## **INTERPRETIVE SUMMARY**

## 2 Foot and leg conformation traits have a small impact on genomic predictions of claw

3 **disorders in Norwegian Red cows.** Ødegård et al. Predictive correlations of genomic breeding

4 values (GEBV) for corkscrew claw, infectious claw disorder and laminitis related claw disorder

5 were calculated using information on claw disorders only (recorded at claw trimming); and by

6 analyzing claw disorders together with genetically correlated foot and leg conformation traits.

7 Including the correlated traits gave a slight increase in the predictive correlation of GEBV for

8 corkscrew claw, but had no effect on the other claw disorders.

## PREDICTIVE CORRELATION OF GEBV FOR CLAW DISORDERS

10	Foot and leg conformation traits have a small impact on genomic predictions of claw
11	disorders in Norwegian Red cows.
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## 29 ABSTRACT

The aim of this study was to evaluate whether the predictive correlation of genomic breeding 30 values (GEBV) for claw disorders increased by including genetically correlated traits as additional 31 information in the analyses. Predictive correlations of GEBV for claw disorders were calculated 32 based on claw disorders only and by analyzing claw disorders together with genetically correlated 33 foot and leg conformation traits. The claw disorders analyzed were: corkscrew claw (CSC); 34 infectious claw disorder (**INF**), including dermatitis, heel horn erosion and interdigital phlegmon; 35 and laminitis related claw disorder (LAM), including sole ulcer, white line disorder and 36 37 hemorrhage of sole and white line. The foot and leg conformation traits included were: hoof quality (HQ), foot angle (FA), rear leg rear view new (RLRV\_N) and rear leg rear view old (RLRV\_O). 38 The data consisted of 183,728 daughters with claw health records and 421,319 daughters with foot 39 and leg conformation scores. A 25K/54K SNP dataset containing 48,249 SNP was available for 40 the analyses. The number of genotyped sires with daughter information in the analyses was 1,093 41 including claw disorders, and 3,111 including claw disorders and foot and leg conformation traits. 42 Predictive correlations of GEBV for CSC, INF and LAM were calculated from a 10-fold cross-43 validation and from an additional validation set including the youngest sires. Only sires having 44 daughters with claw health records were in the validation sets, thus increasing the reference 45 population when adding foot and leg conformation traits. The results showed marginal 46 improvement in the predictive correlation of GEBV for CSC when including HQ and FA, both in 47 48 10-fold cross-validation (from 0.35 to 0.37) and in the validation including the youngest sires (from 0.38 to 0.49). For INF and LAM, including foot and leg conformation traits had no effect 49 on the predictive correlation of GEBV. Claw disorders are novel traits with a limited amount of 50 51 historical data and therefore a small reference population. Increasing the reference population by

including sires with daughter information on foot and leg conformation traits had small impact on
the predictive correlation of GEBV. However, the small increase in predictive correlation of
GEBV for CSC show a possible gain when including moderate to high genetically correlated traits.

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56 Keywords: dairy cow, genomic breeding value, claw health, Norwegian Red

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## INTRODUCTION

Claw health is important for animal welfare (Bruijnis et al., 2012) and for dairy production 59 economy (Bruijnis et al., 2010) by influencing milk production (Sogstad et al., 2007), fertility and 60 production diseases (Sogstad et al., 2006). In Norway, claw health status at claw trimming has 61 been reported to the Norwegian Dairy Herd Recording System since 2004. The frequencies of claw 62 disorders in Norwegian Red are in general low, ranging from 0.2% (interdigital phlegmon) to 10% 63 (corkscrew claw (CSC)) (Ødegård et al., 2013). Heritabilities (on the underlying scale) of claw 64 disorders in different breeds ranged from 0.06 to 0.23 (e.g. Swalve et al., 2008; Buch et al., 2011; 65 Ødegård et al., 2013). Estimated genetic correlations between claw disorders and foot and leg 66 conformation traits are low to moderate, but with some variations between studies and breeds (e.g. 67 Uggla et al., 2008; Van der Waaij et al., 2005; Ødegård et al., 2014a). Ødegård et al. (2014a) 68 showed that 7 out of 15 genetic correlations between claw disorders and foot and leg conformation 69 traits in Norwegian Red were significantly different from zero, ranging from -0.86 to 0.26. The 70 strongest genetic correlation was found between CSC from claw trimming and hoof quality (HQ) 71 from conformation score, which are supposed to measure the same trait. Hoof quality has the same 72 definition as CSC, but are recorded by breeding advisors when the cow is standing. 73

75 Genomic selection has a huge potential to increase genetic gain (Meuwissen et al., 2001). In the selection program for Norwegian Red, the accuracy of genomic breeding values (GEBV) are low 76 compared to the accuracy of estimated breeding values (EBV) from progeny testing, especially for 77 health and fertility traits (e.g. Luan et al., 2009; Svendsen et al., 2013; Haugaard et al., 2014). The 78 accuracy of GEBV, calculated as the correlation between EBV and GEBV, ranged from 0.16 79 (stillbirth, direct) to 0.77 (slaughter classification) in Norwegian Red (Svendsen et al., 2013). 80 Similar results were found in other studies, where production traits showed higher accuracy or 81 reliability of GEBV than functional traits (e.g. Solberg et al., 2011; Gao et al., 2013; Zhou et al., 82 2014). With novel traits such as claw disorders, the historical data and reference population is 83 limited making genomic selection challenging. One way to improve the accuracy of GEBV is to 84 increase the size of the reference population (e.g. Hayes et al., 2009) by including genetically 85 correlated traits. Svendsen et al. (2013) calculated relatively high accuracy of GEBV for foot and 86 leg conformation traits, ranging from 0.60 to 0.71. Foot and leg conformation traits that are 87 genetically correlated to claw disorders may contribute additional information and thereby 88 improve the predictive correlation of GEBV for claw disorders. 89

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The aims were: 1) genomic analyses of claw disorders in Norwegian Red. First to evaluate predictive correlation of GEBV for CSC, infectious claw disorder (**INF**) and laminitis related claw disorder (**LAM**) and 2) to examine whether including genetically correlated foot and leg conformation traits in the analyses increased the genomic prediction of CSC, INF and LAM.

