-0.082)	74,168 (61,560-88,026)	3,567 (2,648-4,773)	1,564 **
Croatia	4,410	0.0098	EURO/Eurasia	0	0.002 (0.001-0.006)	137 (22-572)	1 (0-10)	0 *
Cuba	11,204	0.0096	AMRO/Americas	1	0.030 (0.017-0.054)	6,578 (4,194-9,570)	240 (111-465)	259
Cyprus	846	0.0115	EURO/Eurasia	1	0.007 (0.004-0.013)	147 (61-308)	2 (0-6)	0 *
Czech Republic	10,410	0.0109	EURO/Eurasia	0	0.001 (0.000-0.003)	59 (4-512)	0 (0-5)	0
Denmark	5,482	0.0113	EURO/Eurasia	0	0.000 (0.000-0.002)	11 (0-136)	0 (0-1)	2 *
Djibouti	879	0.0284	EMRO/Sub-Saharan Africa	0	0.001 (0.000-0.001)	3 (0-29)	0 (0-0)	0
Dominican Republic	10,225	0.0209	AMRO/Americas	0	0.019 (0.009-0.040)	9,461 (4,334-17,804)	247 (78-672)	722 *
Ecuador	13,775	0.0198	AMRO/Americas	0	0.000 (0.000-0.001)	45 (3-389)	0 (0-3)	12 *
Egypt	84,472	0.0223	EMRO/Eurasia	5	0.012 (0.006-0.024)	47,754 (21,874-92,324)	1,196 (395-3,096)	19 **
El Salvador	6,201	0.0200	AMRO/Americas	1	0.014 (0.008-0.022)	4,039 (2,370-6,240)	80 (33-168)	8 *
Equatorial Guinea	693	0.0359	AFRO/Sub-Saharan Africa	2	0.192 (0.144-0.253)	5,059 (4,441-5,668)	385 (297-493)	182 **
Eritrea	5,204	0.0344	AFRO/Sub-Saharan Africa	0	0.003 (0.001-0.006)	767 (265-2,109)	9 (2-34)	0 **
Estonia	1,340	0.0121	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Ethiopia	84,996	0.0300	AFRO/Sub-Saharan Africa	8	0.003 (0.002-0.004)	11,197 (6,431-18,883)	137 (60-294)	1 **
Fiji	854	0.0206	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Finland	5,346	0.0114	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
France	62,618	0.0124	EURO/Eurasia	7	0.013 (0.009-0.018)	22,040 (15,621-30,597)	520 (290-915)	79 **
French Guiana	231	0.0240	AMRO/Americas	6	0.015 (0.011-0.021)	186 (133-251)	5 (3-8)	4
French Polynesia	272	0.0164	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Gabon	1,501	0.0270	AFRO/Sub-Saharan Africa	3	0.280 (0.223-0.342)	9,342 (8,503-10,189)	864 (705-1,059)	736
Gambia	1,751	0.0369	AFRO/Sub-Saharan Africa	29	0.075 (0.065-0.087)	8,657 (7,977-9,355)	418 (359-487)	305 *
Georgia	4,221	0.0115	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-4)	0 (0-0)	0
Germany	82,014	0.0087	EURO/Eurasia	0	0.004 (0.002-0.008)	5,940 (2,480-12,680)	85 (22-267)	23
Ghana	24,339	0.0303	AFRO/Sub-Saharan Africa	52	0.087 (0.079-0.097)	102,229 (96,569-107,899)	5,474 (4,924-6,051)	4,847 *
Greece	11,194	0.0101	EURO/Eurasia	31	0.026 (0.020-0.037)	6,667 (5,142-8,664)	206 (142-322)	1 **
Grenada	104	0.0193	AMRO/Americas	0	0.037 (0.016-0.079)	174 (84-284)	6 (2-14)	5
Guadeloupe	467	0.0132	AMRO/Americas	1	0.031 (0.022-0.044)	459 (343-594)	14 (8-22)	16
Guam	180	0.0176	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Guatemala	14,378	0.0315	AMRO/Americas	0	0.011 (0.006-0.020)	10,285 (5,208-19,137)	252 (105-570)	0 **
Guinea	10,324	0.0376	AFRO/Sub-Saharan Africa	1	0.168 (0.128-0.213)	70,965 (62,302-79,040)	5,232 (4,063-6,564)	4,325
Guinea-Bissau	1,647	0.0374	AFRO/Sub-Saharan Africa	4	0.041 (0.029-0.057)	5,011 (3,938-6,175)	192 (123-284)	0 **
Guyana	761	0.0174	AMRO/Americas	0	0.021 (0.009-0.049)	622 (251-1,247)	17 (5-53)	11
Haiti	10,189	0.0257	AMRO/Americas	8	0.051 (0.038-0.069)	26,801 (21,627-32,317)	1,088 (756-1,531)	622 *
Honduras	7,609	0.0258	AMRO/Americas	0	0.049 (0.027-0.092)	17,783 (11,179-25,925)	772 (368-1,473)	0 **
Hong Kong	7,288	0.0088	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Hungary	9,979	0.0101	EURO/Eurasia	1	0.001 (0.000-0.003)	59 (4-406)	0 (0-4)	0
Iceland	329	0.0147	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
India	1,196,940	0.0213	SEAR/Arab-India	113	0.023 (0.020-0.027)	996,563 (883,210-1,135,290)	42,016 (35,347-50,919)	25,768 **
Indonesia	232,546	0.0174	SEAR/Southeast Asia	14	0.000 (0.000-0.000)	328 (79-1,206)	1 (0-7)	0 *
Iran, Islamic Republic of	75,084	0.0162	EMRO/Arab-India	0	0.011 (0.007-0.018)	25,768 (16,613-39,359)	862 (444-1,581)	96 **

Country ¹	Population ²	CBR ³	WHO/Hbs region ⁴	Surveys	Hbs AF (IQR ⁵)	AS neonates/year (IQR5)	SS neonates/year (IQR5)	M&D ⁶
Iraq	31,443	0.0341	EMRO/Arab-India	1	0.010 (0.005-0.022)	23,488 (11,617-47,957)	629 (255-1,595)	499
Ireland	4,588	0.0156	EURO/Eurasia	0	0.001 (0.000-0.003)	60 (5-447)	0 (0-4)	0
Italy	60,119	0.0091	EURO/Eurasia	41	0.005 (0.003-0.009)	5,635 (3,028-10,363)	91 (37-236)	3 **
Jamaica	2,730	0.0179	AMRO/Americas	4	0.037 (0.029-0.048)	3,998 (3,282-4,820)	137 (99-189)	172
Japan	126,995	0.0085	WPRO/Southeast Asia	1	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Jordan	6,461	0.0237	EMRO/Eurasia	2	0.017 (0.012-0.026)	6,365 (4,245-9,313)	145 (80-260)	4 **
Kazakhstan	15,758	0.0206	EURO/Eurasia	0	0.000 (0.000-0.000)	1 (0-11)	0 (0-0)	0
Kenya	40,835	0.0369	AFRO/Sub-Saharan Africa	51	0.038 (0.034-0.043)	98,471 (89,175-108,467)	4,475 (3,879-5,137)	7,872 **
Korea, Democratic People's Republic of	23,963	0.0142	SEAR/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Korea, Republic of	48,517	0.0099	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Kuwait	3,050	0.0174	EMRO/Arab-India	0	0.031 (0.014-0.065)	3,824 (1,878-6,476)	116 (36-293)	37
Kyrgyzstan	5,545	0.0240	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-1)	0 (0-0)	0
Lao People's Democratic Republic	6,434	0.0218	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Latvia	2,241	0.0110	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Lebanon	4,237	0.0149	EMRO/Eurasia	1	0.049 (0.033-0.071)	6,382 (4,808-8,044)	251 (152-382)	6 **
Lesotho	2,064	0.0271	AFRO/Sub-Saharan Africa	0	0.001 (0.000-0.003)	40 (4-244)	0 (0-2)	0
Liberia	4,102	0.0376	AFRO/Sub-Saharan Africa	15	0.046 (0.038-0.057)	13,217 (11,487-14,919)	562 (443-714)	105 **
Libyan Arab Jamahiriya	6,546	0.0217	EMRO/Eurasia	4	0.021 (0.015-0.027)	6,020 (4,673-7,559)	188 (128-267)	34 **
Lithuania	3,256	0.0108	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-1)	0 (0-0)	0
Luxembourg	493	0.0118	EURO/Eurasia	0	0.005 (0.003-0.008)	48 (20-105)	0 (0-1)	0
Madagascar	20,146	0.0346	AFRO/Sub-Saharan Africa	37	0.061 (0.053-0.070)	69,295 (63,138-75,644)	3,379 (2,902-3,915)	2,084 **
Malawi	15,690	0.0445	AFRO/Sub-Saharan Africa	15	0.033 (0.023-0.050)	47,370 (35,308-61,509)	1,688 (1,029-2,765)	917 *
Malaysia	27,949	0.0198	WPRO/Southeast Asia	6	0.000 (0.000-0.000)	1 (0-14)	0 (0-0)	0
Maldives	314	0.0165	SEAR/Eurasia	0	0.000 (0.000-0.000)	0 (0-1)	0 (0-0)	0 *
Mali	13,362	0.0450	AFRO/Sub-Saharan Africa	8	0.057 (0.044-0.072)	56,236 (47,355-65,909)	2,701 (2,068-3,509)	4,421 *
Malta	410	0.0091	EURO/Eurasia	0	0.015 (0.007-0.031)	143 (61-284)	2 (1-8)	0 **
Martinique	406	0.0116	AMRO/Americas	1	0.035 (0.025-0.049)	391 (300-496)	12 (8-19)	9
Mauritania	3,359	0.0327	AFRO/Sub-Saharan Africa	1	0.050 (0.030-0.085)	9,505 (6,585-12,929)	433 (240-748)	159 *
Mauritius	1,297	0.0125	AFRO/Sub-Saharan Africa	0	0.000 (0.000-0.001)	0 (0-19)	0 (0-0)	1 *
Mayotte	199	0.0326	AFRO/Sub-Saharan Africa	0	0.027 (0.010-0.070)	447 (168-863)	12 (2-40)	0 *
Mexico	110,567	0.0185	AMRO/Americas	34	0.007 (0.005-0.009)	31,120 (23,289-41,145)	641 (422-975)	105 **
Micronesia, Federated States of	112	0.0242	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Moldova, Republic of	3,538	0.0121	*/Eurasia	0	0.000 (0.000-0.001)	1 (0-22)	0 (0-0)	0
Mongolia	2,706	0.0227	WPRO/Eurasia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Montenegro	654	0.0120	EURO/Eurasia	0	0.005 (0.002-0.012)	72 (16-226)	1 (0-4)	0 *
Morocco	31,943	0.0187	EMRO/Eurasia	0	0.003 (0.001-0.008)	3,202 (824-10,051)	40 (6-215)	118
Mozambique	23,418	0.0363	AFRO/Sub-Saharan Africa	14	0.027 (0.021-0.035)	42,588 (34,837-51,562)	1,645 (1,193-2,235)	170 **
Myanmar	50,503	0.0167	SEAR/Southeast Asia	6	0.000 (0.000-0.000)	40 (5-222)	0 (0-1)	0
Namibia	2,212	0.0252	AFRO/Sub-Saharan Africa	8	0.010 (0.006-0.015)	1,138 (730-1,806)	30 (17-55)	10 *
Nepal	29,950	0.0232	SEAR/Arab-India	1	0.002 (0.001-0.004)	1,558 (357-4,805)	11 (1-67)	0 **
Netherlands	16,649	0.0108	EURO/Eurasia	2	0.011 (0.008-0.016)	4,759 (3,163-6,819)	82 (43-149)	10 **
Netherlands Antilles	201	0.0131	AMRO/Americas	2	0.023 (0.016-0.032)	150 (105-203)	4 (2-6)	9 *
New Caledonia	254	0.0159	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
New Zealand	4,303	0.0143	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	9 **