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#### **MATERIALS AND METHODS**

97 Data and editing

98 *Claw health.* Claw health status at claw trimming reported to the Norwegian Dairy Herd Recording System from 2004 to 2013 were included in the analyses. Nine different claw disorders were 99 recorded at claw trimming; CSC, dermatitis, heel horn erosion, interdigital phlegmon, sole ulcer, 100 101 white line disorder, hemorrhage of sole and white line, lameness and acute trauma. Cows with no claw disorders present at claw trimming were recorded as having normal claws. Based on 102 frequencies of and genetic correlations among claw disorders (Ødegård et al., 2013); 1 claw 103 disorder and 2 groups of claw disorders were included in the analyses: CSC, INF (including 104 dermatitis, heel horn erosion and interdigital phlegmon) and LAM (including sole ulcer, white line 105 disorder and hemorrhage of sole and white line). A cow was defined as unaffected (0) or affected 106 (1) for CSC, INF and LAM in each parity in which the cow had at least 1 record from claw 107 trimming. The claw trimming practice varies among herds; in some herds all cows are routinely 108 109 claw trimmed once a year, whereas in others, claw trimming is carried out occasionally on selected cows only. In Norway, claw trimming is performed by: professional claw trimmers (with 110 certification), other claw trimmers (working as claw trimmers without certification), farmers or 111 112 others (e.g. veterinarians). More details of claw health data in Norway can be found in Ødegård et al. (2013). 113

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Data was edited as described by Ødegård et al. (2013): only lactating cows with recorded claw health records; daughters of Norwegian Red AI sires; at least 1 claw health record in a parity; and herds reporting at least 10% or 10 normal claw records from 2004 to 2013 (this to exclude herds reporting only affected cows) were included in the analyses. Sires were required to have at least 30 daughters with claw health records. Data included in the analyses consisted of 281,835 claw health records from 183,728 daughters of 1,093 sires and the number of herds were 6,976. The
mean frequencies of CSC, INF and LAM after editing were 11%, 7% and 8%, respectively.

*Foot and leg conformation.* Foot and leg conformation was scored on 1<sup>st</sup> parity cows and reported 123 to the Norwegian Dairy Herd Recording System. Breeding advisors, at present about 50 people, 124 score 4 defined foot and leg conformation traits: HQ, foot angle (FA), rear leg rear view (RLRV) 125 and rear leg side view (**RLSV**) on a linear scale from 1 to 9. The definition and optimal value of 126 RLRV changed in 2010, hence 2 traits were defined: RLRV new (RLRV N) and RLRV old 127 (RLRV O). The optimum values were: 9 for HQ, 8 for RLRV N, and 5 for FA, RLRV O and 128 RLSV. Based on results from Ødegård et al. (2014a), the foot and leg conformation traits included 129 in the analyses were: HQ, FA, RLRV\_N and RLRV\_O (these traits had a genetic correlation 130 131 significantly different from zero for at least one claw disorder). Available foot and leg conformation score were: HQ from 1996 to 2013, FA from 1987 to 2013, RLRV\_N from 2010 to 132 2013, and RLRV O from 1987 to 2009. 133

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Data was edited as described in Ødegård et al. (2014a): only daughters of Norwegian Red AI sires; age at first calving between 18 and 33 months; and conformation scored within a defined time period (months after calving) were included. The data analyzed consisted of 305,195 daughters of 2,183 sires for HQ; 421,319 daughters of 3,111 sires for FA; 52,330 daughters of 571 sires for RLRV\_N; and 368,834 daughters of 2,710 sires for RLRV\_O. Number of records for each combination of claw disorders and foot and leg conformation traits are given in Table 1.

142 SNP dataset. An imputed 25K/54K SNP dataset was available for the analyses. Not all SNPs included in the 25K SNP-chip are in the 54K SNP-chip, so to exploit all available SNPs the dataset 143 was imputed from 25K to 54K and vice versa. For details of the imputation refer to Solberg et al. 144 (2011). After standard editing: removal of animals with an individual call rate < 97 %, deletion of 145 Mendelian errors for animals with known parents, removal of SNP with Mendelian error rate > 2.5 146 %, deletion of SNP with a call rate < 25 %, and removal of SNP with MAF < 0.05, the dataset 147 contained 48,249 SNP for a total of 3,768 Norwegian Red AI sires. Sires with genotype and 148 informative daughters (with data on claw disorders, foot and leg conformation traits or both) were 149 included in the analyses. Number of sires for each trait combination are given in Table 1. 150

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#### 152 Statistical analyses

Three sets of trait combinations were analyzed: (1) CSC, INF and LAM (CH); (2) CSC, INF, LAM, HQ and FA (CF1); and (3) CSC, INF, LAM, RLRV\_N and RLRV\_O (CF2). Because of convergence issues it was not possible to analyze all the claw disorders and foot and leg conformation traits together.