Web Table 1

Country ¹	Population ²	CBR ³	WHO/Hbs region ⁴	Surveys	Hbs AF (IQR ⁵)	AS neonates/year (IQR5)	SS neonates/year (IQR5)	M&D ⁶
Nicaragua	5,822	0.0228	AMRO/Americas	0	0.023 (0.009-0.054)	6,514 (2,641-12,964)	204 (53-576)	4 *
Niger	15,885	0.0477	AFRO/Sub-Saharan Africa	7	0.080 (0.058-0.111)	90,671 (73,050-109,417)	4,965 (3,543-6,894)	6,046
Nigeria	158,255	0.0393	AFRO/Sub-Saharan Africa	24	0.171 (0.153-0.192)	1,136,909 (1,070,320-1,200,907)	85,186 (76,413-94,945)	77,192
Norway	4,856	0.0123	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-13)	0 (0-0)	2 *
Oman	2,962	0.0169	EMRO/Arab-India	2	0.001 (0.000-0.001)	32 (9-94)	0 (0-1)	222 **
Pakistan	184,754	0.0263	EMRO/Arab-India	1	0.002 (0.001-0.005)	18,962 (7,260-47,537)	228 (53-892)	101
Panama	3,508	0.0191	AMRO/Americas	3	0.030 (0.019-0.046)	4,457 (3,053-6,176)	140 (77-244)	8 **
Papua New Guinea	6,886	0.0290	WPRO/Southeast Asia	1	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Paraguay	6,462	0.0237	AMRO/Americas	0	0.003 (0.001-0.006)	664 (178-2,093)	7 (1-36)	0 **
Peru	29,493	0.0196	AMRO/Americas	10	0.000 (0.000-0.000)	2 (0-22)	0 (0-0)	25 **
Philippines	93,617	0.0245	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Poland	38,049	0.0109	EURO/Eurasia	0	0.000 (0.000-0.001)	59 (3-590)	0 (0-4)	0
Portugal	10,229	0.0088	EURO/Eurasia	2	0.010 (0.007-0.017)	2,170 (1,303-3,598)	41 (19-90)	3 **
Puerto Rico	3,998	0.0129	AMRO/Americas	1	0.004 (0.002-0.006)	326 (127-710)	2 (1-8)	23 *
Qatar	1,508	0.0116	EMRO/Arab-India	0	0.032 (0.015-0.068)	1,365 (668-2,262)	41 (12-104)	7 *
Réunion	837	0.0169	*/	0	0.001 (0.000-0.004)	6 (0-110)	0 (0-1)	1 *
Romania	21,209	0.0103	EURO/Eurasia	0	0.001 (0.000-0.004)	320 (45-1,500)	2 (0-21)	0 *
Russian Federation	140,324	0.0118	EURO/Eurasia	0	0.000 (0.000-0.000)	14 (1-135)	0 (0-0)	0
Rwanda	10,277	0.0406	AFRO/Sub-Saharan Africa	2	0.023 (0.018-0.029)	22,857 (18,400-28,098)	601 (423-846)	125 **
Saint Lucia	174	0.0169	AMRO/Americas	1	0.049 (0.031-0.074)	311 (223-402)	12 (6-19)	7
Saint Vincent and the Grenadines	109	0.0164	AMRO/Americas	0	0.045 (0.022-0.086)	176 (99-262)	6 (2-13)	2
Samoa	530	0.0237	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Saudi Arabia	26,207	0.0214	EMRO/Arab-India	28	0.018 (0.016-0.021)	19,447 (17,012-22,221)	691 (572-842)	5,397 **
Sao Tome and Principe	165	0.0299	AFRO/Sub-Saharan Africa	0	0.094 (0.033-0.236)	763 (376-1,132)	41 (12-92)	49
Senegal	12,866	0.0359	AFRO/Sub-Saharan Africa	20	0.067 (0.054-0.082)	53,002 (46,123-60,123)	2,535 (1,998-3,163)	1,895 *
Serbia	9,842	0.0110	EURO/Eurasia	0	0.004 (0.002-0.010)	830 (225-2,417)	10 (1-49)	0 **
Sierra Leone	5,837	0.0365	AFRO/Sub-Saharan Africa	0	0.164 (0.108-0.240)	39,530 (31,971-47,084)	2,838 (1,906-4,018)	3,243
Singapore	4,777	0.0095	WPRO/Southeast Asia	1	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Slovakia	5,406	0.0107	EURO/Eurasia	0	0.001 (0.000-0.002)	12 (0-125)	0 (0-1)	0
Slovenia	2,027	0.0099	EURO/Eurasia	0	0.001 (0.000-0.005)	22 (2-176)	0 (0-2)	0
Solomon Islands	536	0.0304	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Somalia	9,359	0.0430	EMRO/Sub-Saharan Africa	2	0.002 (0.001-0.005)	1,192 (321-3,777)	13 (2-69)	0 *
South Africa	50,523	0.0205	AFRO/Sub-Saharan Africa	3	0.003 (0.002-0.004)	4,946 (2,617-8,872)	73 (28-177)	0 **
Spain	45,322	0.0106	EURO/Eurasia	4	0.007 (0.004-0.012)	7,324 (4,253-12,071)	149 (64-341)	0 **
Sri Lanka	20,410	0.0170	SEAR/Arab-India	28	0.001 (0.001-0.002)	425 (295-590)	2 (1-3)	0 **
Sudan	43,182	0.0319	AFRO/Sub-Saharan Africa	32	0.043 (0.034-0.055)	98,244 (84,697-113,554)	4,567 (3,527-5,894)	483 **
Suriname	524	0.0178	AMRO/Americas	0	0.030 (0.013-0.066)	653 (300-1,162)	19 (6-53)	13
Swaziland	1,195	0.0287	AFRO/Sub-Saharan Africa	0	0.006 (0.003-0.010)	398 (173-813)	4 (1-13)	1
Sweden	9,292	0.0120	EURO/Eurasia	0	0.000 (0.000-0.001)	2 (0-41)	0 (0-0)	4 *
Switzerland	7,597	0.0100	EURO/Eurasia	0	0.004 (0.002-0.011)	622 (145-1,907)	6 (1-36)	0 *
Syrian Arab Republic	22,627	0.0219	EMRO/Eurasia	4	0.025 (0.019-0.033)	24,448 (19,056-31,267)	901 (664-1,261)	66 **
Tajikistan	7,078	0.0277	EURO/Eurasia	0	0.000 (0.000-0.000)	0 (0-6)	0 (0-0)	0
Tanzania, United Republic of	45,028	0.0410	AFRO/Sub-Saharan Africa	57	0.074 (0.063-0.087)	203,747 (185,237-222,866)	11,022 (9,327-12,897)	8,655 *
Thailand	68,141	0.0115	SEAR/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-10)	0 (0-0)	0

Country ¹	Population ²	CBR ³	WHO/Hbs region ⁴	Surveys	Hbs AF (IQR ⁵)	AS neonates/year (IQR5)	SS neonates/year (IQR5)	M&D ⁶
Timor-Leste	1,167	0.0376	SEAR/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Togo	6,774	0.0310	AFRO/Sub-Saharan Africa	4	0.125 (0.105-0.149)	34,057 (31,180-36,958)	2,175 (1,846-2,540)	1,746 *
Trinidad and Tobago	1,344	0.0143	AMRO/Americas	1	0.038 (0.026-0.056)	1,648 (1,207-2,154)	56 (33-91)	30 *
Tonga	104	0.0257	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Tunisia	10,374	0.0165	EMRO/Eurasia	12	0.011 (0.009-0.013)	4,420 (3,623-5,372)	81 (58-113)	63
Turkey	75,699	0.0169	EURO/Eurasia	45	0.005 (0.004-0.007)	12,852 (9,281-20,051)	330 (248-526)	116 **
Turkmenistan	5,178	0.0212	EURO/Eurasia	0	0.000 (0.000-0.001)	4 (0-54)	0 (0-0)	0
Uganda	33,798	0.0439	AFRO/Sub-Saharan Africa	11	0.082 (0.067-0.100)	189,801 (169,157-210,719)	10,143 (8,223-12,325)	8,216 *
Ukraine	45,454	0.0109	EURO/Eurasia	0	0.000 (0.000-0.001)	33 (2-293)	0 (0-2)	0
United Arab Emirates	4,647	0.0123	EMRO/Arab-India	2	0.004 (0.002-0.007)	402 (189-827)	4 (1-11)	24 *
United Kingdom	61,900	0.0121	EURO/Eurasia	4	0.009 (0.007-0.013)	15,746 (10,982-21,949)	296 (163-517)	108 *
United States of America	317,736	0.0137	AMRO/Americas	25	0.011 (0.009-0.014)	100,150 (82,111-122,520)	2,843 (2,038-3,968)	838 **
Uruguay	3,373	0.0145	AMRO/Americas	0	0.001 (0.001-0.003)	82 (26-231)	1 (0-3)	0 **
Uzbekistan	27,790	0.0208	EURO/Eurasia	0	0.000 (0.000-0.000)	1 (0-24)	0 (0-0)	0
Vanuatu	246	0.0287	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-0)	0 (0-0)	0
Venezuela, Bolivarian Republic of	29,044	0.0198	AMRO/Americas	40	0.014 (0.012-0.017)	19,272 (16,150-22,799)	435 (325-591)	108 **
Viet Nam	89,009	0.0159	WPRO/Southeast Asia	0	0.000 (0.000-0.000)	0 (0-4)	0 (0-0)	0
Virgin Islands, U.S.	109	0.0121	*/	0	0.008 (0.004-0.018)	25 (8-59)	0 (0-1)	4 *
Western Sahara	441	0.0213	*/	0	0.007 (0.003-0.017)	144 (51-330)	3 (1-10)	2
Yemen	24,325	0.0372	EMRO/Sub-Saharan Africa	1	0.004 (0.002-0.006)	6,411 (3,355-12,043)	80 (29-207)	119
Zambia	13,254	0.0465	AFRO/Sub-Saharan Africa	11	0.112 (0.095-0.133)	88,801 (80,963-96,982)	5,652 (4,801-6,632)	3,346 **
Zimbabwe	12,645	0.0287	AFRO/Sub-Saharan Africa	7	0.021 (0.016-0.028)	16,345 (12,706-20,982)	483 (326-704)	162 **

1. Official country names consistent with ISO 3166-1. The following countries are not shown: Åland Islands; Aksai Chin; American Samoa; Andorra; Anguilla; Antarctica; Antigua and Barbuda; Arunachal Pradesh; Ashmore and Cartier Islands; Azores Islands; Baker Island; Bassas da India; Bermuda; Bird Island; Bonaire; Sint Eustatius and Saba; Bouvet Island; British Indian Ocean Territory; British Virgin Islands; Cayman Islands; China/India; Christmas Island; Clipperton Island; Cocos (Keeling) Islands; Cook Islands; Curaçao; Dhekelia and Akrotiri SBA; Dominica; Europa Island; Falkland Islands (Malvinas); Faroe Islands; French Southern Territories; Gaza Strip; Gibraltar; Glorioso Island; Greenland; Guernsey; Hala'ib triangle; Heard Island and McDonald Islands; Holy See (Vatican City State); Howland Island; Ilemi triangle; Isle of Man; Jammu Kashmir; Jarvis Island; Jersey; Johnston Atoll; Juan de Nova Island; Kashmir; Kingman Reef; Kiribati; Kuril islands; Liancourt Rock; Liechtenstein; Madeira Islands; Marshall Islands; Ma'tan al-Sarra; Midway Island; Monaco; Montserrat; Nauru; Navassa Island; Niue; Norfolk Island; Northern Mariana Islands; Palau; Palestinian Territory, Occupied; Palmyra Atoll; Paracel Islands; Pitcairn; Saint Bathélemy; Saint Helena, Ascension and Tristan de Cunha; Saint Kitts and Nevis; Saint Martin (French Part); Saint Pierre and Miquelon; San Marino; Scarborough Reef; Senkaku Islands; Seychelles; Sint Maarten (Dutch part); South Georgia and the South Sandwich Islands; Svalbard and Jan Mayen; Taiwan, Province of China; Tokelau; Trinidad and Tobago; Tromelin Island; Turks and Caicos islands; Tuvalu; United States Minor Outlying Islands; Virgin Islands, British; Wake Island; Wallis and Futuna; West Bank.