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*Estimated breeding values.* Breeding values for CSC, INF and LAM were predicted using a linear
sire model including effects as described in Ødegård et al. (2013). The model in matrix notation
was:

## 161 $y = X\beta + Z_hh + Z_ss + e$

where **y** is a vector of observations on the trait,  $\beta$  is a vector of systematic effects, **h** is a vector of random herd effects, **s** is a vector of sire effects, **e** is a vector of residuals, and **X**, **Z**<sub>h</sub> and **Z**<sub>s</sub> are the corresponding incidence matrices. The systematic effects were: parity with 4 classes, where the 4<sup>th</sup>

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class included parity 4 to 13; year and month of calving with 119 classes; time of claw trimming
(in months after calving) with 12 classes; and claw trimmer with 4 classes: (1) professional claw
trimmer, (2) other claw trimmer, (3) farmer and (4) other person (e.g. veterinarian). The herd
effects included 6,976 levels.

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Breeding values for HQ, FA, RLRV\_N and RLRV\_O were predicted using a linear sire model
including effects described in Ødegård et al. (2014a). The model in matrix notation was:

# 172 $y = X\beta + Z_{hy}hy + Z_{s}s + e$

where y is a vector of observations of the trait;  $\beta$  is a vector of systematic effects including year 173 and month of calving, time from calving (months) and time from milking (hours) to scoring, and 174 175 age at scoring (in months); hy is a vector of random herd-year effects; s is a vector of sire effects; 176 e is a vector of residuals; and X, Z<sub>hy</sub> and Z<sub>s</sub> are the corresponding incidence matrices. Year and 177 month of calving had 216 levels for HQ, 315 levels for FA, 51 levels for RLRV\_N and 275 levels for RLRV\_O; time from calving (months) and time from milking (hours) to scoring had 96 levels 178 179 for HQ, FA, RLRV\_N and RLRV\_O; and age at scoring (in months) had 7 levels for HQ, FA, 180 RLRV\_N and RLRV\_O. The herd-year effect included 98,820 levels for HQ; 149,249 levels for FA; 12,661 for RLRV\_N; and 136,566 for RLRV O. 181

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The 3 datasets were analyzed using multivariate models with (co)variances: var(h) =  $\mathbf{H} \otimes \mathbf{I}$ , var(hy) =  $\mathbf{HY} \otimes \mathbf{I}$ , var(s) =  $\mathbf{G_0} \otimes \mathbf{A}$ , and var(e) =  $\mathbf{R} \otimes \mathbf{I}$ , where  $\mathbf{H}$  is the 3×3 herd (co)variance matrix; **HY** is the 2×2 herd-year variance matrix (co-variances were assumed to be zero);  $\mathbf{A}$  is the additive genetic relationship matrix;  $\mathbf{I}$  are identity matrices;  $\mathbf{G_0}$  and  $\mathbf{R}$  are the 3×3, 5×5 and 5×5 corresponding genetic and residual (co)variance matrices for the datasets CH, CF1 and CF2, respectively. The residual covariance between RLRV\_N and RLRV\_O was assumed zero, because
no cows had observation on both traits. The pedigree of sires were traced as far as possible resulting
in a pedigree file of 15,172 animals for CH and 26,120 animals for CF1 and CF2.

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*Deregressed proofs.* To calculate deregressed proofs (DRP) (Lidauer and Strandén, 1999; Vuori
et al., 2006), the EBV was used as the response variable and the residuals were weighted by
effective daughter contribution (Fikse and Banos, 2001) calculated from reliabilities of EBV.

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*Genomic breeding values.* Genomic breeding values were predicted using GBLUP (Meuwissen
et al., 2001). Deregressed proofs were used as response variables for genomic predictions. The
model in matrix notation was:

199  $y = 1\mu + Zg + e$ 

200 where y is a vector of DRP, 1 is a vector of ones,  $\mu$  is the overall mean, g is a vector of genomic effects, Z is the incidence matrix of  $\mathbf{g}$ , and  $\mathbf{e}$  is a vector of residuals. It was assumed that  $var(\mathbf{g}) =$ 201  $G_0 \otimes G$  and  $var(e) = \mathbf{R} \otimes \mathbf{D}$ ; where G is the genomic relationship matrix; **D** is a diagonal matrix 202 containing weighting factors for the residuals; G<sub>0</sub> and **R** are the  $3\times3$ ,  $5\times5$  and  $5\times5$  corresponding 203 genetic and residual (co)variance matrix for CH, CF1 and CF2, respectively. The residual 204 covariances between claw disorders and foot and leg conformation traits in CF1 and CF2 were set 205 to zero. The residuals were weighted by reliabilities of EBV. The inverse G-matrix used in 206 207 prediction of GEBV was obtained using the G-matrix package (Su and Madsen, 2012) and consisted of 1,093 sires in CH and 3,111 sire in CF1 and CF2. 208

*Predictive correlation of GEBV.* Predictive correlation of GEBV was calculated as the correlation
between GEBV and DRP, where DRP was calculated from EBV predicted with all available
information for each of the 3 datasets (CH, CF1 and CF2).

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10-fold cross-validation. A 10-fold cross-validation (CV) was performed to assess predictive 214 correlations of GEBV for CSC, INF and LAM from the datasets CH, CF1 and CF2. The 1,093 215 sires with daughter information on claw health were randomly assigned to 10 groups, including 216 109 or 110 sires. Therefore a sire was only represented in 1 group. In the CV, 1 group was used as 217 validation set and the remaining 9 constituted the reference population. Sires having daughters 218 with only foot and leg conformation scores were included in the reference population. The 219 reference populations consisted of 983(984), 3,001(3,002) and 3,001(3,002) sires for the datasets 220 221 CH, CF1 and CF2, respectively.

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*Validation by youngest sires.* An additional validation set (VAL) consisting of the youngest sires
having daughters with claw health information was analyzed. This validation set included 190 sires
(born in 2007, 2008 and 2009), and the reference populations (sires born before 2007) consisted
of 903, 2,797 and 2,797 sires for CH, CF1 and CF2, respectively.

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The DMU software (Madsen and Jensen, 2010) was used to estimate (co)variances and predict EBV and GEBV. (Co)variances estimated from the full datasets were used in prediction of EBV for each of the reference populations in CV and VAL. Estimated heritabilities and genetic correlations are given in Table 2. The MiX99 software (Lidauer and Strandén, 1999; Vuori et al., 2006) was used to calculate DRP and reliabilities of EBV.

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## **RESULTS AND DISCUSSION**

235 **Predictive correlation of GEBV** 

10-fold cross-validation. The mean predictive correlations of GEBV for CSC, INF and LAM were 236 low, varying from 0.27 to 0.37 (Table 3). The mean predictive correlation of GEBV increased 237 slightly, from 0.35 to 0.37, for CSC when including HQ and FA as correlated traits (CF1), whereas 238 including RLRV N and RLRV O (CF2) slightly decreased the mean predictive correlation of 239 GEBV. Including foot and leg conformation traits (CF1 and CF2) decreased the mean predictive 240 correlations of GEBV for INF and LAM compared to using CH (Table 3). The results suggest that 241 these genetically correlated traits may introduce more noise than additional information to the 242 prediction of GEBV. This may be because few cows had records on both claw disorders and foot 243 and leg conformation traits (Table 1), and that the genetic correlations among these traits were in 244 general low (Table 2). The standard deviation (SD) of predictive correlations of GEBV ranged 245 from 0.06 to 0.13 among the traits and datasets (Table 3), showing relatively large variation among 246 247 the folds (Figure 1). The highest SD for CSC, INF and LAM occurred using CF2, which had the lowest mean predictive correlation of GEBV and lowest number of cows with records on both 248 claw disorders and foot and leg conformation traits (Table 1). All mean predictive correlations of 249 GEBV for CSC, INF and LAM using CF1 and CF2 were within the range of one SD of the mean 250 predictive correlation of GEBV using the dataset CH. The overall best result for CSC was obtained 251 using dataset CF1 (Figure 1), whereas for INF it was obtained using dataset CH (Figure 1). For 252 LAM, datasets CH and CF1 gave very similar results over all folds (Figure 1). The large 253 differences in predictive correlations of GEBVs among validation sets in CV could be due to 254 255 unequal amount of information for sires in the validation set, or differences in the relationship of

a sire to the reference population. By using CV, and randomly assign sires in groups, some sires
in a validation set may be older elite sires having sons with information in the reference population,
and thereby gaining a lot of information in the CV compared to young sires with less data. This
could lead to overestimation of predictive correlation of GEBV, therefore an additional validation
set including the youngest sires were analyzed.

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Validation by youngest sires. Including foot and leg conformation traits increased the predictive 262 correlation of GEBV for CSC in VAL (Table 4), and the highest correlation was achieved using 263 the dataset CF1 (0.49) which included HQ that had strong genetic correlation to CSC (Table 2). 264 For INF the predictive correlation of GEBV was 0.33 to 0.34 in all 3 datasets, whereas for LAM 265 the predictive correlation of GEBV decreased when including foot and leg conformation traits 266 267 (Table 4). The predictive correlations of GEBV for INF and LAM from VAL (Table 4) were within the range of values found in CV (Table 3). For CSC the predictive correlations of GEBV from 268 CF1 and CF2 were above the maximum value in CV. Infectious claw disorder had low genetic 269 270 correlation with foot and leg conformation traits (Table 2), and was therefore expected to benefit less from including these as correlated traits in genomic prediction. This is reflected by the results, 271 where INF had the lowest predictive correlation of GEBV among the claw disorders and no gain 272 from correlated traits. The predictive correlations of GEBV for CSC, INF and LAM from 273 validation based on the youngest sires were similar as those obtained in CV, indicating that 274 overestimation was not a problem in this study. A benefit of using CV, compared to VAL, was the 275 obtained variance of the predictive correlation of GEBV, which is a measure of precision. 276