2. In thousands. Population data were obtained for the year 2000 from the Global Rural Urban Mapping Project (GRUMP) alpha version and projected to 2010 by applying United Nations national, medium variant, inter-censal growth rates by country [24].

3. CBR = Crude birth rate. Source: Population Division of the Department of Economic and Social Affairs of the United Nations Secretariat, World Population Prospects: The 2010 Revision, <http://esa.un.org/unpd/wpp/index.htm>, Monday, July 25, 2011.

4. World Health Organisation Regional Offices: AFRO, African; SEARO, South East Asian; WPRO, Western Pacific; EMRO, Eastern Mediterranean; AMRO, American; EURO, European.

5. IQR: interquartile range of the posterior predictive distribution for each population at risk estimate. The color code indicates high uncertainty (IQR>1.5*median) in red; medium-high uncertainty (IQR>median) in orange; medium low uncertainty (IQR>0.5*median) in blue; and low uncertainty (IQR<0.5*median) in green.

6. SS neonate estimates from Modell B, Darlison M (2008) Global epidemiology of haemoglobin disorders and derived service indicators. Bull World Health Organ 86: 480-487 [7]. C54

Web Table 2: National areal prediction summaries and Monte Carlo standard errors (SE) for AS and SS neonate estimates

COUNTRY ¹	HbS heterozygote neonates (AS)								HbS homozygote neonates (SS)							
	Mean	SE	Median	SE	Q25%	SE	Q75%	SE	Mean	SE	Median	SE	Q25%	SE	Q75%	SE
Afghanistan	1,889	37	144	8	13	1	1,078	39	40	2	0	0	0	0	8	0
Albania	1,619	6	1,391	8	863	9	2,112	12	44	0	30	0	14	0	58	1
Algeria	11,939	84	10,486	88	7,351	94	15,100	113	267	3	191	3	115	2	332	5
Angola	114,473	451	112,443	369	90,764	554	135,659	578	9,761	73	8,364	51	5,880	49	11,735	76
Argentina	1,436	32	444	20	131	7	1,273	38	38	2	4	0	1	0	22	1
Armenia	60	4	0	0	0	0	8	1	1	0	0	0	0	0	0	0
Aruba	54	0	48	0	30	0	72	0	1	0	1	0	0	0	2	0
Australia	1,101	20	663	14	281	9	1,415	27	17	0	5	0	1	0		0
Austria	727	11	143	3	19	1	691	18	23	1	1	0	0	0	10	0
Azerbaijan	398	13	4	0	0	0	92	8	10	1	0	0	0	0	0	0
Bahamas	376	3	287	4	130	2	559	6	19	0	8	0	3	0	24	0
Bahrain	2,468	8	1,548	13	1,104	16	2,016	9	71	0	59	1	32	1	98	1
Bangladesh	10,409	234	3,459	113	847	32	11,777	323	195	8	16	1	2	0	101	4
Barbados	243	2	179	2	66	1	350	5	12	0	4	0	1	0	15	0
Belarus	94	5	0	0	0	0	7	0	3	0	0	0	0	0	0	0
Belgium	2,499	9	2,302	8	1,695	8	3,094	17	38	0	29	0	17	0	49	1
Belize	950	2	937	3	719	3	1,178	3	54	0	47	0	31	0	70	0
Benin	61,586	149	64,207	138	58,876	181	69,623	145	4,637	17	4,543	21	3,808	22	5,362	19
Bhutan	33	1	4	0	0	0	26	1	0	0	0	0	0	0	0	0
Bolivia (Plurinational State of)	136	5	6	0	0	0	45	2	3	0	0	0	0	0	0	0
Bosnia and Herzegovina	487	12	120	5	18	1	509	16	16	1	1	0	0	0	8	0
Botswana	557	6	300	5	120	3	693	11	16	0	4	0	1	0	15	0
Brazil	105,051	455	101,138	532	81,605	771	124,941	488	3,697	25	3,244	28	2,312	28	4,570	38
Brunei Darussalam	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Bulgaria	1,840	30	1,076	29	377	12	2,510	38	61	1	17	1	4	0	64	1
Burkina Faso	72,279	126	71,083	100	65,562	96	76,864	214	3,179	9	3,124	8	2,681	8	3,609	15
Burundi	24,599	52	24,365	56	19,070	65	30,317	108	1,017	5	889	2	577	4	1,322	8
Cambodia	21	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Cameroon	107,877	219	109,857	203	102,502	238	117,187	268	7,033	24	6,915	24	6,069	24	7,853	37
Canada	3,971	50	3,004	50	1,678	32	5,163	68	90	2	45	1	17	0	104	2
Cape Verde	622	2	625	2	463	2	800	3	21	0	18	0	11	0	28	0
Central African Republic	18,179	67	18,315	79	14,060	97	22,742	59	1,126	7	976	8	638	7	1,437	10
Chad	43,331	156	43,553	189	35,604	150	52,360	191	2,224	14	2,045	14	1,459	9	2,788	18
Chile	4	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
China	941	48	43	6	3	1	384	43	19	2	0	0	0	0	1	0
Colombia	18,814	58	14,464	175	7,675	109	25,504	207	591	3	321	5	126	3	742	8
Comoros	1,364	7	1,105	12	660	8	1,736	9	44	0	25	0	10	0	56	0

COUNTRY¹	Mean	SE	Median	SE	Q25%	SE	Q75%	SE	Mean	SE	Median	SE	Q25%	SE	Q75%	SE
Congo	24,627	38	22,773	58	19,572	60	25,999	77	1,729	10	1,560	6	1,218	4	2,008	11
Costa Rica	3,685	22	2,731	27	1,211	32	5,281	46	141	2	60	1	18	1	173	3
Côte d'Ivoire	69,104	194	74,168	243	61,560	194	88,026	221	3,890	20	3,567	16	2,648	19	4,773	27
Croatia	542	7	137	3	22	1	572	12	18	0	1	0	0	0	10	0
Cuba	7,187	35	6,578	29	4,194	40	9,570	56	345	3	240	2	111	2	465	3
Cyprus	296	2	147	3	61	1	308	5	5	0	2	0	0	0	6	0
Czech Republic	748	18	59	3	4	0	512	22	22	1	0	0	0	0	5	0
Democratic Republic of the Congo	477,016	1,714	489,745	2,002	455,733	1,504	524,014	2,012	39,965	271	38,217	286	32,838	235	44,870	371
Denmark	318	8	11	0	0	0	136	6	10	0	0	0	0	0	1	0
Djibouti	62	2	3	0	0	0	29	1	1	0	0	0	0	0	0	0
Dominican Republic	12,092	71	9,461	79	4,334	47	17,804	121	540	7	247	3	78	1	672	10
Ecuador	877	28	45	1	3	0	389	16	23	1	0	0	0	0	3	0
Egypt	64,466	478	47,754	751	21,874	287	92,324	703	2,570	34	1,196	31	395	9	3,096	41
El Salvador	4,668	20	4,039	28	2,370	17	6,240	60	129	1	80	1	33	1	168	2
Equatorial Guinea	5,099	7	5,059	13	4,441	8	5,668	9	409	1	385	2	297	1	493	2
Eritrea	1,799	34	767	19	265	5	2,109	46	42	1	9	0	2	0	34	1
Estonia	6	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Ethiopia	14,263	106	11,197	113	6,431	71	18,883	148	259	3	137	2	60	1	294	3
Fiji	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Finland	18	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0
France	24,435	104	22,040	175	15,621	124	30,597	208	727	5	520	5	290	3	915	11
French Guiana	201	2	186	2	133	2	251	2	6	0	5	0	3	0	8	0
French Polynesia	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Gabon	9,374	16	9,342	16	8,503	18	10,189	21	905	2	864	2	705	2	1,059	4
Georgia	38	2	0	0	0	0	4	0	1	0	0	0	0	0	0	0
Germany	9,880	143	5,940	88	2,480	58	12,680	156	265	6	85	2	22	1	267	5
Ghana	102,545	248	102,229	264	96,569	261	107,899	250	5,519	19	5,474	21	4,924	20	6,051	23
Greece	7,187	40	6,667	51	5,142	47	8,664	46	259	2	206	2	142	2	322	3
Grenada	196	1	174	2	84	2	284	2	10	0	6	0	2	0	14	0
Guadeloupe	469	1	459	2	343	1	594	2	16	0	14	0	8	0	22	0
Guam	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Guatemala	14,159	184	10,285	172	5,208	96	19,137	258	470	9	252	6	105	3	570	14
Guinea	68,464	115	70,965	124	62,302	156	79,040	176	5,518	19	5,232	24	4,063	16	6,564	29
Guinea-Bissau	4,708	19	5,011	23	3,938	11	6,175	32	218	2	192	2	123	1	284	3
Guyana	837	7	622	7	251	5	1,247	18	43	1	17	0	5	0	53	1
Haiti	26,659	38	26,801	75	21,627	72	32,317	57	1,197	3	1,088	5	756	5	1,531	6
Honduras	19,204	100	17,783	182	11,179	171	25,925	104	1,083	8	772	11	368	8	1,473	10
China, Hong Kong SAR	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Hungary	553	16	59	3	4	0	406	9	14	1	0	0	0	0	4	0
Iceland	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

COUNTRY ¹	Mean	SE	Median	SE	Q25%	SE	Q75%	SE	Mean	SE	Median	SE	Q25%	SE	Q75%	SE
India	1,044,394	10,547	996,563	11,472	883,210	10,488	1,135,290	10,399	44,252	671	42,016	706	35,347	616	50,919	770
Indonesia	1,907	140	328	23	79	8	1,206	76	32	6	1	0	0	0	7	1
Iran (Islamic Republic of)	30,617	411	25,768	441	16,613	353	39,359	433	1,252	21	862	18	444	11	1,581	19
Iraq	36,723	386	23,488	420	11,617	176	47,957	761	1,532	27	629	14	255	6	1,595	35
Ireland	526	13	60	2	5	0	447	13	15	1	0	0	0	0	4	0
Israel	7,856	48	7,117	58	5,302	36	9,248	83	252	2	218	3	136	2	324	6
Italy	8,051	67	5,635	48	3,028	45	10,363	107	209	4	91	2	37	1	236	4
Jamaica	4,128	13	3,998	16	3,282	14	4,820	18	152	1	137	1	99	1	189	1
Japan	3	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Jordan	6,850	28	6,365	37	4,245	22	9,313	50	202	1	145	1	80	1	260	3
Kazakhstan	63	2	1	0	0	0	11	1	1	0	0	0	0	0	0	0
Kenya	98,657	956	98,471	1,055	89,175	915	108,467	1,039	4,566	63	4,475	63	3,879	59	5,137	68
Dem. People's Republic of Korea	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Republic of Korea	3	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Kuwait	3,982	30	3,824	34	1,878	39	6,476	71	212	3	116	2	36	1	293	7
Kyrgyzstan	52	3	0	0	0	0	1	0	1	0	0	0	0	0	0	0
Lao People's Democratic Republic	17	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Latvia	15	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Lebanon	6,502	37	6,382	41	4,808	37	8,044	46	290	3	251	3	152	2	382	3
Lesotho	343	8	40	2	4	0	244	13	7	0	0	0	0	0	2	0
Liberia	12,890	54	13,217	60	11,487	50	14,919	59	594	3	562	4	443	3	714	4
Libyan Arab Jamahiriya	6,087	29	6,020	41	4,673	37	7,559	36	210	1	188	2	128	1	267	2
Lithuania	32	2	0	0	0	0	1	0	1	0	0	0	0	0	0	0
Luxembourg	83	1	48	1	20	0	105	1	1	0	0	0	0	0	1	0
China, Macao SAR	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
TFYR Macedonia	736	5	510	6	230	5	1,016	14	22	0	9	0	3	0	25	0
Madagascar	71,583	336	69,295	362	63,138	282	75,644	372	3,457	28	3,379	28	2,902	23	3,915	33
Malawi	47,002	194	47,370	249	35,308	251	61,509	264	2,133	16	1,688	18	1,029	14	2,765	25
Malaysia	56	2	1	0	0	0	14	0	1	0	0	0	0	0	0	0
Maldives	6	0	0	0	0	0	1	0	0	0	0	0	0	0	0	0
Mali	65,849	235	56,236	231	47,355	268	65,909	301	2,904	18	2,701	20	2,068	17	3,509	20
Malta	197	2	143	2	61	1	284	4	7	0	2	0	1	0	8	0
Martinique	403	1	391	1	300	1	496	1	14	0	12	0	8	0	19	0
Mauritania	10,335	47	9,505	72	6,585	50	12,929	109	573	5	433	5	240	3	748	10
Mauritius	105	4	0	0	0	0	19	1	4	0	0	0	0	0	0	0
Mayotte	575	3	447	7	168	3	863	8	30	0	12	0	2	0	40	1
Mexico	34,594	214	31,120	235	23,289	256	41,145	242	800	10	641	9	422	7	975	10
Micronesia (Fed. States of)	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Republic of Moldova	124	5	1	0	0	0	22	1	3	0	0	0	0	0	0	0
Mongolia	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0