278 It was beneficial to include the foot and leg conformation traits HO and FA in genomic predictions 279 for CSC, whereas for INF and LAM including foot and leg conformation traits introduced more noise than additional information. Ødegård et al. (2014b) calculated the predictive ability of GEBV 280 281 (correlation between GEBV and daughter yield deviation) for CSC in a univariate (0.29) and bivariate model (0.32), including CSC and HQ, showing similar results as in the present study. 282 The higher predictive correlation of GEBV for CSC found in the present study (Tables 3 and 4) 283 compared to Ødegård et al. (2014b) could be due to different response variables and additional 284 traits included in the analyses. Karoui et al. (2012) showed that accuracy of GEBV increased 285 slightly in small breeds when highly genetic correlated traits from larger breeds were included in 286 the analyses. The low genetic correlation among most of the claw disorders and foot and leg 287 conformation traits (Table 2) could explain the small effect on predictive correlation of GEBV in 288 289 the present study. Buitenhuis et al. (2007) detected 4 QTL associated with lameness (group of claw disorders), and these had small overlap with QTL found for foot and leg conformation traits. This 290 indicate that different genes affect claw disorders and foot and leg conformation traits, which is 291 292 also consistent with the low genetic correlations among these traits (e.g. van der Waaij et al., 2005; Ødegård et al., 2014a). 293

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The accuracy of GEBV for other low heritability traits in Norwegian Red (e.g. Solberg et al., 2011; Svendsen et al., 2013; Haugaard et al., 2014) were in the same range as the predictive correlation of GEBV calculated in the present study. Haugaard et al. (2014) found accuracy of genomic predictions (correlation between EBV and GEBV) for 4 fertility related disorders in Norwegian Red ranging from 0.17 to 0.65. In Norwegian Red, correlations between GEBV and EBV were predicted for milk production traits to be around 0.6, whereas for health and fertility traits the

correlations ranged from 0.2 to 0.4 (Svendsen et al., 2013). Similar results were found in other
breeds (e.g. Karoui et al., 2012; Pintus et al., 2012; Zhou et al. 2014), where the accuracy of GEBV
were lowest for low heritable traits. Despite the limited historical data and the small reference
population available for claw disorders, the predictive correlations of GEBV for CSC, INF and
LAM were in the same range as accuracies of GEBV obtained for other low heritable traits in
Norwegian Red.

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### 308 Increasing the predictive correlation of GEBV

309 Claw disorders are novel traits with limited historical data and therefore fewer animals in the reference population. Including foot and leg conformation traits had little or no effect on the 310 predictive correlations of GEBV for CSC, INF and LAM, despite the increased number of sires in 311 the reference population. This could partly be because most sires had few daughters with claw 312 health information (average 168, minimum 30) and few cows had information on both claw health 313 and foot and leg conformation score. The high effective population size in Norwegian Red (Geno, 314 2013) and the low genetic correlations among the traits also affected the results. Better predictive 315 correlations of GEBV could possibly be obtained by increasing the number of animals in the 316 reference population, increasing the number of phenotypic records (claw health records) and by 317 genotyping of cows. 318

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Genomic predictions across breeds and populations is one approach to obtain larger reference populations (e.g. Brøndum et al., 2011; Heringstad et al., 2011; Lund et al., 2011) and thereby increase predictive correlation of GEBV. Reliabilities of GEBV for Norwegian Red calculated in a joint Nordic reference population (including Norwegian Red, Swedish Red, Finnish Ayrshire 324 and Danish Red) increased slightly for production traits compared to a reference population 325 consisting of only Norwegian Red. However, for health traits there was no gain in reliability and for fertility traits the reliability of GEBV decreased (Heringstad et al., 2011). Lund et al. (2011) 326 327 showed increased reliability of genomic prediction using a common reference population within breed, and Hozé et al. (2014) found increased gain in accuracy of genomic evaluation methods 328 using a multi-breed reference population in a small breed where bulls had missing sires in the 329 reference population. The results in these studies varied among breeds and populations, which 330 partly could be explained by variation in relationship among animals, as confirmed by Brøndum 331 et al. (2011) who concluded that reliabilities of direct breeding values increased when strong 332 genetic links between animals in a multi-breed reference population were present. 333

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The number of yearly claw health records has increased since national recording started in 2004, 335 to approximately 70,000 records per year. There is however a huge potential to further increase 336 the recording of claw health in Norway, as only 33% of the herds recorded claw health at claw 337 338 trimming in 2013. Number of daughters with claw health records for the 1,093 Norwegian Red sires in the present study varied from 30 to 6,524, and reliabilities of their EBV for CSC, INF and 339 LAM varied from 0.20 to 0.99. Mean reliability of EBV for CSC increased from 0.67 (using CH 340 and CF1) to 0.72 using CH1, whereas for INF and LAM it did not change between the 3 datasets. 341 The increased reliability of EBV for CSC using CF1 can be explained by more informative 342 daughters available for analyses, because of the strong genetic correlation between CSC and HQ. 343 In the present analyses only sires having at least 30 daughters with information were included, 344 whereas in routine genetic evaluations most sires have less than 30 daughters with claw health 345 346 records at the time of their first official proof. However, claw health information from more herds

can contribute with more information per sire and increased reliability of EBV, and therebyimproved phenotypes for genomic prediction.

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Genotyping of females to be included in the reference population is another possibility to increase 350 the predictive correlation of GEBV. Several studies have shown that genotyping of females are 351 beneficial in genomic predictions (e.g. Mc Hugh et al., 2011; Pryce et al., 2012; Egger-Danner et 352 al., 2014), especially in breeds with small reference populations or for novel traits. In a study where 353 the reference population consisted of genotyped cows with phenotypic records on new traits, 354 including genotyped bulls in the reference population with records on a positive genetic correlated 355 index increased the accuracy of selection (Calus et al., 2013). Egger-Danner et al. (2014) stated 356 that for novel traits, the reliability of GEBV would increase if genotyped cows with reliable 357 phenotypes were added to a small reference population, because bulls in the reference population 358 would have few daughters with records on the novel traits, and thereby less reliable GEBV. For 359 claw disorders in Norwegian Red, it might be beneficial to genotype cows with claw health records 360 361 to increase the reference population and thereby improve genomic predictions.

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This was the first genomic analyses of claw disorders in Norwegian Red. Although claw disorders are novel traits with limited historical data and small reference population, the predictive correlations of GEBV for CSC, INF and LAM were in the same range as for other health traits in Norwegian Red. Further increase in predictive correlation of GEBV may be achieved by getting more herds to record claw health, and by genotyping cows to be included in the reference population.

## CONCLUSION

The predictive correlations of GEBV for CSC, INF and LAM were in general low, and including 371 genetically correlated foot and leg conformation traits had little or no effect, despite the increased 372 373 reference population. The predictive correlation of GEBV for CSC increased slightly when including HQ and FA, whereas for other traits a small decrease were observed when including the 374 correlated traits. The results illustrate the challenges related to genomic selection of novel traits 375 376 with limited historical data and a small reference population. Including traits with strong genetic correlation may have some slight, positive influence on the predictive correlation of GEBV. 377 378 ACKNOWLEDGEMENTS 379 The authors would like to thank the Norwegian Dairy Herd Recording System and the Norwegian 380 Cattle Health Service (Ås, Norway) for access to data, Geno SA for access to SNP data, Harald 381 Grove, Cigene, NMBU, for imputation of SNP data, and the Norwegian Research Council and 382

383 Geno SA for funding (project 212864).

385	REFERENCES
386	Bruijnis, M. R. N., B. Beerda, H. Hogeveen, and E. N. Stassen. 2012. Assessing the welfare impact
387	of foot disorders in dairy cattle by modeling approach. Animal 6:962-970.
388	Bruijnis, M. R. N., H. Hogeveen, and E. N. Stassen. 2010. Assessing economic consequences of
389	foot disorders in dairy cattle using a dynamic stochastic simulation model. J. Dairy Sci.
390	93:2419-2432.
391	Buch, L. H., A. C. Sørensen, J. Lassen, P. Berg, JÅ. Eriksson, J. H. Jakobsen, and M. K. Sørensen.
392	2011. Hygiene-related and feed-related hoof diseases show different patterns of genetic
393	correlations to clinical mastitis and female fertility. J. Dairy Sci. 94:1540-1551.
394	Buitenhuis, A. J., M. S. Lund, J. R. Thomasen, B. Thomsen, V. Hunnicke Nielsen, C. Bendixen,
395	and B. Guldbrandtsen. 2007. Detection of quantitative trait loci affecting lameness and leg
396	conformation traits in Danish Holstein cattle. J. Dairy Sci. 90:472-481.
397	Brøndum, R. F., E. Rius-Vilarrasa, I. Strandén, G. Su, B. Guldbrandtsen, W. F. Fikse, and M. S.
398	Lund. 2011. Reliabilities of genomic prediction using combined reference data of the
399	Nordic Red dairy cattle populations. J. Dairy Sci. 94:4700-4707.
400	Calus, M. P. L., Y. de Haas, M. Pszczola, and R. F. Veerkamp. 2013. Predicted accuracy of and
401	response to genomic selection for new traits in dairy cattle. Animal 7:183-191.
402	Egger-Danner, C., H. Schwarzenbacher, and A. Willam. 2014. Short communication: Genotyping
403	of cows to speed up availability of genomic estimated breeding values for direct health
404	traits in Austrian Fleckvieh (Simmental) cattle – Genetic and economic aspects. J. Dairy
405	Sci. 97:4552-4556.
406	Fikse, W. F. and G. Banos. 2001. Weighting factors of sire daughter information in international

genetic evaluations. J. Dairy Sci. 84:1759-1767. 407

- Gao, H., M. S. Lund, Y. Zhang, and G. Su. 2013. Accuracy of genomic prediction using different
  models and response variables in the Nordic Red cattle population. J. Anim. Breed. Genet.
  130:333-340.
- 411 Geno, 2013. Årsberetning og regnskap 2013. Page 13. Accessed Feb. 10, 2015.
  412 http://viewer.zmags.com/publication/d5a081bc#/d5a081bc/1. (In Norwegian).
- 413 Haugaard, K., M. Svendsen, and B. Heringstad. 2014. Genomic predictions of fertility related
- disorders in Norwegian Red using 30 years of data. Proc. 10<sup>th</sup> World Congress on Genetics
   Applied to Livestock Production (WCGALP), Vancouver, Canada.
- 416 Hayes B. J., P. J. Bowman, A. J. Chamberlain, and M. E. Goddard. 2009. Invited review: Genomic
- 417 selection in dairy cattle: Progress and challenges. J. Dairy Sci. 92:433-443.
- Heringstad, B., G. Su, T. R. Solberg, B. Guldbrantsen, M. Svendsen, and M. S. Lund. 2011.
- 419 Genomic predictions based on a joint reference population for Scandinavian red breeds. Proc.
- 420 62<sup>th</sup> European Federation for Animal Science (EAAP) Annual Meeting, Stavanger, Norway.
- 421 Hozé, C., S. Fritz, F. Phocas, D. Boichard, V. Ducrocq, and P. Croiseau. 2014. Efficiency of multi-
- 422 breed genomic selection for dairy cattle breeds with different sizes of reference population. J.
- 423 Dairy Sci. 97:3918-3929.
- Karoui, S., M. J. Carabaño, C. Díaz, and A. Legarra. 2012. Joint genomic evaluation of French
  dairy cattle breeds using multiple-trait models. Genet. Sel. Evol. 44:39
- Lidauer, M., and I. Strandén, 1999. Fast and flexible program for genetic evaluation in dairy cattle.
- 427 International workshop in high performance computing and new statistical methods in dairy
- 428 cattle breeding, Tuusula, Finland. Interbull bull. 20:20-25.