COUNTRY¹	Mean	SE	Median	SE	Q25%	SE	Q75%	SE	Mean	SE	Median	SE	Q25%	SE	Q75%	SE
Montenegro	170	3	72	1	16	0	226	3	5	0	1	0	0	0	4	0
Morocco	8,269	92	3,202	54	824	23	10,051	139	278	6	40	1	6	0	215	7
Mozambique	43,902	241	42,588	253	34,837	195	51,562	312	1,818	19	1,645	19	1,193	13	2,235	30
Myanmar	305	12	40	2	5	0	222	7	4	0	0	0	0	0	1	0
Namibia	1,442	11	1,138	8	730	8	1,806	22	46	0	30	1	17	0	55	1
Nepal	4,150	77	1,558	33	357	16	4,805	80	105	4	11	0	1	0	67	2
Netherlands	5,281	19	4,759	19	3,163	22	6,819	41	115	1	82	1	43	0	149	2
Netherlands Antilles	159	0	150	1	105	1	203	1	5	0	4	0	2	0	6	0
New Caledonia	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
New Zealand	3	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Nicaragua	8,715	57	6,514	77	2,641	40	12,964	125	471	7	204	4	53	1	576	10
Niger	89,801	231	90,671	325	73,050	326	109,417	469	5,503	24	4,965	32	3,543	25	6,894	38
Nigeria	1,138,493	1,665	1,136,909	1,640	1,070,320	2,018	1,200,907	1,719	86,420	211	85,186	247	76,413	272	94,945	190
Norway	77	4	0	0	0	0	13	1	2	0	0	0	0	0	0	0
Oman	81	3	32	1	9	0	94	2	2	0	0	0	0	0	1	0
Pakistan	37,554	616	18,962	275	7,260	92	47,537	540	1,123	42	228	6	53	2	892	9
Panama	4,793	16	4,457	23	3,053	16	6,176	30	184	1	140	1	77	1	244	2
Papua New Guinea	4	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Paraguay	1,878	40	664	19	178	4	2,093	57	55	2	7	0	1	0	36	1
Peru	80	5	2	0	0	0	22	1	1	0	0	0	0	0	0	0
Philippines	55	9	0	0	0	0	0	0	1	0	0	0	0	0	0	0
Poland	1,110	26	59	3	3	0	590	12	28	1	0	0	0	0	4	0
Portugal	2,921	21	2,170	14	1,303	13	3,598	34	81	1	41	0	19	0	90	1
Puerto Rico	498	4	326	4	127	1	710	11	8	0	2	0	1	0	8	0
Qatar	1,809	13	1,365	28	668	11	2,262	26	74	1	41	1	12	0	104	2
Réunion	193	4	6	0	0	0	110	7	8	1	0	0	0	0	1	0
Romania	1,543	33	320	7	45	2	1,500	48	44	2	2	0	0	0	21	1
Russian Federation	381	22	14	1	1	0	135	9	9	1	0	0	0	0	0	0
Rwanda	24,532	75	22,857	103	18,400	58	28,098	124	680	4	601	6	423	2	846	7
Saint Lucia	317	1	311	2	223	1	402	1	14	0	12	0	6	0	19	0
Saint Vincent and the Grenadines	187	1	176	1	99	2	262	1	9	0	6	0	2	0	13	0
Samoa	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Sao Tome and Principe	783	6	763	11	376	7	1,132	7	72	1	41	1	12	0	92	1
Saudi Arabia	20,767	187	19,447	190	17,012	181	22,221	213	723	9	691	9	572	8	842	10
Senegal	51,572	142	53,002	154	46,123	146	60,123	205	2,652	15	2,535	16	1,998	14	3,163	22
Serbia	1,891	28	830	20	225	9	2,417	59	59	2	10	0	1	0	49	2
Sierra Leone	39,730	128	39,530	197	31,971	123	47,084	116	3,170	21	2,838	23	1,906	15	4,018	24
Singapore	4	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Slovakia	221	7	12	1	0	0	125	7	5	0	0	0	0	0	1	0
Slovenia	228	5	22	1	2	0	176	5	7	0	0	0	0	0	2	0

COUNTRY¹	Mean	SE	Median	SE	Q25%	SE	Q75%	SE	Mean	SE	Median	SE	Q25%	SE	Q75%	SE
Solomon Islands	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Somalia	3,423	51	1,192	37	321	12	3,777	70	109	2	13	1	2	0	69	2
South Africa	6,803	58	4,946	39	2,617	43	8,872	109	157	3	73	1	28	1	177	3
Spain	9,231	45	7,324	47	4,253	43	12,071	103	281	2	149	2	64	1	341	5
Sri Lanka	479	5	425	6	295	4	590	6	2	0	2	0	1	0	3	0
Sudan	100,612	596	98,244	714	84,697	688	113,554	511	4,921	40	4,567	52	3,527	43	5,894	43
Suriname	783	7	653	10	300	7	1,162	13	39	1	19	0	6	0	53	1
Swaziland	605	4	398	3	173	2	813	7	12	0	4	0	1	0	13	0
Sweden	186	10	2	0	0	0	41	2	5	0	0	0	0	0	0	0
Switzerland	1,510	24	622	20	145	5	1,907	51	46	1	6	0	1	0	36	1
Syrian Arab Republic	23,406	124	24,448	123	19,056	163	31,267	71	1,030	10	901	9	664	11	1,261	8
Tajikistan	109	8	0	0	0	0	6	0	2	0	0	0	0	0	0	0
United Republic of Tanzania	203,814	696	203,747	685	185,237	738	222,866	831	11,304	71	11,022	77	9,327	68	12,897	82
Thailand	108	11	0	0	0	0	10	0	2	0	0	0	0	0	0	0
Gambia	8,557	6	8,657	12	7,977	9	9,355	10	427	1	418	1	359	1	487	1
Timor-Leste	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Togo	30,324	89	34,057	93	31,180	109	36,958	92	2,211	10	2,175	8	1,846	9	2,540	12
Tonga	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Trinidad and Tobago	1,713	5	1,648	9	1,207	6	2,154	9	68	0	56	1	33	0	91	1
Tunisia	4,619	17	4,420	20	3,623	16	5,372	25	90	1	81	1	58	0	113	1
Turkey	16,384	449	12,852	363	9,281	327	20,051	656	519	20	330	17	248	14	526	23
Turkmenistan	173	7	4	0	0	0	54	3	5	0	0	0	0	0	0	0
Uganda	187,866	618	189,801	754	169,157	604	210,719	728	10,454	54	10,143	65	8,223	51	12,325	65
Ukraine	600	16	33	1	2	0	293	11	14	1	0	0	0	0	2	0
United Arab Emirates	1,043	16	402	10	189	7	827	22	11	0	4	0	1	0	11	0
United Kingdom	17,273	41	15,746	79	10,982	52	21,949	114	395	2	296	1	163	2	517	5
United States of America	102,887	512	100,150	600	82,111	505	122,520	631	3,289	25	2,843	20	2,038	16	3,968	34
Uruguay	263	3	82	2	26	1	231	6	7	0	1	0	0	0	3	0
Uzbekistan	255	23	1	0	0	0	24	2	5	1	0	0	0	0	0	0
Vanuatu	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Venezuela (Bolivarian Republic of)	19,721	107	19,272	104	16,150	93	22,799	156	485	4	435	4	325	2	591	6
Viet Nam	190	19	0	0	0	0	4	0	3	1	0	0	0	0	0	0
United States Virgin Islands	42	0	25	0	8	0	59	0	1	0	0	0	0	0	1	0
Western Sahara	295	4	144	3	51	1	330	8	10	0	3	0	1	0	10	0
Yemen	9,318	58	6,411	77	3,355	41	12,043	78	194	2	80	1	29	1	207	2
Zambia	87,955	297	88,801	344	80,963	327	96,982	344	5,806	29	5,652	32	4,801	28	6,632	34
Zimbabwe	17,166	50	16,345	48	12,706	57	20,982	70	560	4	483	4	326	4	704	4

1. Official country names consistent with ISO 3166-1. See Supplementary Table S1 for list of countries not shown.

Web Table 3: Sub-national areal prediction summaries and Monte Carlo standard errors (SE) for HbAS and HbSS neonate estimates

AFRICA						HbAS								HbSS							
Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE
Algeria	Illizi	362	7	47,940	0.0192	60	0	56	1	43	1	74	1	2	0	2	0	1	0	3	0
Algeria	Ouargla	373	2	636,715	0.0192	416	7	387	3	266	2	530	18	10	0	8	0	4	0	14	1
Angola	Cuando Cubango	402	5	220,749	0.0399	249	4	230	2	166	3	313	8	7	0	6	0	3	0	9	0
Angola	Lunda Norte	409	7	530,093	0.0399	4,550	3	4,553	10	4,250	9	4,846	21	388	0	381	1	334	0	434	1
Angola	Lunda Sul	410	2	262,521	0.0399	1,659	10	1,656	23	1,397	13	1,928	17	105	1	98	3	73	1	133	2
Angola	Malanje	411	4	1,453,300	0.0399	14,035	8	14,033	4	12,783	34	15,344	95	1,471	6	1,364	4	1,096	10	1,722	35
Benin	Atakora	610	7	737,962	0.0383	2,989	16	2,969	18	2,553	11	3,413	17	128	2	122	2	92	0	157	2
Botswana	Ngamiland	660	2	139,642	0.0229	17	0	10	1	4	0	22	0	0	0	0	0	0	0	0	0
Burkina Faso	Boucle Du Mouhoun	40276	3	1,795,040	0.0424	5,474	42	5,355	44	4,403	43	6,438	18	192	2	177	2	128	2	242	1
Burkina Faso	Cascades	40277	3	546,281	0.0424	2,768	11	2,757	27	2,189	0	3,312	4	135	1	124	3	81	1	173	2
Burkina Faso	Centre-est	40279	2	1,353,980	0.0424	6,155	9	6,121	26	5,083	30	7,131	3	267	0	251	3	182	0	335	0
Burkina Faso	Centre-sud	40282	8	799,269	0.0424	3,313	9	3,287	21	2,896	8	3,712	12	128	1	123	2	97	1	153	1
Burkina Faso	Hauts-bassins	40284	26	1,663,650	0.0424	5,326	1	5,305	5	4,822	0	5,803	6	167	0	163	0	139	0	190	0
Burkina Faso	Nord	40285	2	1,520,520	0.0424	7,267	67	7,237	41	5,746	101	8,666	76	338	5	311	2	209	4	436	8
Burkina Faso	Sahel	40287	4	1,008,040	0.0424	5,405	49	5,385	44	4,594	51	6,203	33	284	4	271	1	201	3	353	1
Burundi	Bujumbura-Mairie	40543	2	498,769	0.0333	1,440	5	1,395	6	1,089	8	1,743	12	49	0	43	0	27	0	64	1
Cote d'Ivoire	Lagunes	1044	3	5,733,330	0.0330	19,069	48	18,876	95	15,669	64	22,253	75	756	2	704	5	502	1	946	3
Cote d'Ivoire	Zanzan	1053	4	928,182	0.0330	4,976	36	4,996	34	4,416	57	5,574	18	314	4	304	6	242	6	375	1
Cameroon	Centre	816	2	3,394,960	0.0349	23,598	12	23,650	57	21,407	15	25,981	44	1,731	1	1,687	13	1,373	8	2,060	13
Cameroon	Est	817	2	877,748	0.0349	4,618	41	4,604	28	3,321	36	5,818	105	319	7	272	0	151	0	426	16
Cameroon	Extreme-Nord	818	6	3,170,660	0.0349	11,777	142	11,722	121	10,244	155	13,211	142	532	12	512	9	401	8	642	13
Cameroon	Littoral	819	4	3,257,190	0.0349	18,928	63	19,112	121	16,178	86	21,801	89	1,175	7	1,122	11	808	4	1,482	14
Cameroon	Ouest	822	2	2,365,110	0.0349	12,515	53	12,531	49	10,704	68	14,365	11	725	2	691	1	510	7	897	2
Cameroon	Sud	823	2	650,268	0.0349	4,224	1	4,219	6	3,742	3	4,732	12	303	1	290	3	229	2	367	1
Cameroon	Sud-Ouest	824	15	1,590,850	0.0349	10,125	35	10,159	19	9,400	42	10,831	25	682	2	676	2	580	6	774	2
Chad	Mandoul	12925	2	422,971	0.0434	1,466	1	1,453	0	1,084	5	1,815	4	56	0	49	0	30	0	74	1
Comoros	Ngazidja	969	2	360,557	0.0357	518	1	472	2	298	3	682	1	12	0	8	0	4	0	15	0
Congo	Pool	978	3	1,664,670	0.0346	10,867	54	10,920	25	9,907	69	11,894	79	736	4	725	1	598	5	863	12
DRC	Equateur	1068	2	7,264,900	0.0421	52,038	84	51,681	524	43,694	183	60,396	40	4,082	16	3,730	47	2,805	7	5,019	34
DRC	Katanga	1071	2	8,788,270	0.0421	70,148	581	69,811	596	59,978	21	79,793	1,259	6,215	117	5,514	144	4,175	8	7,350	151
DRC	Province Orientale	1075	3	8,980,980	0.0421	56,452	277	56,200	157	48,710	357	63,712	266	3,922	27	3,711	9	2,945	15	4,687	43

Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE
Egypt	Al Wadi/al Jadid	1169	2	1,147,300	0.0223	1,427	29	1,171	12	680	20	1,969	22	62	2	40	1	18	1	82	1
Ethiopia	Addis Ababa	1227	2	3,923,150	0.0300	489	5	282	5	99	1	670	5	3	0	1	0	0	0	3	0
Ethiopia	Amhara	1229	3	21,959,100	0.0300	3,153	5	1,917	26	829	6	4,052	18	46	0	17	0	5	0	48	2
Gambia	Kombo Saint Mary	1286	5	492,564	0.0369	2,456	1	2,447	1	2,194	10	2,710	1	116	0	113	1	91	0	135	1
Gambia	Q2Ser River	1287	13	87,850	0.0369	432	2	433	1	384	1	482	2	21	0	20	0	16	0	25	0
Gambia	North Bank	1288	4	209,418	0.0369	1,056	0	1,055	1	970	4	1,141	2	52	0	50	0	43	0	59	0
Gambia	Western	1290	4	464,194	0.0369	2,185	3	2,169	11	1,905	13	2,459	19	101	0	96	1	77	1	122	2
Ghana	Ashanti	1324	18	4,872,310	0.0303	20,087	15	19,982	16	18,179	49	21,934	23	1,006	4	969	2	812	8	1,165	6
Ghana	Central	1326	4	1,928,780	0.0303	9,172	19	9,176	17	8,122	58	10,248	14	531	2	513	1	409	3	637	3
Ghana	Eastern	1327	4	2,693,720	0.0303	13,261	3	13,263	39	11,897	3	14,664	14	790	0	767	3	624	2	945	1
Ghana	Greater Accra	1328	3	4,345,450	0.0303	19,670	2	19,689	35	17,716	44	21,647	36	1,044	0	1,021	3	829	1	1,231	5
Ghana	Upper East	1330	16	1,069,720	0.0303	2,779	16	2,737	6	2,303	7	3,229	22	100	1	91	0	67	0	125	1
Ghana	Volta	1332	2	2,034,190	0.0303	10,257	88	10,263	103	9,279	93	11,213	53	647	6	635	9	526	5	755	4
Guinea-Bissau	Biombo	1386	3	103,423	0.0374	187	0	180	1	138	1	228	1	4	0	4	0	2	0	5	0
Kenya	Coast	51326	28	3,507,020	0.0369	16,124	78	16,142	83	14,764	78	17,487	128	840	7	817	11	690	8	969	7
Kenya	Nairobi	51328	2	3,501,660	0.0369	2,808	18	2,532	11	1,610	2	3,689	27	38	0	26	0	12	0	51	0
Kenya	Nyanza	51330	8	6,161,510	0.0369	33,401	129	33,319	68	29,437	142	37,418	251	1,934	12	1,865	4	1,491	18	2,291	24
Kenya	Rift Valley	51331	6	9,919,120	0.0369	16,861	293	15,967	266	11,894	408	20,726	413	533	6	436	10	288	8	683	0
Kenya	Western	51332	4	4,689,570	0.0369	21,830	24	21,742	40	19,229	34	24,485	24	1,069	3	1,030	1	829	3	1,278	3
Liberia	Grand Bassa	1817	3	303,787	0.0376	977	1	964	1	797	2	1,143	6	35	0	32	0	23	0	44	0
Liberia	Grand Cape Mount	1818	2	319,087	0.0376	1,772	0	1,765	11	1,388	15	2,150	28	102	1	90	1	57	0	133	2
Liberia	Lofa	1821	2	159,054	0.0376	1,019	9	1,023	9	885	7	1,149	11	67	1	65	0	50	1	82	1
Liberia	Nimba	1825	2	355,946	0.0376	777	6	743	6	560	3	959	6	24	0	20	0	12	0	31	1
Liberia	Sinoe	1828	2	424,039	0.0376	882	4	832	4	606	5	1,105	5	26	0	20	0	12	0	33	0
Libya	Banghazi	1839	2	888,002	0.0217	896	10	823	1	561	12	1,152	15	23	0	17	0	9	0	30	1
Madagascar	Analamanga	41752	2	3,038,530	0.0346	7,368	61	7,080	47	4,889	73	9,424	28	244	1	197	1	109	0	325	2
Madagascar	Analanjirifo	41753	2	1,188,460	0.0346	3,806	2	3,709	22	2,747	25	4,717	38	167	1	145	2	89	0	224	0
Madagascar	Anosy	41755	4	639,336	0.0346	2,072	3	2,034	13	1,718	4	2,413	8	89	0	83	0	63	0	110	1
Madagascar	Atsimo Andrefana	41756	5	1,222,760	0.0346	2,789	13	2,670	35	2,007	1	3,420	7	98	1	84	2	55	1	125	1
Madagascar	Atsimo Atsinanana	41757	3	685,524	0.0346	4,119	9	4,118	15	3,695	12	4,530	16	265	0	258	1	209	1	313	1
Madagascar	Haute Matsiatra	41763	2	1,242,210	0.0346	6,579	19	6,573	4	5,392	5	7,753	48	399	0	367	2	255	1	510	1
Madagascar	Menabe	41767	4	452,022	0.0346	1,265	4	1,233	6	970	26	1,543	6	48	0	43	1	29	1	63	1
Madagascar	Vakinankaratra	41770	2	1,812,550	0.0346	4,265	52	4,074	12	2,937	23	5,370	20	148	3	122	1	68	1	197	1
Madagascar	Vatovavy Fitovinany	41771	6	1,258,170	0.0346	8,084	11	8,094	15	7,380	20	8,794	16	555	2	546	1	453	3	639	0
Malawi	Southern Region	1890	14	7,304,820	0.0445	15,087	64	14,649	34	11,882	133	17,829	9	383	2	345	4	239	3	484	2

Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE
Mali	Koulikoro	1930	4	2,972,940	0.0450	14,006	65	13,707	68	11,350	44	16,591	55	653	11	595	2	433	5	822	11
Mali	Sikasso	1933	2	2,748,970	0.0450	13,633	34	13,503	37	11,197	54	15,825	38	707	5	656	3	471	1	875	8
Mozambique	Cabo Delgado	2112	5	1,915,120	0.0363	4,567	44	4,326	55	3,204	37	5,668	76	162	2	128	1	81	1	204	2
Mozambique	Gaza	2113	2	1,627,090	0.0363	651	15	398	12	171	4	838	16	11	1	4	0	1	0	11	0
Mozambique	Maputo (city)	2117	2	1,275,780	0.0363	954	10	828	22	540	11	1,248	9	12	0	8	0	4	0	16	0
Mozambique	Nampula	2118	2	4,290,360	0.0363	15,143	79	14,443	86	10,873	79	18,853	189	817	12	699	26	442	11	1,080	11
Namibia	Caprivi	2137	2	100,425	0.0252	45	0	27	1	9	0	61	1	1	0	0	0	0	0	1	0
Namibia	Kavango	2141	6	239,368	0.0252	374	0	360	3	276	2	466	1	13	0	11	0	7	0	18	0
Niger	Diffa	2203	3	411,442	0.0477	1,648	34	1,545	41	1,056	29	2,128	83	73	1	56	2	31	1	97	5
Niger	Tillaberi	2208	2	3,251,270	0.0477	22,880	123	23,048	68	19,469	41	26,191	110	1,387	8	1,321	10	998	6	1,702	7
Nigeria	Jigawa	2223	2	5,125,070	0.0393	20,174	61	19,359	103	14,431	98	25,126	98	973	3	809	12	518	6	1,264	8
Nigeria	Kaduna	2224	2	7,174,530	0.0393	53,425	146	53,496	278	47,227	275	59,490	135	3,893	7	3,722	31	2,956	12	4,624	22
Nigeria	Katsina	2226	4	6,545,700	0.0393	41,483	9	41,625	71	33,571	444	49,337	188	2,763	11	2,541	10	1,801	13	3,464	19
Nigeria	Lagos	2230	2	12,494,500	0.0393	108,491	21	108,425	73	99,496	131	117,602	464	8,699	2	8,457	10	7,047	6	10,143	102
Nigeria	Ogun	2232	3	4,460,800	0.0393	40,858	39	40,870	38	38,706	122	42,955	88	3,539	10	3,484	10	3,105	6	3,929	5
Nigeria	Osun	2234	3	4,466,810	0.0393	38,165	9	38,353	24	35,132	84	41,354	2	3,059	1	3,005	12	2,507	11	3,559	9
Nigeria	Oyo	2235	3	7,186,760	0.0393	63,733	233	63,863	158	58,902	611	68,586	14	5,443	32	5,315	23	4,520	81	6,272	12
Senegal	Fatick	47586	4	785,769	0.0359	3,554	12	3,558	24	3,051	11	4,039	0	170	1	163	2	124	1	207	3
Senegal	Tambacounda	2643	15	668,232	0.0359	3,798	48	3,788	58	3,038	47	4,486	68	257	5	234	7	164	3	317	9
Sudan	Central Ekuatoria	2748	5	779,795	0.0319	2,026	8	2,002	14	1,651	16	2,378	4	72	0	66	0	48	1	91	1
Sudan	Eastern Ekuatoria	2750	4	549,064	0.0319	1,020	8	956	7	723	3	1,249	13	35	0	29	0	20	0	44	0
Sudan	Khartoum	2753	6	7,313,860	0.0319	8,999	66	8,646	78	6,527	73	11,121	134	189	3	161	1	98	2	249	5
Sudan	Sennar	2762	2	1,932,200	0.0319	7,929	39	7,789	12	6,082	34	9,695	6	445	3	389	0	251	0	585	1
Sudan	Southern Kordofan	68785	3	2,294,240	0.0319	8,516	77	8,455	121	7,065	69	9,862	22	458	6	428	8	316	9	568	8
Sudan	Unity	2747	2	684,019	0.0319	938	24	846	33	559	14	1,216	31	29	1	20	1	10	1	39	2
Sudan	Upper Nile	37021	5	1,094,950	0.0319	421	16	360	15	212	12	570	26	9	0	5	0	2	0	12	1
Togo	Maritime	2972	3	2,878,020	0.0310	17,738	75	17,811	81	16,135	73	19,428	58	1,273	9	1,251	13	1,024	9	1,501	12
Togo	Plateaux	2973	2	1,510,980	0.0310	8,088	0	8,145	52	7,027	35	9,102	10	535	2	516	3	390	2	649	3
Tunisia	Tunis	3015	2	813,223	0.0165	322	3	291	5	196	1	413	1	5	0	3	0	2	0	6	0
Uganda	Kampala	42182	3	1,515,680	0.0439	10,104	9	10,128	34	9,095	47	11,189	27	535	2	523	3	421	0	630	2
Tanzania	Kagera	48361	4	2,567,560	0.0410	13,507	5	13,477	24	11,420	59	15,593	38	709	2	670	0	497	1	891	2
Tanzania	Kilimanjaro	48363	10	1,892,730	0.0410	3,100	5	2,965	9	2,329	3	3,778	9	72	0	62	0	42	0	93	0
Tanzania	Morogoro	48368	3	2,314,940	0.0410	12,546	10	12,456	57	9,853	22	15,118	9	714	3	641	0	420	2	915	7
Tanzania	Mwanza	48370	2	3,811,750	0.0410	25,644	176	25,676	147	20,971	156	30,296	299	1,703	26	1,592	13	1,090	13	2,166	49
Tanzania	Pwani	48373	3	1,132,920	0.0410	6,835	48	6,788	58	5,905	112	7,806	28	400	5	377	7	289	9	488	5

Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE	
Tanzania	Tanga	48381	15	2,148,450	0.0410	9,090	68	9,046	43	7,975	4	10,165	80	393	5	380	4	302	2	468	8	
Tanzania	Unguja North	48372	2	170,008	0.0410	739	3	730	1	620	5	854	2	29	0	28	0	20	0	37	0	
Tanzania	Unguja South	48379	9	156,851	0.0410	525	1	522	1	460	0	586	2	16	0	16	0	13	0	20	0	
Tanzania	Unguja Urban West	48382	2	522,047	0.0410	1,915	1	1,884	13	1,565	7	2,241	9	65	0	60	0	42	0	83	0	
Zambia	Luapula	3429	2	996,460	0.0465	8,839	35	8,878	9	7,723	39	10,004	39	673	5	646	2	494	2	821	6	
Zambia	Northern	3432	2	1,707,160	0.0465	12,757	136	12,637	60	10,301	40	15,020	343	900	25	815	11	566	8	1,119	42	
Zambia	North-Western	3431	2	769,142	0.0465	6,241	27	6,277	62	5,209	46	7,273	30	480	2	438	8	313	7	593	5	
Zambia	Western	3434	2	1,021,440	0.0465	3,538	7	3,468	31	2,625	15	4,340	6	161	1	140	1	87	0	212	1	
Zimbabwe	Mashonaland East	69550	2	1,261,370	0.0287	2,011	25	1,918	8	1,441	6	2,471	62	62	2	53	1	33	1	80	4	
ASIA						HbAS								HbSS								
Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE	
China	Taiwan Sheng	925	7	27,603,700	0.0119	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
India	Andaman and Nicobar	1484	8	404,656	0.0213	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
India	Andhra Pradesh	1485	21	90,604,700	0.0213	107,874	484	104,699	594	85,674	170	127,179	1,030	4,237	5	3,783	8	2,767	20	5,263	13	
India	Chhattisgarh	70075	8	24,165,500	0.0213	39,033	974	38,019	858	31,681	1,041	45,487	1,292	1,667	68	1,538	60	1,208	59	1,994	93	
India	Gujarat	1491	9	60,421,700	0.0213	88,628	1,232	84,780	1,478	69,441	1,339	104,503	1,357	3,530	79	3,234	125	2,451	85	4,319	73	
India	Jharkhand	70078	3	32,676,100	0.0213	8,250	104	5,867	78	3,463	251	10,581	23	146	7	68	0	30	3	170	7	
India	Karnataka	1494	6	62,125,300	0.0213	155,842	3,975	150,473	2,359	120,782	2,940	184,620	5,729	10,388	374	9,394	277	6,936	160	12,890	551	
India	Kerala	1495	8	37,121,100	0.0213	28,072	1,380	26,739	1,168	21,265	976	33,472	1,796	1,065	72	964	60	695	37	1,335	112	
India	Madhya Pradesh	70079	5	70,788,900	0.0213	91,872	868	86,888	355	66,685	195	111,513	1,021	3,871	90	3,346	55	2,273	19	4,854	99	
India	Maharashtra	1498	14	115,681,000	0.0213	132,957	1,982	127,739	932	103,303	1,164	157,602	3,352	5,387	96	4,678	52	3,511	31	6,644	182	
India	Orissa	1504	10	43,291,600	0.0213	55,843	470	54,318	958	46,360	1,248	62,997	256	2,323	24	2,205	43	1,794	22	2,736	36	
India	Rajasthan	1506	2	66,017,300	0.0213	37,251	1,550	29,852	689	17,925	1,129	49,064	2,014	1,114	83	689	38	356	31	1,362	87	
India	Tamil Nadu	1508	5	75,463,700	0.0213	120,340	739	113,504	1,512	81,234	1,251	152,672	838	6,326	86	5,221	24	3,271	123	8,169	200	
India	West Bengal	1511	4	94,360,600	0.0213	9,178	70	6,069	244	3,097	87	12,053	270	105	1	44	2	15	1	118	0	
Indonesia	NUSA TENGGARA TIMUR	1013683	7	3,897,850	0.0174	2	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
Indonesia	SUMATERA UTARA	1013681	2	13,056,600	0.0174	26	3	0	0	0	0	2	0	0	0	0	0	0	0	0	0	
Malaysia	Perak	1898	4	2,413,540	0.0198	4	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
Myanmar	Yangon Division	1012330	2	6,925,730	0.0167	10	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	
Saudi Arabia	Asir	2622	3	1,965,050	0.0214	1,721	70	1,606	75	1,198	53	2,131	89	53	3	44	4	27	2	69	5	
Saudi Arabia	Eastern Province	2624	12	3,702,700	0.0214	8,546	16	8,514	15	7,594	41	9,451	12	417	5	405	5	335	5	481	5	
Saudi Arabia	Madinah	2628	2	1,746,360	0.0214	1,075	7	737	12	386	11	1,437	56	31	1	13	0	6	0	34	2	
Saudi Arabia	Makkah	2629	5	7,054,850	0.0214	1,559	25	1,275	39	761	13	2,000	61	31	1	20	1	10	0	42	3	
Saudi Arabia	Riyad	2633	2	6,111,720	0.0214	4,032	6	3,672	11	2,526	43	5,166	24	103	1	82	1	46	1	136	1	

Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE	
Sri Lanka	Central	2736	3	2,618,530	0.0170	41	0	28	0	13	0	53	0	0	0	0	0	0	0	0	0	0
Sri Lanka	Eastern	2737	3	1,531,360	0.0170	81	0	61	1	34	0	106	1	1	0	0	0	0	0	0	1	0
Sri Lanka	North Central	2738	2	1,224,280	0.0170	40	1	29	0	16	0	51	0	0	0	0	0	0	0	0	0	0
Sri Lanka	North Western	2739	4	2,375,170	0.0170	37	0	23	0	10	0	47	1	0	0	0	0	0	0	0	0	0
Sri Lanka	Northern	2740	5	1,180,100	0.0170	97	0	74	1	43	0	124	0	1	0	0	0	0	0	0	1	0
Sri Lanka	Sabaragamuwa	2741	2	2,005,920	0.0170	36	1	26	0	13	1	49	1	0	0	0	0	0	0	0	0	0
Sri Lanka	Southern	2742	3	2,514,960	0.0170	53	1	33	1	14	0	73	2	0	0	0	0	0	0	0	0	0
Sri Lanka	Uva	2743	2	1,259,060	0.0170	30	1	21	0	10	0	40	1	0	0	0	0	0	0	0	0	0
Sri Lanka	Western	2744	4	5,700,630	0.0170	59	0	30	0	10	0	76	1	0	0	0	0	0	0	0	0	0
Syria	Damascus	2837	3	1,726,370	0.0219	5,394	36	5,348	68	4,693	47	6,105	8	284	3	269	5	212	4	347	3	3
Turkey	Adana	3017	4	1,943,940	0.0169	1,898	1	1,861	7	1,536	2	2,229	9	65	0	61	1	44	0	81	1	1
Turkey	Hatay	3052	14	1,350,070	0.0169	1,552	41	1,546	39	1,376	42	1,724	43	50	2	49	2	40	2	59	3	3
Turkey	Icel	3053	16	2,016,630	0.0169	2,104	55	2,081	70	1,873	55	2,317	64	78	3	76	3	62	3	92	3	3
Turkey	K.maras	3058	9	1,113,540	0.0169	16	0	9	0	3	0	21	0	0	0	0	0	0	0	0	0	0
UAE	Abu Dhabi	3173	2	1,737,000	0.0123	301	5	221	9	111	4	402	17	5	0	2	0	1	0	6	0	0

NORTH AMERICA						HbAS								HbSS								
Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE	
Canada	Saskatchewan	836	2	1,102,480	0.0113	17	0	4	0	1	0	14	0	0	0	0	0	0	0	0	0	0
USA	Alaska	3215	2	682,465	0.0137	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
USA	California	3218	5	39,245,000	0.0137	11,911	236	11,030	264	7,832	353	15,032	204	260	7	199	8	114	6	336	8	8
USA	Florida	3223	3	18,312,800	0.0137	9,196	183	7,639	255	5,045	114	11,799	281	290	4	181	8	95	3	346	12	12
USA	New York	3246	7	21,636,900	0.0137	11,519	95	11,029	115	8,774	12	13,627	236	259	3	224	1	153	2	323	10	10
USA	North Carolina	3247	3	8,821,450	0.0137	3,088	111	2,211	64	1,137	78	4,272	91	94	5	40	3	14	2	114	3	3

SOUTH AMERICA						HbAS								HbSS								
Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE	
Belize	Stann Creek	607	3	32,388	0.0240	139	0	140	0	122	1	157	0	9	0	9	0	7	0	11	0	0
Brazil	Amazonas	668	5	2,642,080	0.0150	264	1	143	6	61	6	326	13	5	0	2	0	0	0	5	0	0
Brazil	Bahia	669	9	13,676,400	0.0150	14,817	60	14,078	12	11,465	2	17,488	300	640	4	558	1	393	3	780	12	12
Brazil	Rio Grande Do Sul	685	8	12,029,500	0.0150	3,721	25	3,493	7	2,643	41	4,484	17	65	1	56	1	35	0	82	1	1
Brazil	Rondonia	686	2	1,540,980	0.0150	75	4	12	1	1	0	62	1	2	0	0	0	0	0	0	0	0
Brazil	Sao Paulo	689	15	45,722,000	0.0150	26,450	512	23,546	718	15,886	433	33,552	590	700	24	505	26	290	18	867	36	36
Chile	Antofagasta (ii)	883	3	601,091	0.0140	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
French Guiana	Cayenne	1271	3	167,704	0.0240	77	2	57	0	28	1	104	2	2	0	1	0	0	0	2	0	0