- Luan, T., J. A. Woolliams, S. Lien, M. Kent, M. Svendsen, and T. H. E. Meuwissen. 2009. The
  accuracy of genomic selection in Norwegian Red cattle assessed by cross-validation. Genetics
  183:1119-1126.
- 432 Lund, M. S., A. P. W. de Roos, A. G. de Vries, T. Druet, V. Ducrocq, S. Fritz, F. Guillaume, B.
- 433 Guldbrandtsen, Z. Liu, R. Reents, C. Schrooten, F. Seefried, and G. Su. 2011. A common
- reference population from four European Holstein populations increases reliability of genomic
  predictions. Genet. Sel. Evol. 43:43.
- 436 Madsen, P., and J. Jensen. 2010. An user's guide to DMU. A package for analysing multivariate
- 437 mixed models. Version 6, release 5.0. University of Aarhus, Faculty Agricultural Science
- 438 (DJF), Dept. of Genetics and Biotechnology, Research Center Foulum, Tjele, Denmark.
- Mc Hugh, N., T. H. E. Meuwissen, A. R. Cromie, and A. K. Sonesson. 2011. Use of female
  information in dairy cattle genomic breeding programs. J. Dairy Sci. 94:4109-4118.
- Meuwissen, T. H. E., B. J. Hayes, and M. E. Goddard. 2001. Prediction of total genetic value using
  genome-wide dense marker maps. Genetics 157:1819-1829.
- 443 Pintus, M. A., E. L. Nicolazzi, J. B. C. H. M. Van Kaam, S. Biffani, A. Stella, G. Gaspa, C.
- Dimauro, and N. P. P. Macciotta. 2012. Use of different statistical models to predict direct
  genomic values for productive and functional traits in Italian Holsteins. J. Anim. Breed. Genet.
- 446 130:32-40.
- Pryce, J. E., B. J. Hayes, and M. E. Goddard. 2012. Genotyping dairy females can improve the
  reliability of genomic selection for young bulls and heifers and provide farmers with new
  management tools. Proc. International Committee for Animal Recording (ICAR), Cork,
  Irland.

- Sogstad, Å. M., O. Østerås, and T. Fjeldaas. 2006. Bovine claw and limb disorders related to
  reproductive performance and production diseases. J. Dairy Sci. 89:2519-2528.
- Sogstad, Å. M., O. Østerås, T. Fjeldaas, and A. O. Refsdal. 2007. Bovine claw and limb disorders
  at claw trimming related to milk yield. J. Dairy Sci. 90:749-759.
- Solberg, T. R., B. Heringstad, M. Svendsen, H. Grove, and T. H. E. Meuwissen. 2011. Genomic
  predictions for production- and functional traits in Norwegian Red from BLUP analyses of
- 457 imputed 54K and 777K SNP data. Interbull Bull. 44:240-243.
- Su, G., and P. Madsen. 2012. User's guide for Gmatrix. A program for computing genomic
  relationship matrix. Departments of Genetics and Biotechnology, Aarhus University.
- 460 Svendsen, M., B. Heringstad, and T. R. Solberg. 2013. Bruk av genomisk avlsverdi ved innkjøp
  461 av seminokseemner i NRF avlen. Husdyrforsøksmøte. Lillestrøm, Norway. (In Norwegian).
- 462 Swalve, H. H., H. Alkhoder, and R. Pijl. 2008. Estimates of breeding values for sires based on
- diagnoses recorded at hoof trimming: Relationships with EBV for conformation traits.
  Interbull Bull. 38:87-90.
- Uggla, E., J. H. Jakobsen, C. Bergsten, J.-Å. Eriksson, and E. Strandberg. 2008. Genetic
  correlations between claw health and feet and leg conformation traits in Swedish dairy cows.
  Interbull Bull. 38:91-95.
- Van der Waaij, E. H., M. Holzhauer, E. Ellen, C. Kamphuis, and G. de Jong. 2005. Genetic
  parameters for claw disorders in Dutch dairy cattle and correlations with conformation traits.
  J. Dairy Sci. 88:3672-3678.
- Vuori, K., I. Strandén, M. Lidauer, and E. A. Mäntysaari. 2006. MiX99 Effective solver for large
  and complex linear models. Proc. 8<sup>th</sup> World Congress on Genetics Applied to Livestock
  Production (WCGALP), Belo Horizonte, MG, Brazil.

- 474 Zhou, L., B. Heringstad, G. Su, B. Guldbrandtsen, T. H. E. Meuwissen, M. Svendsen, H. Grove,
- U. S. Nielsen, and M. S. Lund. 2014. Genomic predictions based on a joint reference
  population for the Nordic Red cattle breeds. J. Dairy Sci. 97:4485-4496.
- 477 Ødegård, C., M. Svendsen, and B. Heringstad. 2013. Genetic analyses of claw health in Norwegian
  478 Red cows. J. Dairy Sci. 96:7274-7283.
- Ødegård, C., M. Svendsen, and B. Heringstad. 2014a. Genetic correlations between claw health
  and feet and leg conformation in Norwegian Red cows. J. Dairy Sci. 97:4522-4529.
- 481 Ødegård, C., M. Svendsen, and B. Heringstad. 2014b. Predictive ability of genomic breeding
- 482 values for corkscrew claw in Norwegian Red. Proc. 10<sup>th</sup> World Congress on Genetics Applied
- 483 to Livestock Production (WCGALP), Vancouver, Canada.

485	Table 1. Number of claw health records above diagonal and number of Norwegian Red sires
486	with genotype and informative daughters (claw health records, foot and leg conformation scores
487	or both) below diagonal for each combination of corkscrew claw (CSC), infectious claw disorder
488	(INF), laminitis related claw disorder (LAM), hoof quality (HQ), foot angle (FA), rear leg rear
489	view new (RLRV_N) and rear leg rear view old (RLRV_O).

	CSC	INF	LAM	HQ	FA	RLRV_N	RLRV_O
CSC	281,835 1,093	281,835	281,835	25,598	3 25,598	11,803	13,795
INF	1,093	281,835 1,093	281,835	25,598	3 25,598	11,803	13,795
LAM	1,093	1,093	281,835 1,093	25,598	3 25,598	11,803	13,795
HQ	1,093	1,093	1,093	305,195 2,183	305,195	52,330	252,865
FA	1,093	1,093	1,093	2,183	421,319 3,111	52,330	368,834
RLRV_N	447	447	447	571	571	52,330 571	0
RLRV_O	816	816	816	1.782	2.710	170	368,834
				-,	_,, ~~		

Table 2. Estimated heritability of corkscrew claw (CSC), infectious claw disorder (INF), laminitis
related claw disorder (LAM), hoof quality (HQ), foot angle (FA), rear leg rear view new
(RLRV\_N) and rear leg rear view old (RLRV\_O) and their genetic correlation (standard errors) to
claw disorders.

		Genetic correlation						
Trait	Heritability	CSC	INF	LAM				
CSC	0.06							
INF	0.03	0.09 (0.07)						
LAM	0.03	0.26 (0.06)	0.25 (0.08)					
HQ	0.03	-0.79 (0.04)	-0.09 (0.07)	-0.27 (0.07)				
FA	0.09	0.08 (0.05)	0.10 (0.06)	0.11 (0.06)				
RLRV_N	0.08	0.03 (0.08)	-0.09 (0.09)	0.15 (0.09)				
RLRV_O	0.07	0.14 (0.06)	-0.02 (0.07)	0.14 (0.07)				

498	Table 3. Mean, standard deviation (SD), minimum value (Min) and maximum value (Max) of
499	predictive correlation of genomic breeding values (GEBV) for corkscrew claw (CSC), infectious
500	claw disorder (INF) and laminitis related claw disorder (LAM) from a 10-fold cross-validation.
501	Correlation between GEBV and deregressed proofs from multivariate models using 3 datasets:
502	$CH^1$ , $CF1^2$ and $CF2^3$ .

	CSC				INF				LAM			
Dataset	Mean	SD	Min	Max	Mean	SD	Min	Max	Mean	SD	Min	Max
CH <sup>1</sup>	0.35	0.07	0.25	0.45	0.32	0.10	0.13	0.52	0.33	0.06	0.24	0.42
CF1 <sup>2</sup>	0.37	0.07	0.28	0.47	0.29	0.08	0.17	0.49	0.32	0.06	0.22	0.41
CF2 <sup>3</sup>	0.31	0.10	0.15	0.42	0.27	0.13	0.00	0.53	0.29	0.07	0.16	0.36

 $^{1}$ CH – dataset including CSC, INF and LAM.

 $^{2}$ CF1 – dataset including CSC, INF, LAM, hoof quality and foot angle.

 $^{3}$ CF2 – dataset including CSC, INF, LAM, rear leg rear view new and rear leg rear view old.

Table 4. Predictive correlation of genomic breeding values (GEBV) for corkscrew claw, infectious
claw disorder and laminitis related claw disorder from validation by the 190 youngest sires.
Correlation between GEBV and deregressed proofs from multivariate models using 3 datasets:
CH<sup>1</sup>, CF1<sup>2</sup> and CF2<sup>3</sup>.

	$CH^1$	CF1 <sup>2</sup>	CF2 <sup>3</sup>
Corkscrew claw	0.38	0.49	0.43
Infectious claw disorder	0.33	0.34	0.33
Laminitis related claw disorder	0.41	0.36	0.36

<sup>1</sup>CH – dataset including corkscrew claw, infectious claw disorder and laminitis related claw
disorder.

<sup>515</sup> <sup>2</sup>CF1 – dataset including corkscrew claw, infectious claw disorder, laminitis related claw

516 disorder, hoof quality and foot angle.

<sup>517</sup> <sup>3</sup>CF2 – dataset including corkscrew claw, infectious claw disorder, laminitis related claw

518 disorder, rear leg rear view new and rear leg rear view old.

519









- 523 (INF) and laminitis related claw disorder (LAM) from 10-fold cross-validation using 3 datasets:
- 524 CH (CSC, INF and LAM); CF1 (CSC, INF, LAM, hoof quality and foot angle); and CF2 (CSC,
- 525 INF, LAM, rear leg rear view new and rear leg rear view old).