Country	Admin1 unit	GAUL	Surveys	Population	CBR	Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE
French Guiana	Saint-laurent-du-maroni	1272	3	63,209	0.0240	122	0	115	1	84	1	156	1	5	0	4	0	2	0	6	0
Haiti	Grand'Anse	72911	5	390,714	0.0257	1,144	1	1,124	8	827	3	1,431	5	54	0	45	1	27	0	71	1
Haiti	Nord-Ouest	1414	2	491,890	0.0257	1,205	1	1,168	4	913	6	1,479	5	48	0	41	0	26	0	62	0
Jamaica	Saint Andrew & Kingston	1640	3	688,272	0.0179	1,241	7	1,227	8	1,025	15	1,442	9	47	0	44	0	32	1	59	1
Mexico	Chiapas	1006348	2	4,239,470	0.0185	1,165	18	501	7	184	1	1,335	20	29	1	5	1	1	0	22	1
Mexico	Tabasco	1006373	2	2,125,840	0.0185	1,060	17	567	3	173	4	1,443	41	28	2	5	0	1	0	24	0
Nths Antilles	Not available	2157	2	200,555	0.0131	159	0	152	1	103	0	204	1	5	0	4	0	2	0	6	0
Peru	LORETO	1012328	3	1,032,140	0.0196	10	1	0	0	0	0	3	0	0	0	0	0	0	0	0	0
Peru	PASCO	1012373	2	280,795	0.0196	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Peru	UCAYALI	1012374	2	461,896	0.0196	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Venezuela	Amazonas	3302	14	59,011	0.0198	19	0	15	0	8	0	25	0	0	0	0	0	0	0	0	0
Venezuela	Anzoategui	3303	2	1,432,080	0.0198	888	1	718	3	388	8	1,196	9	22	0	12	0	5	0	27	0
Venezuela	Aragua	3305	3	1,716,840	0.0198	1,634	2	1,551	6	1,160	12	2,036	12	39	0	32	0	20	1	51	0
Venezuela	Bolivar	3307	3	1,543,730	0.0198	719	6	624	9	397	9	927	11	14	0	9	0	5	0	17	0
Venezuela	Falcon	3313	2	755,559	0.0198	911	0	865	8	611	9	1,160	7	30	0	24	0	14	0	39	0
Venezuela	Miranda	3317	2	3,079,300	0.0198	2,342	3	2,204	2	1,641	28	2,907	29	49	0	40	0	25	1	64	1
Venezuela	Sucre	3321	3	902,407	0.0198	787	3	729	4	531	7	981	9	21	0	17	0	11	0	27	0
Venezuela	Zulia	3325	4	4,230,060	0.0198	1,884	13	1,702	12	1,180	29	2,381	0	30	0	22	0	12	0	38	0

EUROPE

Country	Admin1 unit	GAUL	Surveys	Population	CBR	HbAS								HbSS							
						Mean	SE	Median	SE	Q25	SE	Q75	SE	Mean	SE	Median	SE	Q25	SE	Q75	SE
Belgium	Region wallonne	602	17	3,480,780	0.0114	668	2	641	2	516	3	784	3	8	0	7	0	5	0	10	0
France	Ile-de-France	1259	3	12,074,800	0.0124	5,827	15	5,574	21	4,318	20	7,110	26	116	1	99	1	64	1	151	1
France	Prov-Alpes-Cote-d'Azur	1269	2	4,902,190	0.0124	1,110	6	951	8	587	5	1,464	12	19	0	12	0	5	0	24	0
Greece	Ipeiros	1341	3	342,146	0.0101	292	1	280	2	207	1	363	2	11	0	9	0	6	0	14	0
Greece	Kentriki Makedonia	1342	19	1,923,720	0.0101	1,064	5	1,018	6	776	6	1,301	7	31	0	27	0	18	0	39	0
Greece	Kriti	1343	2	604,300	0.0101	35	1	16	0	4	0	46	1	1	0	0	0	0	0	0	0
Greece	Notio Aigaio	40766	3	305,049	0.0101	41	0	28	0	11	0	58	1	1	0	0	0	0	0	1	0
Greece	Stereia Ellada	1346	2	562,603	0.0101	502	1	490	2	381	1	609	2	20	0	18	0	11	0	26	0
Italy	Sardegna	1629	13	1,707,900	0.0091	30	0	26	0	18	0	37	0	0	0	0	0	0	0	0	0
Italy	Sicilia	1630	41	5,269,760	0.0091	792	10	765	10	644	10	910	12	11	0	10	0	8	0	14	0
Spain	Madrid, Comunidad De	2729	2	6,090,270	0.0106	705	3	576	5	336	3	934	6	7	0	4	0	2	0	9	0
U.K.	England	3182	5	51,767,000	0.0121	15,809	135	14,509	116	10,263	75	20,013	214	357	4	269	3	150	2	466	7

Web Table 4: Global, regional and national rates of neonates affected (HbAS and HbSS) based on median estimates presented in this report (termed MAP) and on data calculated by Modell and Darlison (termed M&D). Births for 2003 and 2010 are shown in thousands and are based on the 2003 UN Demographic Yearbook and the 2010 UN World Population Prospects, respectively.

Country	Births 2003	Births 2010	MAP				M&D		
			AS		SS		SS		
			n	%	n	%	n (2003)	n (2010)	%
Afghanistan	1,176	1,232	144	0.12	0	0.00	0	0	0.00
Albania	47	40	1,391	34.41	30	0.74	11	10	0.24
Algeria	627	680	10,486	15.42	191	0.28	50	54	0.08
Angola	604	758	112,443	148.37	8,364	11.04	9,560	11,996	15.83
Argentina	697	683	444	0.65	4	0.01	0	0	0.00
Armenia	36	46	0	0.00	0	0.00	0	0	0.00
Aruba	2	1	48	40.41	1	0.84	0	0	0.00
Australia	250	290	663	2.28	5	0.02	0	0	0.00
Austria	77	72	143	1.98	1	0.01	0	0	0.00
Azerbaijan	114	174	4	0.02	0	0.00	4	7	0.04
Bahamas	7	5	287	54.57	8	1.52	34	26	4.90
Bahrain	14	14	1,548	107.15	59	4.08	81	83	5.72
Bangladesh	3,536	3,206	3,459	1.08	16	0.00	0	0	0.00
Barbados	3	3	179	63.90	4	1.43	4	3	1.23
Belarus	89	107	0	0.00	0	0.00	0	0	0.00
Belgium	115	122	2,302	18.85	29	0.24	5	6	0.05
Belize	9	8	937	124.57	47	6.25	5	5	0.63
Benin	264	353	64,207	181.84	4,543	12.87	3,561	4,768	13.50
Bhutan	27	14	4	0.28	0	0.00	0	0	0.00
Bolivia (Plurinational State of)	300	258	6	0.02	0	0.00	0	0	0.00
Bosnia and Herzegovina	36	31	120	3.91	1	0.03	0	0	0.00
Botswana	53	45	300	6.63	4	0.09	0	0	0.00
Brazil	3,633	2,932	101,138	34.50	3,244	1.11	764	617	0.21
Brunei Darussalam	7	7	0	0.00	0	0.00	0	0	0.00
Bulgaria	67	75	1,076	14.36	17	0.23	0	0	0.00
Burkina Faso	491	689	71,083	103.17	3,124	4.53	1,065	1,493	2.17
Burundi	274	284	24,365	85.88	889	3.13	2,449	2,533	8.93
Cambodia	460	325	0	0.00	0	0.00	0	0	0.00
Cameroon	569	697	109,857	157.72	6,915	9.93	5,906	7,231	10.38
Canada	364	383	3,004	7.85	45	0.12	43	45	0.12
Cape Verde	13	10	625	61.22	18	1.76	6	5	0.44
Central African Republic	119	155	18,315	117.80	976	6.28	677	886	5.70
Chad	366	499	43,553	87.20	2,045	4.09	908	1,239	2.48
Chile	234	240	0	0.00	0	0.00	0	0	0.00
China	15,976	16,442	43	0.00	0	0.00	0	0	0.00
Colombia	1,060	875	14,464	16.53	321	0.37	537	444	0.51
Comoros	25	25	1,105	44.79	25	1.01	0	0	0.02
Congo	130	130	22,773	175.05	1,560	11.99	2,042	2,049	15.75
Congo, the Democratic Republic of the	2,325	2,856	489,745	171.50	38,217	13.38	20,113	24,698	8.65
Costa Rica	73	71	2,731	38.47	60	0.85	8	8	0.11
Côte d'Ivoire	720	712	74,168	104.19	3,567	5.01	1,564	1,546	2.17
Croatia	40	43	137	3.17	1	0.02	0	0	0.00
Cuba	151	108	6,578	61.16	240	2.23	259	185	1.72
Cyprus	8	10	147	15.11	2	0.21	0	0	0.03
Czech Republic	94	113	59	0.52	0	0.00	0	0	0.00
Denmark	65	62	11	0.18	0	0.00	2	2	0.03
Djibouti	31	25	3	0.12	0	0.00	0	0	0.00
Dominican Republic	201	214	9,461	44.27	247	1.16	722	769	3.60
Ecuador	329	273	45	0.16	0	0.00	12	10	0.04

Country	Births 2003	Births 2010	MAP				M&D		
			AS		SS		SS		
			n	%	n	%	n (2003)	n (2010)	%
Egypt	1,774	1,884	47,754	25.35	1,196	0.63	19	21	0.01
El Salvador	125	124	4,039	32.57	80	0.65	8	8	0.07
Equatorial Guinea	18	25	5,059	203.35	385	15.48	182	249	10.00
Eritrea	151	179	767	4.28	9	0.05	0	0	0.00
Estonia	13	16	0	0.00	0	0.00	0	0	0.00
Ethiopia	2,380	2,550	11,197	4.39	137	0.05	1	2	0.00
Fiji	17	18	0	0.00	0	0.00	0	0	0.00
Finland	57	61	0	0.00	0	0.00	0	0	0.00
France	759	776	22,040	28.39	520	0.67	79	81	0.10
French Guiana	6	6	186	33.56	5	0.90	4	4	0.68
French Polynesia	4	4	0	0.00	0	0.00	0	0	0.00
Gabon	46	41	9,342	230.51	864	21.32	736	643	15.88
Gambia	58	65	8,657	133.98	418	6.47	305	342	5.30
Georgia	46	49	0	0.00	0	0.00	0	0	0.00
Germany	710	714	5,940	8.32	85	0.12	23	23	0.03
Ghana	726	737	102,229	138.62	5,474	7.42	4,847	4,924	6.68
Greece	105	113	6,667	58.97	206	1.82	1	1	0.01
Grenada	2	2	174	86.69	6	2.99	5	5	2.50
Guadeloupe	8	6	459	74.39	14	2.27	16	13	2.16
Guam	3	3	0	0.00	0	0.00	0	0	0.00
Guatemala	375	453	10,285	22.71	252	0.56	0	0	0.00
Guinea	310	388	70,965	182.82	5,232	13.48	4,325	5,418	13.96
Guinea-Bissau	53	62	5,011	81.35	192	3.12	0	0	1.83
Guyana	16	13	622	46.95	17	1.28	11	9	0.68
Haiti	249	262	26,801	102.35	1,088	4.16	622	655	2.50
Honduras	230	196	17,783	90.58	772	3.93	0	0	0.00
Hong Kong	47	64	0	0.00	0	0.00	0	0	0.00
Hungary	94	101	59	0.59	0	0.00	0	0	0.00
Iceland	4	5	0	0.00	0	0.00	0	0	0.00
India	26,492	25,495	996,563	39.09	42,016	1.65	25,768	24,798	0.97
Indonesia	4,863	4,046	328	0.08	1	0.00	0	0	0.00
Iran, Islamic Republic of	1,170	1,216	25,768	21.18	862	0.71	96	100	0.08
Iraq	903	1,072	23,488	21.91	629	0.59	499	592	0.55
Ireland	62	72	60	0.84	0	0.00	0	0	0.00
Italy	541	547	5,635	10.30	91	0.17	3	3	0.01
Jamaica	57	49	3,998	81.81	137	2.80	172	148	3.03
Japan	1,123	1,079	0	0.00	0	0.00	0	0	0.00
Jordan	148	153	6,365	41.57	145	0.95	4	4	0.03
Kazakhstan	247	325	1	0.00	0	0.00	0	0	0.00
Kenya	1,121	1,507	98,471	65.35	4,475	2.97	7,872	10,578	7.02
Korea, Democratic People's Republic of	495	340	0	0.00	0	0.00	0	0	0.00
Korea, Republic of	494	480	0	0.00	0	0.00	0	0	0.00
Kuwait	45	53	3,824	72.04	116	2.19	37	44	0.83
Kyrgyzstan	105	133	0	0.00	0	0.00	0	0	0.00
Lao People's Democratic Republic	218	140	0	0.00	0	0.00	0	0	0.00
Latvia	21	25	0	0.00	0	0.00	0	0	0.00
Lebanon	77	63	6,382	101.09	251	3.98	6	5	0.08
Lesotho	75	56	40	0.72	0	0.00	0	0	0.00
Liberia	127	154	13,217	85.69	562	3.64	105	127	0.82
Libyan Arab Jamahiriya	111	142	6,020	42.38	188	1.32	34	44	0.31
Lithuania	31	35	0	0.00	0	0.00	0	0	0.00
Luxembourg	5	6	48	8.26	0	0.00	0	0	0.06
Madagascar	609	697	69,295	99.41	3,379	4.85	2,084	2,383	3.42
Malawi	579	698	47,370	67.85	1,688	2.42	917	1,106	1.58

Country	Births 2003	Births 2010	MAP				M&D		
			AS		SS		SS		
			n	%	n	%	n (2003)	n (2010)	%
Malaysia	534	553	1	0.00	0	0.00	0	0	0.00
Maldives	5	5	0	0.00	0	0.00	0	0	0.01
Mali	494	601	56,236	93.53	2,701	4.49	4,421	5,385	8.96
Malta	4	4	143	38.33	2	0.54	0	0	0.00
Martinique	6	5	391	83.02	12	2.55	9	7	1.52
Mauritania	110	110	9,505	86.54	433	3.94	159	158	1.44
Mauritius	19	16	0	0.00	0	0.00	1	1	0.05
Mayotte	2	6	447	68.90	12	1.85	0	1	0.23
Mexico	2,564	2,045	31,120	15.21	641	0.31	105	84	0.04
Micronesia, Federated States of	3	3	0	0.00	0	0.00	0	0	0.00
Moldova, Republic of	36	43	1	0.02	0	0.00	0	0	0.00
Mongolia	46	61	0	0.00	0	0.00	0	0	0.00
Montenegro	87	8	72	9.18	1	0.13	0	0	0.00
Morocco	560	597	3,202	5.36	40	0.07	118	126	0.21
Mozambique	754	850	42,588	50.10	1,645	1.94	170	191	0.23
Myanmar	984	843	40	0.05	0	0.00	0	0	0.00
Namibia	65	56	1,138	20.42	30	0.54	10	9	0.15
Nepal	788	695	1,558	2.24	11	0.02	0	0	0.00
Netherlands	200	180	4,759	26.47	82	0.46	10	9	0.05
Netherlands Antilles	4	3	150	57.09	4	1.52	9	7	2.50
New Caledonia	4	4	0	0.00	0	0.00	0	0	0.00
New Zealand	56	62	0	0.00	0	0.00	9	10	0.16
Nicaragua	190	133	6,514	49.08	204	1.54	4	3	0.02
Niger	506	758	90,671	119.66	4,965	6.55	6,046	9,045	11.94
Nigeria	4,895	6,219	1,136,909	182.80	85,186	13.70	77,192	98,083	15.77
Norway	57	60	0	0.00	0	0.00	2	2	0.04
Oman	90	50	32	0.64	0	0.00	222	124	2.47
Pakistan	3,913	4,859	18,962	3.90	228	0.05	101	126	0.03
Panama	63	67	4,457	66.53	140	2.09	8	9	0.13
Papua New Guinea	175	200	0	0.00	0	0.00	0	0	0.00
Paraguay	168	153	664	4.34	7	0.05	0	0	0.00
Peru	649	578	2	0.00	0	0.00	25	22	0.04
Philippines	1,703	2,294	0	0.00	0	0.00	0	0	0.00
Poland	351	415	59	0.14	0	0.00	0	0	0.00
Portugal	113	90	2,170	24.11	41	0.46	3	2	0.03
Puerto Rico	59	52	326	6.32	2	0.04	23	21	0.40
Qatar	13	17	1,365	78.02	41	2.34	7	10	0.55
Réunion	14	14	6	0.42	0	0.00	1	1	0.10
Romania	213	218	320	1.46	2	0.01	0	0	0.00
Russian Federation	1,475	1,656	14	0.01	0	0.00	0	0	0.00
Rwanda	313	417	22,857	54.78	601	1.44	125	167	0.40
Saint Lucia	3	3	311	105.76	12	4.08	7	7	2.50
Saint Vincent and the Grenadines	2	2	176	98.46	6	3.36	2	2	1.16
Samoa	5	13	0	0.00	0	0.00	0	0	0.00
Saudi Arabia	766	561	19,447	34.68	691	1.23	5,397	3,950	7.04
Sao Tome and Principe	6	5	763	154.66	41	8.31	49	40	8.10
Senegal	407	462	53,002	114.75	2,535	5.49	1,895	2,153	4.66
Serbia	87	108	830	7.67	10	0.09	0	0	0.00
Sierra Leone	246	213	39,530	185.56	2,838	13.32	3,243	2,808	13.18
Singapore	46	45	0	0.00	0	0.00	0	0	0.00
Slovakia	52	58	12	0.21	0	0.00	0	0	0.00
Slovenia	17	20	22	1.10	0	0.00	0	0	0.00
Solomon Islands	15	16	0	0.00	0	0.00	0	0	0.00
Somalia	505	402	1,192	2.96	13	0.03	0	0	0.00

Country	Births 2003	Births 2010	MAP				M&D		
			AS		SS		SS		
			n	%	n	%	n (2003)	n (2010)	%
South Africa	1,258	1,036	4,946	4.78	73	0.07	0	0	0.00
Spain	440	480	7,324	15.25	149	0.31	0	1	0.00
Sri Lanka	363	347	425	1.22	2	0.01	0	0	0.00
Sudan	1,157	1,378	98,244	71.32	4,567	3.32	483	575	0.42
Suriname	10	9	653	69.97	19	2.04	13	12	1.30
Swaziland	36	34	398	11.61	4	0.12	1	1	0.03
Sweden	99	112	2	0.02	0	0.00	4	5	0.04
Switzerland	81	76	622	8.19	6	0.08	0	0	0.00
Syrian Arab Republic	493	496	24,448	49.34	901	1.82	66	66	0.13
Tajikistan	178	196	0	0.00	0	0.00	0	0	0.00
Tanzania, United Republic of	1,232	1,846	203,747	110.36	11,022	5.97	8,655	12,965	7.02
Thailand	1,054	784	0	0.00	0	0.00	0	0	0.00
Timor-Leste	28	44	0	0.00	0	0.00	0	0	0.00
Togo	202	210	34,057	162.19	2,175	10.36	1,746	1,816	8.65
Trinidad and Tobago	19	19	1,648	85.75	56	2.91	30	31	1.60
Tonga	2	3	0	0.00	0	0.00	0	0	0.00
Tunisia	166	171	4,420	25.82	81	0.47	63	64	0.38
Turkey	1,478	1,279	12,852	10.05	330	0.26	116	100	0.08
Turkmenistan	139	110	4	0.04	0	0.00	0	0	0.00
Uganda	1,164	1,484	189,801	127.92	10,143	6.84	8,216	10,469	7.06
Ukraine	410	495	33	0.07	0	0.00	0	0	0.00
United Arab Emirates	75	57	402	7.03	4	0.07	24	18	0.32
United Kingdom	697	749	15,746	21.02	296	0.40	108	116	0.16
United States of America	4,194	4,353	100,150	23.01	2,843	0.65	838	869	0.20
Uruguay	51	49	82	1.68	1	0.02	0	0	0.00
Uzbekistan	520	578	1	0.00	0	0.00	0	0	0.00
Vanuatu	6	7	0	0.00	0	0.00	0	0	0.00
Venezuela, Bolivarian Republic of	492	575	19,272	33.51	435	0.76	108	126	0.22
Viet Nam	1,807	1,415	0	0.00	0	0.00	0	0	0.00
Virgin Islands, U.S.	2	1	25	18.96	0	0.00	4	2	1.60
Western Sahara	9	9	144	15.32	3	0.32	2	2	0.21
Yemen	930	905	6,411	7.08	80	0.09	119	116	0.13
Zambia	457	616	88,801	144.09	5,652	9.17	3,346	4,517	7.33
Zimbabwe	408	363	16,345	45.04	483	1.33	162	144	0.40

WHO regions	Births 2003	Births 2010	MAP				M&D		
			AS		SS		SS		
			n	%	n	%	n (2003)	n (2010)	% (2003/2010)
AFRO	22,895	31,755	3,610,851	113.71	238,083	7.50	184,812	232,336	8.07/7.32
AMRO	16,609	15,210	391,257	25.72	13,104	0.86	4,432	4,156	0.27/0.27
EMRO	16,798	13,971	256,643	18.37	8,239	0.59	7,389	6,072	0.44/0.43
EURO	10,459	10,976	121,601	11.08	3,271	0.30	376	368	0.04/0.03
SEARO	38,139	35,820	1,020,489	28.49	42,597	1.19	25,768	24,798	0.68/0.69
WPRO	23,914	23,535	1,150	0.05	9	0.00	9	10	0.00/0.00

World	Births 2003	Births 2010	MAP				M&D		
			AS		SS		SS		
			n	%	n	%	n (2003)	n (2010)	% (2003/2010)
WORLD	128,814	131,266	5,476,407	41.72	312,302	2.38	222,785	267,740	1.73/2.04

Web Table 5. MCMC output parameter values. Summary statistics presented are the mean and median values, standard deviation (std), interquartile range (IQR) and 95% Bayesian credible interval (95 BCI). The scale parameter is measured in units of earth radii. Values are presented to two significant figures.

Parameter	Symbol	Mean	Median	Std	IQR	95 BCI
Nugget variance	V	0.450	0.443	0.079	0.062	0.204
Amplitude (or partial sill)	φ	2.715	2.665	0.348	0.174	0.836
Scale (or range)	θ	0.199	0.199	0.001	0.001	0.003

Web Table 6. Summary of data selection, mapping and modelling assumptions and suggestions for future improvements.

Data

- We assumed that surveys not reporting any selection bias in the sampling methodology were representative of the local population.*
A complete and consistent description of the sampling method used would remove any ambiguity about the representativeness of the sample.
- We assumed that surveys targeting specific ethnic groups could not be included.*
Studies targeting a specific ethnic group cannot be considered representative of the overall local population. Provided that the sample size is not too small, random or universal surveys testing all ethnic groups without selection bias can be considered as representative.
- We assumed that surveys georeferenced at the province or country level were too imprecise to be included.*
Micromapping studies have shown that HbS allele frequency can highly vary over short distances. Surveys testing individuals from all over a region or country without giving the individuals' origin could not be used. Similarly, surveys not reporting the location of the survey also had to be excluded. Systematic reporting of basic geographical information would increase the number of included studies.
- We assumed that the diagnostic method had a minimum effect on the testing results.*
The diagnostic method is not consistently reported. Most diagnostic methods for sickle haemoglobin can be considered as reliable.
- We assumed that interactions between HbS, HbC and β -thalassaemias were complex.*
The number of individuals with compound status is relatively small and data available is very limited. Calculating newborn estimates for these compound individuals would require both the availability of a similar database on HbC and β -thalassaemia, and a good understanding of their interactions.

Mapping and modelling process

- We assumed that the crude birth rate was homogeneous within a country.*
No data on crude birth rate are consistently available at sub-national levels. Local or national studies could allow us to incorporate more precise information.
- We assumed that populations were in Hardy-Weinberg equilibrium (HWE), and derived HbS allele frequency based on this assumption.*
In some cases, we have observed differences between observed and HWE predicted genotype counts but there is currently not enough information on this relationship to use it within a global model.
- We assumed that information on consanguinity was too crude to be incorporated.*
Information on consanguinity is not consistently reported in surveys. Moreover, although there is some global information (e.g. www.consang.net), only crude national information is available. Including such data in the model would likely have resulted in adding more uncertainty in the predictions.
- We assumed that the amount of data available was not sufficient to develop a spatio-temporal model.*
Large amounts of data are required in order to analyse temporal changes at the global scale. Such a study might be possible in data-rich areas.

Web Dataset. List of published and unpublished sources from which data have been used as input into our model. Other references are ordered alphabetically by surname.

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